SYSTEMATIC REVIEW

Complications Rate and Hip Function After Revision of Infected Hip Arthroplasty with Bone Defects using Bone Allografts: A Systematic Review and Meta-Analysis

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Abstract

Objectives: We performed a systematic review and meta-analysis to evaluate complication rates and hip function following the revision of infected hip arthroplasty with bone defects using bone allografts.

Methods: A comprehensive search of the PubMed, Web of Science, Embase, and Cochrane Library databases was conducted up to January 2024 to identify pre-post clinical trials. The primary outcomes assessed were the risk of reinfection, a critical concern for surgeons, and hip functional scores. The methodological quality of the included studies was also evaluated. A weighted mean difference (WMD) with a 95% confidence interval (CI) was used as the pooled estimate for clinical outcomes through random-effects meta-analysis, accounting for heterogeneity across studies.

Results: Of the 2,189 records retrieved, 12 pre-post clinical trials (with fair to good quality) were included in the meta-analysis, involving a total of 342 participants. The pooled mean difference in Harris Hip Score (HHS) was 36.86 (95% CI: 29.58 to 44.13) post-surgery. In a subgroup analysis of studies employing structural grafts, the HHS increased by 36.99 (95% CI: 29.56 to 44.42). The overall reinfection rate was 6%. Subgroup analysis revealed that in studies utilizing morselized and structural allografts, the reinfection rates were 6% and 3%, respectively. The overall mean rate of aseptic loosening was 5%. Subgroup analysis showed that in studies using morselized grafts, the rate of aseptic loosening was 4%. The incidence of dislocation was 2% in the morselized group and 5% in the structural group.

Conclusion: Revision of infected hip arthroplasty with bone defects using bone allografts may improve hip function. Interestingly, morselized allografts are often associated with higher rates of reinfection. Additionally, our findings suggest that structural allografts are associated with increased dislocation rates compared to morselized allografts. This difference may be attributed to the larger and more complex defects that necessitated the use of structural allografts rather than morselized grafts.

Level of evidence: III

Keywords: Bone allograft, Infection, Meta-analysis, Total hip arthroplasty

Introduction

eriprosthetic joint infection (PJI) represents a highly complex challenge in the field of revision total hip arthroplasty (rTHA). Moreover, when this complication is accompanied by significant bone loss, the complexity of the situation is further compounded.^{1,2} Effectively managing this issue requires not only infection control but also the reconstruction of the deficient bone.

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Several strategies have been proposed to address acetabular bone deficiencies, including bone grafts (bulk or morselized), metal mesh, various cage designs, and tantalum metal augments. Historically, bone loss due to infection was considered a contraindication for reimplantation, primarily due to concerns about the risk of reinfection when using bone allografts. However,



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contemporary surgical approaches have become more cautious regarding the use of bone allografts in cases with a history of prior infection.

Adequate bone stock is the most crucial factor in long-term fixation, and the use of bone allografts to compensate for bone loss has yielded variable results. Acceptable outcomes have been reported in the mid-term when impacted allografts are used for rTHA in non-infected cases. While there have been proposals to use bone allografts in cases of infection. In this practice remains controversial due to concerns about the increased risk of reinfection. To enable evidence-based decision-making, there is a clear need for a systematic review of the available evidence to provide further insight into the safety of bone allografts in these cases.

This systematic review and meta-analysis aims to search for and evaluate studies reporting outcomes related to the use of bone allografts in revision THA for infected hip joints with acetabular or proximal femur defects. Specifically, we aim to determine: 1) the complication rate, particularly reinfection, and 2) the hip function as measured by the Harris Hip Score (HHS).

Materials and Methods

This review was conducted in accordance with the guidelines and checklist of the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA).¹³ This systematic review and meta-analysis were registered with the International Prospective Register of Systematic Reviews (PROSPERO) (CRD42023474113, https://www.crd.york.ac.uk/PROSPERO/).

Study Eligibility (Inclusion and Exclusion Criteria)

Studies were considered acceptable for inclusion in the systematic review if they met the following PICOD criteria: Population (P): rTHA following a previous infected THA with bone defects; Intervention (I): Bone allograft for bone defects (acetabular or proximal femur); Comparison (C): Not applicable; Outcome (O): Complications such as reinfection and hip functional scores; Design (D): Single-group beforeafter clinical trials. The following criteria were used for the exclusion of studies: animal studies, in vitro studies, letters to the editor, case reports, review articles, studies lacking data on the outcomes of interest, non-English language abstracts, and articles with insufficient data [Figure 1].

Literature Search

All authors established the research protocol for this review before the commencement of the literature searches. We conducted a comprehensive search for all pre-post clinical trials that evaluated the surgical outcomes or complication rates following the use of bone allografts in primary or revision THA for infected hip joints with bone (acetabular or femoral) defects. We searched the Cochrane Central Register of Controlled Trials, MEDLINE, Web of Science, and EMBASE for relevant controlled clinical trials from their inception to January 2024. The search strategy included the following MeSH terms and text keywords: ("Total hip arthroplasty" OR "Hip arthroplasty" OR "Hip replacement") AND ("Bone defect" OR "Acetabular defect") AND ("Infection" OR "Reinfection" OR "PJI" OR "infected revision" OR "septic arthritis"). The reference lists of the included studies were

also reviewed to identify relevant controlled clinical trials [Figure 1].

