

Effect of Rheumatoid Arthritis on Postoperative Outcomes in Patients with Lumbar Spinal Disorders: A Systematic Review and Meta-Analysis

Global Spine Journal 2025, Vol. 0(0) 1–9 © The Author(s) 2025 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/21925682251318265 journals.sagepub.com/home/gsj Sage

Shintaro Honda, MD¹[®], Koichi Murata, MD, PhD^{1,2}, Masaki Sakamoto, MD¹, Akihiro Shiroshita, MD, MPH^{3,4}, Natsumi Saka, MD, PhD^{4,5}, Bungo Otsuki, MD, PhD¹, Takayoshi Shimizu, MD, PhD¹[®], Takashi Sono, MD, PhD¹, Soichiro Masuda, MD¹[®], Koichiro Shima, MD¹, and Shuichi Matsuda, MD, PhD¹

Abstract

Study Design: A systematic review and meta-analysis.

Objective: This study aimed to determine whether rheumatoid arthritis (RA) is associated with clinical outcomes following spinal surgery for lumbar spinal disorders.

Methods: MEDLINE, Embase, the Cochrane Library, and the International Clinical Trials Registry Platform were comprehensively searched for observational studies comparing clinical outcomes after lumbar spine surgery in patients with and without RA (>18 years). Quality assessment was conducted using the Quality in Prognosis Studies assessment tool. Pooled odds ratios (ORs) and hazard ratios were calculated for reoperation and surgical site infection by using a random effects model. Subgroup analyses were conducted to examine the effect of surgery type.

Results: Seven studies with 72,969 patients, including 7518 patients with RA, were analyzed. All studies had a moderate risk of bias. Patients with RA had a significantly higher odds of reoperation (OR: 5.57; 95% confidence interval [CI]: 1.10-28.26; I2 = 92%; P = 0.04) and higher odds of surgical site infection (OR: 1.47; 95% CI: 1.28-1.69; I2 = 2%; P < 0.01). No statistically significant difference was found in reoperation-free survival between patients with RA and those without RA (hazard ratio: 1.15; 95% CI: 0.94-1.40; I2 = 100%; P = 0.16). Patients with RA had higher complication rates, with incidence rates ranging from 13.5% to 57%, compared with those without RA.

Conclusion: Compared with patients without RA, patients with RA may be more likely to undergo reoperations and suffer from complications following surgery for lumbar spine lesions.

³ Division of Epidemiology, Department of Medicine, Vanderbilt University School of Medicine, Nashville, TN, USA

Corresponding Author:

Shintaro Honda, MD, Department of Orthopedic Surgery, Kyoto University Graduate School of Medicine, 54 Kawahara-cho, Shogoin, Sakyo, Kyoto 606-8507, Japan.

Email: s_honda726@kuhp.kyoto-u.ac.jp



Creative Commons Non Commercial No Derivs CC BY-NC-ND: This article is distributed under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs 4.0 License (https://creativecommons.org/licenses/by-nc-nd/4.0/) which permits non-commercial use, reproduction and distribution of the work as published without adaptation or alteration, without further permission provided the original work is attributed as specified on the SAGE and Open Access pages (https://us.sagepub.com/en-us/nam/open-access-at-sage).

¹ Department of Orthopaedic Surgery, Kyoto University Graduate School of Medicine, Kyoto, Japan

² Department of Advanced Medicine for Rheumatic Diseases, Kyoto University Graduate School of Medicine, Kyoto, Japan

⁴ Scientific Research Works Peer Support Group (SRWS-PSG), Osaka, Japan

⁵ Department of Orthopaedics, Teikyo University School of Medicine, Tokyo, Japan

Keywords

rheumatoid arthritis, spinal surgery, surgical site infection, clinical outcome

Introduction

Lumbar spinal disorders in patients with rheumatoid arthritis (RA) are complex and challenging, with the disease often affecting the facet joints and discovertebral junctions.¹⁻³ Severe lumbar lesions have serious detrimental effects on the quality of life of patients with RA.⁴ Patients with RA often develop osteoporotic vertebral fractures.⁵ A correlation between disease activity and spinal malalignment was reported.⁶ In a recent longitudinal analysis, we demonstrated that the disease activity of RA contributes to the deterioration of spinal sagittal alignment.⁷