Study Selection and Data Abstraction

Two reviewers (M.S. and M.D.) independently extracted the following data: first author, year of publication, country, sample size, initial diagnosis, patient characteristics (age and sex), follow-up duration, HSS, pathogens isolated, complication rates (e.g., reinfection), type of reconstruction, antibiotic agents, type of bone allograft, duration of oral antibiotics, duration of intravenous antibiotics, and the time between excision arthroplasty and reconstruction THA for each study. A third reviewer (O.S.) resolved any disagreements between the two primary reviewers.

Risk of Bias Assessment

Along with the non-randomized before-and-after clinical trial study design of the included studies, we utilized the National Heart, Lung, and Blood Institute (NHLBI) Study Quality Assessment Tool for pre-post studies. This tool consists of 12 questions that assess the study's aim, sampling and sample size, description of the intervention and outcomes, blinding, follow-up, and statistical methods. The possible answers to these questions include: yes, no, cannot be determined (CD), not reported (NR), and not applicable (NA). The overall score is categorized as Good (score greater than 8), Fair (score between 5 and 8), or Poor (score less than 5). Two reviewers, MS and the first author, assessed the risk of bias in the studies, and the third reviewer, MD, verified their findings.

Data Analysis

The primary outcomes of our meta-analysis were reinfection rate, HHS, aseptic loosening rate, and dislocation rate. Forest plots were used to assess heterogeneity and calculate pooled weighted mean differences (WMD) and prevalence with corresponding 95% confidence intervals (95% CI). To account for heterogeneity in study populations, conducted random-effects meta-analysis. a Heterogeneity across studies was assessed using I² statistics, where $I^2 = 0\%$ indicated no observed heterogeneity and $I^2 \ge$ 50% showed substantial heterogeneity. Cochran's Q statistic was used to analyze the statistical significance of heterogeneity. A sensitivity analysis was performed to evaluate the impact of individual studies on heterogeneity and to assess the robustness of pooled estimates. Additionally, we conducted a meta-regression to determine the effect of participant age and follow-up duration on heterogeneity. Publication bias was not assessed using Egger's regression asymmetry test and Begg's adjusted rank correlation test due to the small number of studies included in the meta-analysis of HHS. All statistical tests were twotailed, and a significance level of less than 0.05 was set for all analyses, except for the heterogeneity test. Statistical analyses were conducted using Stata version 17.0 (Stata Corp., College Station, TX, USA).

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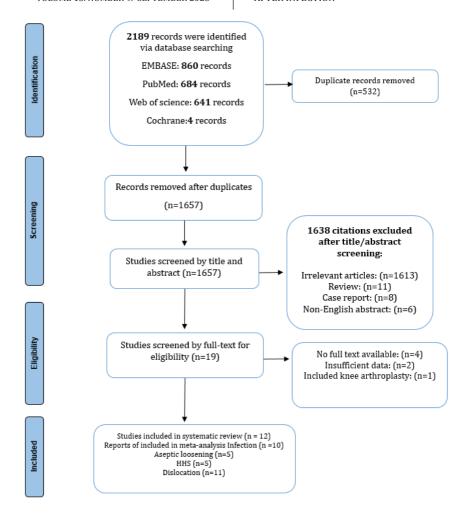


Figure 1. Flowchart of study selection

Results

Literature Search

A total of 2,189 potentially relevant citations were retrieved from the four electronic databases. After reviewing their titles and abstracts, 532 duplicates and 1,613 irrelevant citations were excluded, leaving 19 full-text articles for review. Ultimately, twelve studies published between 1997 and 2022 met the inclusion criteria and were included in the systematic review and meta-analysis. The detailed study selection process is shown in [Figure 1].

Study Selection and Data Abstraction

A total of 12 pre-post clinical trials involving 342 participants met the inclusion criteria, with 218 patients in

the morselized graft group and 91 patients in the structural graft group. The excision-to-reconstruction interval ranged from 0 to 316 weeks. The mean age of the patients ranged from 28 to 86 years, and the maximum follow-up period was 211 months. The study characteristics for each study are presented in [Tables 1 and 2].

Ten studies 9,10,14-21 involving a total of 321 patients have reported the reinfection rate. Five studies 10,15,21-23 involving a total of 167 patients have stated the aseptic loosening rate. Eleven studies,5, 9,10, 15-17, 19-23 involving a total of 324 participants, have evaluated the dislocation rate [Table 3]. Five studies 14,17,18,22,23 involving a total of 84 patients were included in the meta-analysis of HHS [Table 3].

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Table	1. Demographic da	ata of the inclu	ded studies					
Z	Author, Year (Ref)	Country	Study design	Population	Sample size patient (hip)	Age (year) mean (range)	Gender (F/M)	Follow up (month)/ mean (range)
1	Buttaro, 2003(9)	Argentina	RSC	Osteoarthritis/ fracture of the hip/ hip dysplasia/ osteonecrosis/ rheumatoid arthritis/ low grade chondrosarcoma	29 (30)	59 (32-78)	18/11	32.4 (24-60)
2	English, 2001 ¹⁰	England	RSC	Infected primary hip arthroplasty/ infected revision arthroplasty	53	69 (44-83)	27/26	53 (24 to 122)
3	Unfried, 2022 (14)	Brazil	RSC	-	18	(60.09-79.61)	12/6	45.29-82.39
4	Ammon, 2003 ¹⁵	UK	RSC	Osteoarthritis/ hip dysplasia/ Trauma/ Rheumatoid arthritis/ Perthes' disease/ Septic arthritis/ Avascular necrosis/ Ankylosing spondylitis/ Tumour/ Slipped upper femoral epiphysis	57	62 (28-82)	28/29	54 (24-126)
5	Winkler, 2008 ¹⁶	Austria	Prospective Cohort- Pre-post	-	37	68.5 (42-83)	20/17	52.8 (24-96)
6	Nusem, 2006 ¹⁷	Australia	Prospective Cohort- Pre-post	Primary osteoarthrosis/ traumatic osteoarthrosis/ ankylosing spondylitis	18	66 (45-86)	6/12	108 (60-168)
7	Lee, 2011 ¹⁸	Canada	RSC	Osteoarthritis followed by infection , developmental hip dysplasia, fracture , avascular necrosis, Perthes disease, and slipped upper femoral epiphysis	27	62 (28–83)		98.4 (13.2- 201.6)
8	Wang, 1996 ¹⁹	China	Prospective Cohort- Pre-post	Osteonecrosis/ fracture/ osteoarthritis	22	48 (28-75)	4/18	48 (24- 84)
9	Hsieh, 2004 ²⁰	Taiwan	RSC	-	24	59 (34-69)	7/17	50.4 (24-84)
10	Elbers, 2014 ²¹	Netherlands	Historical- prospective	-	36	61 (28-85)	20/16	118 (44-211).
11	Ilyas, 2001 ²²	Australia	RSC	Osteoarthritis/ traumatic arthritis/ ankylosing spondylitis	10	61.3 (42-79)	4/6	65 (36–120)
12	Alexeeff, 1995 ²³	Canada	RSC	Osteoarthritis/ rheumatoid arthritis/ Perthes' disease/ congenital dislocation of the hip/ Trauma	11	66.5 (44 - 83)	5/6	47.8 (24 - 72)

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Tabl	e 2. Surgic	al method	d details of the	included	studies'						
	ear	n stage	ype	ec.	ction interval range)	erial coating	Bone defect	severity	t type	ibiotics days	ws antibiotics op)
N	Author, Year	Reconstruction stage	Prosthesist	Prosthesis type Defect type Excision to reconstruction interval (week) mean (range) Prosthesis antibacterial coating	Prosthesis antibact	Femur type (n)	Acetabular type (n)	Bone allograft type	Duration of oral antibiotics days (Post-op)	Duration of intravenous antibiotics days (Post-op)	
1	Ammon, 2003	Two	Cemented (Palacos R)	A & F	median: 24 (4 - 316)	Vancomycin or gentamicin	-	-	Morselized (45) Structural (12)	-	Cefuroxime (750 mg tds) 2
2	Winkler, 2008	One	Cementless	A & F	NA	Vancomycin or tobramycin		< 2 (Paprosky)	Morselized	-	-
3	Nusem, 2006	Two	Cemented (Palacos R)	A & F	20 (4-32)	Gentamicin	Type I (6) Type III (4) (AAOS)	Type I (2) Type II (2) Type III (5) (AAOS)	Structural	-	-
4	Lee, 2011	Two	Cemented (spacer)	A & F	22 (7.6-56)	-			Structural	-	5
5	Wang, 1996	Two	Both	A & F	26.4 (6-96)	Gentamicin- vancomycin- tobromycin- cefazolin- clindamycin			Both	46 (14-84) (n=15)	16 (7-42) (n=18)
6	English, 2001	Two	Cemented	F	32 (4-240)	Vancomycin, gentamicin, or flucloxacillin			Morselized	-	79
7	Buttaro, 2003	Two	Both	A & F	14.7 (5 to 96)	Vancomycin	Type II(20); Type III (5); Type IV (5); (Endoklinik)	Type I (8); Type II (10); Type III(12); (AAOS)	Morsellised	42 (28 -112) (n=11)	43.4 (35- 56)
8	Hsieh, 2004	Two	Cemented	A & F	13.6 (11 to 17)	Vancomycin + piperacillin, Vancomycin +aztreonam, Vancomycin, Teicoplanin, Aztreonam	Type III (6); Type I (3); (AAOS)	Type III (12); Type IV (4); (AAOS)	Structural	-	7
9	Unfried, 2022	One	Both	A & F	NA	Vancomycin and cefepime		Type II (2); Type III (4); Type I (1) (AAOS)	Morsellised	140	30
10	Elbers, 2014	Two	Both	A & F	14 (0 to 124)	-			Morsellised	Yes	Yes
11	Ilyas, 2001	Two	Cemented (Palacos R)	F	13.6 (6-32)	Gentamicin	Type III (7); Type I (2); (AAOS)	Type III (1); (AAOS)	Structural	-	-
12	Alexeef, 1995	Two	cemented	A & F					Both	84	5