Surgical intervention is a key treatment modality for patients with RA with lumbar lesions refractory to conservative treatment. For these lesions, lumbar fusion or decompression surgery is mainly performed.⁸ However, evidence regarding the association between RA and postoperative outcomes in patients with lumbar spinal disorders is lacking. Several studies suggest that RA has a negative effect on postoperative outcomes, whereas others argue that its effect is limited.⁹⁻¹¹ In addition, systematic reviews on spinal surgery-related complications in patients with RA often integrate cervical and lumbar lesion outcomes.^{12,13} Given the distinct pathologies of cervical and lumbar spine lesions, a separate analysis should be conducted.

Therefore, we aimed to elucidate the association between RA and surgical outcomes by conducting a systematic review and meta-analysis focused on lumbar lesions. Our objective was to determine the incidence of reoperation, surgical site infection (SSI), and other complications after surgery in patients with RA compared with those without RA. This study aimed to contribute to the development of treatment strategies to mitigate the risks in patients with RA undergoing lumbar spine surgery.

Methods

Study Reporting and Protocol Registration

This review was conducted in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines (Supplemental Appendix S1).¹⁴ The requirement for informed consent from the study participants was waived because of the study design. The study protocol was registered on the Open Science Framework (https://osf. io/5t7jp/).

Study Eligibility

The review question posed was as follows: Does RA affect the clinical outcomes (reoperations and complications) of patients

with lumbar spinal disorders who underwent lumbar spine surgery? We included all observational studies that reported clinical outcomes after lumbar spine surgery for lumbar spinal disorders in patients with RA (exposure) and those without RA (control) (≥18 years), including randomized controlled trials, prospective and retrospective cohort studies, casecontrol studies, and cross-sectional studies. Non-English language studies were included if appropriate translations could be obtained. Lumbar spinal disorders included degenerative conditions, such as spinal stenosis, disc herniation, spondylolisthesis, and spondylolysis, as well as inflammatory conditions associated with RA activity, such as synovitis and erosive changes at the facet joints and discovertebral junctions. The types of surgery included decompression surgery (laminectomy, discectomy, and laminoplasty) and lumbar fusion surgery (intervertebral fusion using any approach and posterior lateral fusion without a dynamic stabilization procedure). RA was defined based on the criteria used in each original study, including the American College of Rheumatology criteria, the European League Against Rheumatism criteria, and database codes such as the International Classification of Diseases. We excluded case series and studies that included patients who underwent prior lumbar spine surgery, long spinal fusion from the pelvis to thoracic spine, or spinal osteotomy, as well as patients diagnosed with ankylosing spondylitis.

Outcome of Interest

The primary outcomes assessed in this study were reoperation (survival outcomes) and SSI after lumbar surgery. For survival outcomes, odds ratios (OR) were calculated using the number of events observed in the RA and control groups. Hazard ratios (HR) were calculated if relevant data were available. Reoperation incidence was measured over the entire follow-up period. However, if data were available at several time points, the data closest to 2 years postoperatively were adopted. For SSI, we used the number of event data points. The definition of SSI was based on original studies, using database codes and clinical diagnoses. We reported outcome measures that were adjusted for potential confounding variables over the reported estimates that were not adjusted for potential confounding. The secondary outcome was the incidence of all adverse events related to the index surgery reported in the original study.

Data Screening and Extraction

We performed a comprehensive search of MEDLINE, Embase, the Cochrane Library, and the International Clinical Trials Registry Platform from inception until August 4, 2023. The database-specific search strategies are presented in Supplemental Appendix S2. Two independent reviewers (SH and MS) screened the titles and abstracts of the listed articles. Subsequently, they reviewed the complete texts of potentially included articles. Other relevant studies were identified by manually searching the reference lists of the included studies. Data were extracted by independent reviewers using a prepiloted data extraction form. Any disagreements were resolved by consensus. We attempted to contact the authors of the included studies to clarify any relevant information or request additional data as necessary.