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Table 3	Table 3. Complications rate and hip functional score of the included studies												
	<u>.</u>	НН	IS mean (SD)			_	ing						
Z	Author, Year	Pre	Post	Dislocation N (%)	Fractures N (%)	Re-infection N (%)	Aseptic loosening N (%)	Nerve palsy N (%)	Others N (%)	Failure N (%)			
1	Ammon, 2003	-	-	6 (10.5)	-	7 (15.5) 1 (8.3)	3 (5.2)	1 (1.75)	Symptomatic trochanteric nonunion 3 (5.2)/Pulmonary embolus 1 (1.75) / Substantial haematoma 1(1.75)/ Broken stem of femoral implant 1 (1.75)	-			
2	Winkler, 2008	-	-	0 (0)	-	3 (8.10)	-	-	-	-			
3	Nusem, 2006	34 (19-57)	71 (46-97)	2 (11.1)	1 (5.5)	1 (5.5)	-	-	-	-			
4	Lee, 2011	39.2 (25-60)	1 year: 67.3 (40–91); Last follow: 70.3 (46–81)	1 (3.7)	-	1 (3.70)	-	1 (3.7)	unrelated deaths 2 (7.4)				
5	Wang, 1996	-	-	2 (9.09)	-	2 (9.09)	-	-	-	-			
6	English, 2001	-	-	1 (1.88)	3 (5.6)	4 (7.5)	3 (5.6)	1 (1.88)	Cerebrovascular accident 1(1.88)/ Pulmonary embolism 1(1.88)/ Removal of trochanteric Cablegrip 1(1.88)	-			
7	Buttaro, 2003	-	-	2 (6.89)	1 (3.44)	1(3.3)	-	-	Displacements of the greater trochanter 4(13.3)	-			
8	Hsieh, 2004	-	-	1 (4.16)	2 (8.33)	0 (0)	-	-	-	-			
9	Unfried,	(38.55-73.83)	82.55±11.49	-	-	0 (0)	-		Pruritus and redness of the skin 1(5.5)/Acute renal failure 2(11.1)/Deep vein thrombosis 1(5.5)	·			
10	Elbers, 2014	-	-	2(5.5)	1 (2.7)	4 (11.1)	1(2.7)	-	Persistence of pain 1(2.7)	-			
11	Ilyas, 2001	27.4 (9-58)	73.5 (53-92)	1 (10)	1 (10)	-	1(10)	-	Non-union 1(10)/Haematoma 1(10)	-			
12	Alexeeff, 1995	27.0 (10 - 52)	72.1 (63 - 80)	1 (9.09)	-	-	1 (9.09)	1 (9.09)	Acetabulum revised 2 (18.8)	-			

Risk of Bias Assessment

Of the 12 studies, six (50%) were of good quality, and six (50%) were of fair quality. Nine studies (75%) had insufficient sample sizes. Three studies (25%) considered blinding. Most of these studies had concerns regarding statistical methods and sample size. The majority of studies received good ratings for the clarity of study objectives,

specifying selection criteria, participant enrollment, defining and measuring the intervention, and assessing outcomes and loss to follow-up. The detailed results of this appraisal are presented in [Table 4].

Table 4.	Methodological a	ssessm	ent of	pre-po	st clini	ical tri	al stud	ies wit	h the N	NHLBI :	Study Q	uality A	ssessme	nt Tool
ID	Study	Q1	Q2	Q3	Q4	Q5	Q6	Q7	Q8	Q9	Q10	Q11	Q12	Quality rating
1	Ammon,2003	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes	NA	No	NA	Good
2	Nusem,2005	Yes	No	CD	Yes	No	Yes	Yes	No	Yes	NA	No	NA	Fair
3	Lee 2011	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	No	Yes	Yes	NA	Good
4	Winkler,2008	Yes	Yes	No	Yes	Yes	Yes	Yes	No	Yes	Na	No	NA	Good
5	Ilyas,2001	Yes	Yes	CD	Yes	No	Yes	No	No	Yes	No	No	NA	Fair
6	Buttaro,2004	Yes	No	Yes	Yes	No	Yes	Yes	No	Yes	No	No	NA	Fair

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Table 4. Continued														
7	Elbers, 2014	Yes	Yes	Yes	Yes	No	Yes	Yes	No	Yes	No	Yes	NA	Good
8	English, 2002	Yes	No	CD	No	Yes	Yes	Yes	No	Yes	NR	NR	NA	Fair
9	Hsieh, 2004	No	Yes	Yes	Yes	No	Yes	Yes	No	Yes	NR	NR	NA	Good
10	Unfried, 2022	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	NA	Good
11	Wang, 1997	Yes	Yes	Yes	Yes	No	Yes	No	Yes	Yes	No	Yes	NA	Fair
12	ALEXEEFF, 1995	Yes	Yes	No	Yes	No	Yes	Yes	No	Yes	No	No	NA	Fair

The Questions include 1: Objective clear, 2: Eligibility specified, 3: Participant's representative, 4: All eligible enrolled, 5: Sample size, 6: Intervention, 7: Outcome measures, 8: Blinded assessors, 9: Follow up rate, 10: Statistical analysis, 11: Multiple outcome measures, 12: Individual level data to determine group effect.

Data analysis HHS

Analyzing five studies, the mean difference in HHS was 36.85 points (range, 29.67 to 44.04) after rTHA [Figure 2]. The I^2 statistic showed moderate heterogeneity among the reported data for HHS (I^2 : 66.5%, P = 0.018). To determine which study (if any) had the most impact on heterogeneity and to assess the robustness of the summary findings, a sensitivity analysis was performed by successively removing

one study at a time. In this sensitivity analysis, excluding one study at a time consistently resulted in a significantly improved HHS (range of summary WMDs, 34.49 to 38.41). Analyzing three studies, the mean HHS was 37.02 points (range, 29.46 to 44.55) among the studies using structural bone allografts [Figure 2]. The $\rm I^2$ statistic showed low heterogeneity among the reported data for HHS ($\rm I^2$: 46.7%, P = 0.15).