Quality Assessment

The quality of each included article was assessed by two reviewers using the Quality in Prognosis Studies (QUIPS) assessment tool (Supplemental Appendix S3).¹⁵ To assess the confounding domain of the study, we predefined the following clinically relevant confounders: age, sex, body mass index,¹⁶ smoking,¹⁷ osteoporosis,¹⁸ and comorbidity.¹⁷ If all these confounders were adjusted for in the original study, we rated the study as having a low risk of bias in the confounding domain. The overall risk of bias was rated low if all QUIPS domains were rated low, moderate if one or two domains had an unclear or high risk of bias. During the entire review process, disagreements between the two reviewers were resolved through discussions.

Data Synthesis

Statistical analyses were performed using R (version 4.1.2; R Foundation for Statistical Computing, Vienna, Austria) and RevMan for Macintosh (version 5.4; The Nordic Cochrane Centre, The Cochrane Collaboration, Copenhagen, Denmark). Heterogeneity among the studies was quantified using I2 statistics. We considered an I2 value of <25% to represent low heterogeneity and an I2 value of >75% to indicate high heterogeneity.¹⁹ The Mantel-Haenszel method with a random-effects model was used to calculate the pooled ORs for reoperation and SSI. The inverse variance method with a random-effects model was used to calculate the pooled HRs for reoperation-free survival. If the author did not report the coefficients and variance of the Cox regression model, the HR and variance were estimated using the *P*-value of the log-rank test.²⁰ Forest plots were constructed to illustrate the outcomes of each study. To elucidate the influence of effect modifiers on the results, we conducted subgroup analyses of the primary outcomes according to the type of surgery: fusion surgery vs decompression surgery. If data were insufficient, data were qualitatively reviewed. Publication bias was not assessed using funnel plots and the Egger's test because fewer than 10 studies were included for each outcome. Statistical significance was set at P < 0.05.

Results

Study Characteristics and Patient Demographics

After eliminating duplicates, we screened 384 studies. Subsequently, we incorporated seven studies following the application of the exclusion criteria, as illustrated in Figure 1. Table 1 summarizes the characteristics of the included articles.9-11,21-24 All the included studies were retrospective cohort studies. The seven studies included 72,969 patients with lumbar spinal disorders who underwent lumbar spinal surgery, comprising 7518 (10%) patients with RA and 65,451 (90%) patients without RA. The mean or median age among the analyzed patients with RA ranged from 56.4 to 68.2 years, whereas that of patients without RA varied from 54.2 to 67.7 years. Among the seven studies, two included patients undergoing decompressive surgery, while the remaining five focused on patients after fusion surgery: three involved posterolateral fusion, one pertained to lumbar interbody fusion, and one included any type of fusion surgery. Table 2 shows that all studies had a moderate risk of bias, mainly due to insufficient study attrition and unadjusted confounders.

Reoperation

Five studies reported survival outcomes. Of these, three studies showed that the crude number of reoperations comprising 290 of 2292 patients with RA and 1000 of 11365 patients without RA at their mean follow-up ranged from 3 to 9.5 years.^{10,22,23} All the patients in these studies underwent lumbar fusion surgery. A statistically significant difference was observed in favor of patients without RA (OR: 5.57; 95%) confidence interval [CI]: 1.10-28.26; I2 = 92%; P = 0.04, moderate risk of bias; Figure 2A). Among the five studies, two reported unadjusted survival data. One study only reported the result of the log-rank test at a follow-up of 5 years²⁴; thus, the log HR was estimated from the P-value of the log-rank test. Another study reported the log-rank statistic at a 2-year follow-up,²¹ so we calculated the log HR using these values, following the recommendations of the Cochrane Handbook.² No statistically significant difference was found in survival data (HR: 1.15; 95% CI: 0.94-1.40; I2 = 100%; P = 0.16, moderate risk of bias; Figure 2B). We did not perform sensitivity or subgroup analyses because of the small size of the included studies.