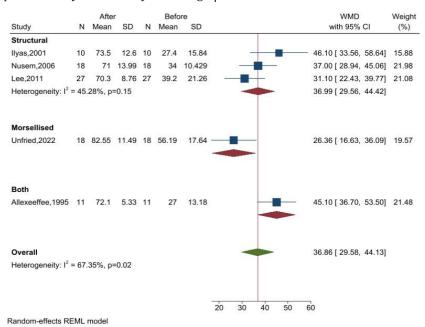


Figure 2. Meta-analysis of Harris Hip Score (HHS) for the effect of bone structural allograft for joint arthroplasty after infected hip. Diamond represents the summary weighted mean difference (pooled WMD) estimate and its width shows corresponding 95% CI with random effects estimate. I-square test and Cochran's Q statistic were used to assess the statistical heterogeneity (P = 0.15) across studies

Aseptic loosening

Meta-analysis of five studies showed that the aseptic loosening rate was 0.04 (range, 0.01 to 0.09) after rTHA [Figure 3]. The I² statistic indicated low heterogeneity among

the reported data for aseptic loosening (I^2 : 0.00%, P = 0.82). To determine which study (if any) had the most impact on heterogeneity and to assess the robustness of the summary findings, a sensitivity analysis was performed by successively

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removing individual studies. In this sensitivity analysis, excluding one study at a time consistently resulted in findings that remained broadly similar (range of summary prevalence: 0.043, 0.058). In the analysis of two studies using bone morselized allografts, the pooled aseptic loosening rate

was 0.04 (range, 0.01 to 0.10) [Figure 3]. Similarly, in two studies using both structural and morselized allografts, the pooled aseptic loosening rate was 0.05 (range, 0.00 to 0.12) [Figure 3].

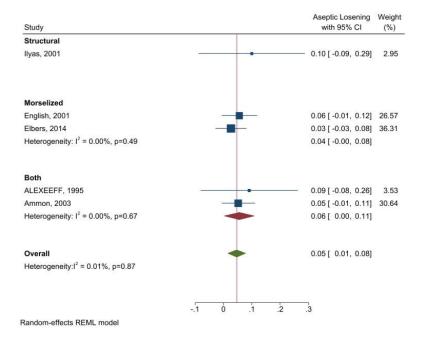


Figure 3. Subgroup meta-analysis of prevalence of aseptic loosening for the effect of bone allograft for joint arthroplasty after infected hip based on the type of allograft

Reinfection

Meta-analysis of ten studies showed that the reinfection rate was 0.06 (range, 0.04 to 0.10) after rTHA [Figure 4]. The I² statistic indicated low heterogeneity among the reported data for reinfection (I^2 : 9.54%, P = 0.35). To determine which study (if any) had the most impact on heterogeneity and to assess the robustness of the summary findings, a sensitivity analysis was performed by successively removing individual studies. In this sensitivity analysis, excluding one study at a time consistently resulted in findings that remained largely similar (range of summary prevalence: 0.06, 0.08). In the analysis of five studies using bone morselized allografts, the reinfection rate was 0.06 (range, 0.03 to 0.11) [Figure 4]. The I² statistic showed low heterogeneity among the reported data for reinfection (I^2 : 0.00%, P = 0.50). In three studies using bone structural allografts, the mean reinfection rate was 0.02 (range, 0.00 to 0.08) [Figure 4].

Dislocation

Meta-analysis of 11 studies showed that the dislocation rate

was 0.05 (range, 0.02 to 0.08) after rTHA [Figure 5]. The I^2 statistic indicated moderate heterogeneity among the reported data for dislocation rate (I^2 : 7.38%, P = 0.37). To determine which study (if any) had the most impact on heterogeneity and to assess the robustness of the summary findings, a sensitivity analysis was performed by successively removing individual studies. In this sensitivity analysis, excluding one study at a time consistently resulted in findings that remained largely similar (range of summary prevalence: 0.04, 0.06). In the analysis of four studies using bone morselized allografts, the dislocation rate was 0.03 (range, 0.00 to 0.07) [Figure 5]. The I² statistic showed low heterogeneity among the reported data for dislocation (I²: 22.35%, P = 0.28). In the analysis of four studies using bone structural allografts, the dislocation rate was 0.06 (range, 0.01 to 0.13) [Figure 5]. The I² statistic showed low heterogeneity among the reported data for dislocation rate $(I^2: 0.00\%, P = 0.72).$

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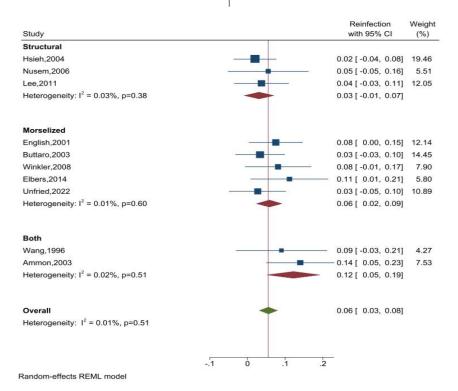


Figure 4. Subgroup meta-analysis of prevalence of Reinfection for the effect of bone allograft for joint arthroplasty after infected hip based on the type of allograft. The pooled prevalence of reinfection in Structural, Morselized, and both types of allograft is 3,6 and 12%, respectively

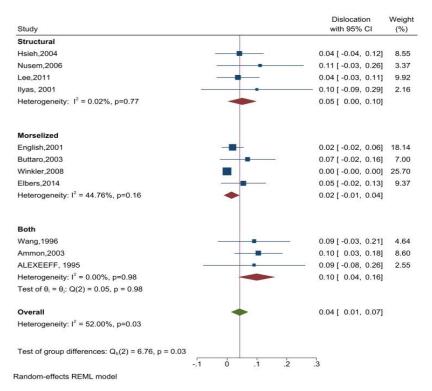


Figure 5. Subgroup meta-analysis of prevalence of dislocation for the effect of bone allograft for joint arthroplasty after infected hip based on the type of allograft. The prevalence of dislocation in Structural, Morselized, and both types of allograft is 5,2 and 10%, respectively

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Discussion

The outcomes of bone allografts in addressing acetabular or femoral bone defects resulting from infection during revision THA remain uncertain. To address this gap in knowledge, we conducted a systematic review and meta-analysis of studies involving patients with hip prosthesis infection and subsequent bone defects who underwent rTHA with bone allograft. Our objective was to evaluate the incidence of complications, particularly reinfection, and assess hip functional scores following this treatment.

Findings from ten studies revealed a pooled reinfection rate of 6%. The primary complications reported included reinfection (6%), aseptic loosening (5%), and dislocation (4%). Reinfection occurred in five studies using morselized allografts (combined estimate: 6%), 9,10,14,16,21 three studies utilizing structural allografts (combined estimate: 3%), 17,18,20 and two studies employing both structural and morselized allografts (combined estimate: 12%). 15,19 The rate of reinfections following aseptic revisions (without prior infection) is slightly higher at 8.13%,²⁴ indicating that revision procedures inherently carry a notable infection risk, regardless of prior infection status or graft usage. While previous literature suggests that the appropriate type of acetabular bone graft is typically selected based on the classification of the bone stock deficit — with cavitary defects often restored using cancellous morselized auto- or allografts, and larger segmental bone stock defects generally addressed with structural (or bulk) cortico-cancellous autoor allografts ²⁵— a recent meta-analysis by Strahl et al. indicates that both graft types were equally utilized across all AAOS and Paprosky classifications for significant bone loss²⁶ Concerns persist regarding the heightened risk of reinfection associated with large allografts, potentially due to impediments to antibiotic delivery and the host immune response. Despite various techniques being employed, a consensus on the optimal treatment approach remains elusive.

The choice between one-stage and two-stage procedures for treating PIIs after multiple unsuccessful surgeries remains a topic of significant debate. Some studies suggest that twostage procedures may offer advantages in reducing the risk of infection recurrence, primarily due to the opportunity for two separate debridement and cleaning interventions²⁷ However, the evidence remains inconclusive regarding the superiority of either approach. For instance, Winkler et al. 16 and Unfried et al.¹⁴ both studied one-stage procedures using morselized allografts, reporting reinfection rates of 8% and 3%, respectively. On the other hand, two-stage revisions have shown varied outcomes. Ammon et al. 15 reported a 14% reinfection rate, while Wang and Chen¹⁹ noted a 9% reinfection rate. These studies involved mixed patient cohorts treated with either morselized or structural allografts, with follow-up periods ranging from 24 to 126 months. Given the variability in reinfection rates across studies and techniques, it remains challenging to definitively conclude whether one-stage or two-stage revisions are more effective in minimizing reinfection rates. Further research is required to provide more robust comparative data.

Revision THA presents specific challenges when managing PJIs. In addition to addressing bone loss, two additional obstacles must be overcome. First, eradicating infection is difficult due to biofilm formation, especially when dealing with "difficult-to-treat" pathogens. Second, patients often present with various systemic diseases, rendering them immunocompromised. This complicates the recovery process and can negatively impact overall outcomes for this patient population. Performing acetabular bone grafting requires a high level of technical skill and should be considered only when other reconstruction methods are unlikely to yield a lasting result. Surgeries for acetabular or femoral bone defects caused by infection during revision THA should be performed only at specialized referral centers. Si

Additionally, it is essential to consider the potential protective role of antibacterial coatings in reducing the risk of infection recurrence. The effectiveness of silver-coated medical implants has been highlighted in oncology patients to the extent that it has become the standard approach for this particular patient subgroup³² However, a limitation of silver coatings is their ability to cover only the outer surface of medical implants. To address this limitation, a rapidly absorbable hydrogel coating known as DAC®, composed of hyaluronan and poly-D, L-lactide covalently linked, has been developed. This coating aims to prevent bacterial colonization in the immediate post-surgery period, offering short-term localized antibiotic release while minimizing potential adverse effects and the development of antibiotic resistance.33,34 Studies by Ammon et al. and Białecki et al., which reported the highest reinfection rates, used a single agent (vancomycin or gentamicin) for the antibacterial coating.

The weighted mean difference for HHS in the structural allograft group was notably high at 36.86, indicating an improvement in hip function post-surgery. Aseptic loosening occurred in two studies using morselized allografts (pooled estimate: 4%),^{10,21} one study using structural allografts²², and two studies using both structural and morselized allografts (pooled estimate: 6%).^{15,23} Since radiographic evidence, such as radiolucent lines and cup migration, could not be consistently included in the success criteria, the rate of aseptic loosening is likely higher. Regarding dislocation, structural allografts demonstrated a pooled prevalence of 5%, morselized allografts 2%, and procedures involving both types of allografts yielded a pooled prevalence of 10%. The dislocation rate after aseptic revision THA is comparable to that of structural allografts (5%) after one year.²⁴

Despite literature documenting favorable clinical and radiological outcomes in the short- to long-term, along with survival rates, for acetabular and femoral revisions employing allografts,^{35,36} the precise factors contributing to successful allograft integration remain not fully understood. It is likely that factors such as defect size, surgical approach, allograft type, use of bone cement, and patient-related variables all contribute to graft incorporation.