SSI

Six studies reported the incidence of SSI. Figure 3 shows a statistically significant difference in favor of patients without RA in total samples (OR: 1.47; 95% CI: 1.28-1.69; I2 = 2%; P < 0.01, moderate risk of bias). The pooled ORs for both surgical procedures showed a trend toward fewer SSIs in patients without RA. A subgroup analysis of the types of surgery was performed, and the test for subgroup differences



Figure 1. Flow diagram. ICTRP: International Clinical Trials Registry Platform.

revealed minimal heterogeneity between the types of surgery (P = 0.52).

Other Complications

Table 3 shows details of complications from the included studies. Six studies reported on complications. The reported prevalence of complications in patients with RA ranged from 13.5% to 57%, compared with 17.1% to 36% in patients without RA. Four studies performed statistical analyses of complication rates between groups: one study found no statistical difference in complication rates,⁹ whereas three other studies found a predominance of complications in patients with RA.^{11,21,24}

Discussion

Lumbar spine lesions in patients with RA have been previously reported.^{1,2} However, evidence of the clinical outcomes following spine surgery is lacking. Our meta-analysis included a large sample of 72,969 patients, 10% of whom (7518 patients) were diagnosed with RA. These findings indicate that patients with RA had higher odds of undergoing reoperation and experienced more postoperative complications. Spine surgeons should properly understand the risks associated with lumbar spine surgery in patients with RA and perform comprehensive perioperative management to improve clinical outcomes.

Reoperation

The finding of a higher incidence of reoperation in patients with RA, indicated by a pooled OR of 5.57 (95% CI: 1.10-28.26), suggests that patients with RA were more likely to undergo reoperation than those without RA. The significant heterogeneity (I2 = 92%) in these results may reflect variations in surgical technique, perioperative care, and patient selection across the included studies. However, the consistently higher odds across these studies cannot be disregarded. Conversely, the HR for reoperation-free survival did not show a statistically significant difference (HR: 1.15, 95% CI: 0.94-1.40), which might suggest that the long-term risk of reoperation may not be as pronounced as initially indicated by the ORs. One of the two studies that provided survival data had a 5-year

Table I. Characteristics of the Included Studies.

				Patients with RA Patients without RA		- /					
Study	Setting	Country	n	Age, year	Female (%)	n	Age, year	Female (%)	Type of Surgery	Follow-up, year	Risk of Bias
Gulati, 2016 ⁹	Clinical registry	Norway	37	68.2 ± 6.9	28 (75.7)	1396	67.7 ± 10	675 (48.4)	Laminectomy	I	Moderate
Ratnasamy, 2023 ²⁴	Claims database	USA	2149	56.4 ± 11.7	1368 (63.7)	8485	56.4 ± 11.8	5398 (63.6)	Discectomy	5	Moderate
Crawford, 2008 ¹⁰	Single center	USA	19	64	17 (89)	19	65	18 (94)	PLF	RA:2, nonRA:2.2	Moderate
Kang, 2016 ¹¹	Single center	Korea	40	64.3 (44-77)	39 (97)	134	65.3 (49-79)	117 (87)	PLF	RA:3, nonRA:3.2	Moderate
Bell, 2021 ²¹	Claims database	USA	3021	NA	2227 (73.7)	44186	NA	32714 (74)	PLF	NA	Moderate
Thiebaut, 2021 ²²	Single center	France	13	57.4	9 (69)	36	54.2	22 (61)	LIF	RA:9.5, nonRA:9.4	Moderate
Park, 2022 ²³	Claims database	Korea	2239	64.5±8.8	603(26.9)	11195	64.5±8.8	3033(27.1)	Fusion surgery	NA	Moderate

PLF, posterolateral fusion; LIF, lateral interbody fusion; NA, not applicable.

Table 2. Risk of Bias Assessment Using Quality in Prognosis Studies (QUIPS) Assessment Tool.