To our knowledge, this is the first systematic review analyzing the reinfection rate of rTHA involving bone

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allograft utilization in both acetabular and femoral defects. Our study has several limitations. The included studies varied in terms of the severity of acetabular or femoral bone defects. Complications across all studies were aggregated without categorization based on Paprosky's bone loss types. Nonetheless, this diversity allowed us to assess the effectiveness of bone allograft techniques across a wide range of bone defects encountered in surgical practice. Another limitation is the lack of detailed procedural data, such as prosthesis features, in some of the included studies. Notably, all included studies were retrospective follow-up studies without control groups. Variations in surgical approaches, postoperative care protocols, age at primary revision, patient demographics, primary disease, number of cases (ranging from 20 to 304), and time to follow-up (ranging from 2 to 22 years) resulted in limited comparability. However, the calculated I² statistics for heterogeneity were low or moderate across all conducted meta-analyses.

Conclusion

The reinfection rate after revision hip arthroplasty for infection using bone allografts ranges from 2% to 12% across the included studies. Morselized allografts, in particular, tend to have higher reinfection rates. Moreover, our analysis indicates that structural allografts are associated with a higher risk of dislocation compared to morselized allografts. Nevertheless, the evidence suggests that using allografts in patients whose infections have been fully resolved is safe and does not increase the risk of reinfection when compared to resections performed without allografts.

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References

- 1. Gross AE, Wong P, Saleh KJ. Don't throw away the ring: Indications and use. J Arthroplasty. 2002;17(4):162-166. doi.org/10.1054/arth.2002.32464.
- 2. Guntin J, Serino J, Rossi D, Boniello A, Gusho CA, Della Valle CJ. Hypoalbuminemia increases mortality after two-stage revision total joint arthroplasty. Arch Bone Jt Surg. 2023;11(3):173. doi.org/10.22038/ABJS.2022.65148.3123.
- 3. Del Gaizo DJ, Kancherla V, Sporer SM, Paprosky WG. Tantalum augment for Paprosky IIIA defects remains stable at midterm follow-up. Clin Orthop Relat Res. 2012;470:395-401. doi: 10.1007/s11999-011-2170-x
- 4. Salvati EA, Chekofsky KM, Brause BD, Wilson Jr PD. Reimplantation in infection: a 12-year experience. Clin Orthop Relat Res. 1982;170:62-75.
- Lee J-M, Nam H-T. Acetabular Revision Total Hip Arthroplasty Using an Impacted Morselized Allograftand a Cementless Cup: Minimum 10-Year Follow-Up. J Arthroplasty. 2011;26(7):1057-1060. doi.org/10.1016/j.arth.2011.03.035.
- 6. Boldt JG, Dilawari P, Agarwal S, Drabu KJ. Revision total hip arthroplasty using impaction bone grafting with cemented nonpolished stems and Charnley cups. J Arthroplasty. 2001;16(8):943-952. doi.org/10.1054/arth.2001.25559.

- Piccaluga F, Della Valle AG, Fernández JE, Pusso R. Revision of the femoral prosthesis with impaction allografting and a Charnley stem: a 2-to 12-year follow-up. J Bone Joint Surg Br. 2002;84(4):544-549. doi.org/10.1302/0301-620X.84B4.0840544.
- 8. Berry DJ, Chandler HP, Reilly DT. The use of bone allografts in two-stage reconstruction after failure of hip replacements due to infection. J Bone Joint Surg Am. 1991;73(10):1460-8.
- Buttaro MA, Pusso R, Piccaluga F. Vancomycin-supplemented impacted bone allografts in infected hip arthroplasty Article. Journal of Bone and Joint Surgery - Series B. 2005;87(3):314-319. doi:10.1302/0301-620X.87B3.14788
- 10. English H, Timperley AJ, Dunlop D, Gie G. Impaction grafting of the femur in two-stage revision for infected total hip replacement. J Bone Joint Surg Br. Jul 2002;84(5):700-5. doi:10.1302/0301-620x.84b5.12504
- 11. Nestor BJ, Hanssen AD, Ferrer-Gonzalez R, Fitzgerald Jr RH. The use of porous prostheses in delayed reconstruction of total hip replacements that have failed because of infection. JBJS. 1994;76(3):349-359.
- 12. Morscher E, Babst R, Jenny H. Treatment of infected joint arthroplasty. Int Orthop. 1990;14:161-165. doi.org/10.1007/BF00180122.
- Moher D, Shamseer L, Clarke M, et al. Preferred reporting items for systematic review and meta-analysis protocols (PRISMA-P) 2015 statement. Systematic reviews. 2015;4:1-9. doi.org/10.1186/2046-4053-4-1.
- 14. Unfried RI, Krause LMF, Cezimbra HM, Pacheco LS, Larangeira JA, Ribeiro TA. A Retrospective Observational Cohort Study of Periprosthetic Hip Infection Treated by one-stage Method Including Cases With Bone Graft Reconstruction. Clin Med Insights Arthritis Musculoskelet Disord. 2022;15:11795441221090344. doi:10.1177/11795441221090344
- Ammon P, Stockley I. Allograft bone in two-stage revision of the hip for infection: is it safe? The Journal of Bone & Joint Surgery British Volume. 2004;86(7):962-5.
- 16. Winkler H, Stoiber A, Kaudela K, Winter F, Menschik F. One stage uncemented revision of infected total hip replacement using cancellous allograft bone impregnated with antibiotics. The Journal of Bone & Joint Surgery British Volume. 2008;90(12):1580-4.
- 17. Nusem I, Morgan DA. Structural allografts for bone stock reconstruction in two-stage revision for infected total hip arthroplasty: good outcome in 16 of 18 patients followed for 5–14 years. Acta orthopaedica. 2006;77(1):92-7.
- 18. Lee PT, Clayton RA, Safir OA, Backstein DJ, Gross AE. Structural allograft as an option for treating infected hip arthroplasty with massive bone loss. Clinical Orthopaedics and Related Research®. 2011;469(4):1016-23.
- Wang J-W, Chen C-E. Reimplantation of infected hip arthroplastics using bone allografts. Clinical Orthopaedics and Related Research (1976-2007). 1997;335:202-10.
- Hsieh P-H, Shih C-H, Chang Y-H, Lee M, Yang W-E, Shih H-N. Treatment of deep infection of the hip associated with massive bone loss: two-stage revision with an antibiotic-loaded interim cement prosthesis followed by reconstruction with allograft. The Journal of Bone & Joint Surgery British Volume. 2005;87(6):770-5.

- 21. Elbers JB, Leijtens B, van Werven HE, Sturm PD, Kullberg BJ, Schreurs BW. Antibiotic mixing through impacted bone grafts does not seem indicated in two-stage cemented hip revisions for septic loosening. Hip International. 2014;24(6):596-603.
- 22. Ilyas I, Morgan D. Massive structural allograft in revision of septic hip arthroplasty. International orthopaedics. 2001;24(6):319-22.
- Alexeeff M, Mahomed N, Morsi E, Garbuz D, Gross A, Latner A. Structural allograft in two-stage revisions for failed septic hip arthroplasty. The Journal of Bone & Joint Surgery British Volume. 1996;78(2):213-6.
- Badarudeen S, Shu AC, Ong KL, Baykal D, Lau E, Malkani AL. Complications after revision total hip arthroplasty in the medicare population. The Journal of Arthroplasty. 2017;32(6):1954-8.
- Gamradt SC, Lieberman JR. Bone graft for revision hip arthroplasty: biology and future applications. Clinical Orthopaedics and Related Research (1976-2007). 2003;417:183-94.
- 26. Strahl A, Boese CK, Ries C, Hubert J, Beil FT, Rolvien T. Outcome of different reconstruction options using allografts in revision total hip arthroplasty for severe acetabular bone loss: a systematic review and meta-analysis. Archives of orthopaedic and trauma surgery. 2023;143(10):6403-22.
- Logoluso N, Pedrini FA, Morelli I, De Vecchi E, Romanò CL, Pellegrini AV. Megaprostheses for the revision of infected hip arthroplasties with severe bone loss. BMC surgery. 2022;22(1):68.
- 28. Wimmer MD, Hischebeth GT, Randau TM, Gathen M, Schildberg FA, Fröschen FS, et al. Difficult-to-treat pathogens significantly reduce infection resolution in periprosthetic joint infections. Diagnostic Microbiology and Infectious Disease. 2020;98(2):115114.
- George DA, Drago L, Scarponi S, Gallazzi E, Haddad FS, Romano CL. Predicting lower limb periprosthetic joint infections: A review of risk factors and their classification. World journal of orthopedics. 2017;8(5):400.
- Akgün D, Perka C, Trampuz A, Renz N. Outcome of hip and knee periprosthetic joint infections caused by pathogens resistant to biofilm-active antibiotics: results from a prospective cohort study. Archives of orthopaedic and trauma surgery. 2018;138(5):635-42.
- 31. Corona PS, Vicente M, Lalanza M, Amat C, Carrera L. Use of modular megaprosthesis in managing chronic end-stage periprosthetic hip and knee infections: Is there an increase in relapse rate? European Journal of Orthopaedic Surgery & Traumatology. 2018;28(4):627-36.
- Wafa H, Grimer R, Reddy K, Jeys L, Abudu A, Carter S, Tillman R. Retrospective evaluation of the incidence of early periprosthetic infection with silver-treated endoprostheses in high-risk patients: case-control study. The bone & joint journal. 2015;97(2):252-7.
- 33. Romanò CL, Malizos K, Capuano N, Mezzoprete R, D'Arienzo M, Der CV, et al. Does an antibiotic-loaded hydrogel coating reduce early post-surgical infection after joint arthroplasty? Journal of bone and joint infection. 2016;1(1):34-41.
- 34. Capuano N, Logoluso N, Gallazzi E, Drago L, Romanò CL.

SAFETY OF BONE ALLOGRAFT FOR ACETABULAR RECONSTRUCTION AFTER INFECTION

One-stage exchange with antibacterial hydrogel coated implants provides similar results to two-stage revision, without the coating, for the treatment of peri-prosthetic infection. Knee Surgery, Sports Traumatology, Arthroscopy. 2018;26(11):3362-7.

- 35. Ullmark G, Hallin G, Nilsson O. Impacted corticocancellous allografts and cement for femoral revision of total hip
- arthroplasty using Lubinus and Charnley prostheses. The Journal of arthroplasty. 2002;17(3):325-34.
- 36. Schreurs B, Keurentjes J, Gardeniers J, Verdonschot N, Slooff T, Veth R. Acetabular revision with impacted morsellised cancellous bone grafting and a cemented acetabular component: a 20-to 25-year follow-up. The Journal of Bone & Joint Surgery British Volume. 2009;91(9):1148-53.