Study	Study Participation	Study Attrition	Prognostic Factor Measurement	Outcome Measurement	Study Confounding	Statistical Analysis and Reporting
Gulati, 2016 ⁹	Moderate	High	Low	Low	High	Low
Ratnasamy, 2023 ²⁴	Moderate	High	Low	Low	Moderate	Low
Crawford, 2008 ¹⁰	High	High	Low	Low	Moderate	Low
Kang, 2016 ¹¹	High	High	Low	Low	Moderate	Low
Bell, 2021 ²¹	Moderate	High	Low	Low	Moderate	Low
Thiebaut, 2021 ²²	Moderate	High	Low	Low	High	Low
Park, 2022 ²³	Moderate	High	Low	Low	Moderate	Low

А								
	RA		Cont	trol		Odds Ratio		Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl		M-H, Random, 95% CI
Kang 2016	15	40	12	134	34.8%	6.10 [2.55, 14.60]]	
Park 2022	265	2239	985	11195	38.5%	1.39 [1.20, 1.61]]	•
Thiebaut 2021	10	13	3	36	26.7%	36.67 [6.37, 210.94]]	
Total (95% CI)		2292		11365	100.0%	5.57 [1.10, 28.26]]	
Total events	290		1000					
Heterogeneity: Tau ² =	1.78; Cl	$ni^2 = 23$	3.77, df	= 2 (P <	0.00001)	; $I^2 = 92\%$	0 002	0 1 1 10 500
Test for overall effect:	Z = 2.02	7 (P = C)	0.04)				0.002	Favours [RA] Favours [control]
R								
Ъ								
						Hazard Ratio		Hazard Ratio
Study or Subgroup	log[H	azard	Ratio]	SE	Weight	IV, Random, 95% CI		IV, Random, 95% CI
Bell 2021		0	.2404	0.005	50.0%	1.27 [1.26, 1.28]		
Ratnasamy 2023		0	.0397	0.0006	50.0%	1.04 [1.04, 1.04]		•
Total (95% CI)					100.0%	1.15 [0.94, 1.40]		•
Heterogeneity: Tau ²	= 0.02;	Chi ² =	1588.3	5, $df = 1$	(P < 0.0	0001); $I^2 = 100\%$	- 0.1	
Test for overall effect	t: Z = 1.	40 (P =	0.16)				0.01	U.I I IO IOO

Figure 2. Comparison between patients with rheumatoid arthritis (RA) and patients without RA (control). Outcome: reoperation. (A) The odds ratios (ORs) were determined using the Mantel–Haenszel method with a random effects model. (B) SE indicates standard error. The hazard ratios (HRs) were determined using a random effects model weighted by the inverse variance estimate. Square data markers represent ORs or HRs, with marker size reflecting the statistical weight of the study. Horizontal lines represent 95% confidence intervals (Cls). Diamond represents overall ORs or HRs, and 95% Cls for the outcome of interest.

	RA		Cont	rol		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	M-H, Random, 95% Cl
3.1.1 Decompression	Í.				0000		
Gulati 2016	0	37	50	1396	0.2%	0.36 [0.02, 5.87]	
Ratnasamy 2023 Subtotal (95% CI)	103	2149 2186	271	8485 9881	33.3% 33.5%	1.53 [1.21, 1.92] 1.47 [0.95, 2.28]	
Total events	103		321				
Heterogeneity: Tau ² =	0.04; Cl	$ni^2 = 1.$	04, df =	1 (P = 0	.31); I ² =	3%	
Test for overall effect:	Z = 1.74	4 (P = 0)	0.08)				
3.1.2 Fusion							
Bell 2021	171	3021	1787	44186	65.1%	1.42 [1.21, 1.67]	
Crawford 2008	2	19	1	19	0.3%	2.12 [0.18, 25.55]	
Kang 2016	4	40	4	134	0.9%	3.61 [0.86, 15.15]	
Thiebaut 2021	2	13	0	36	0.2%	15.87 [0.71, 355.05]	
Subtotal (95% CI)		3093		44375	66.5%	1.93 [0.97, 3.87]	◆
Total events	179	2	1792				
Heterogeneity: $Tau^2 =$	0.17; C	$1i^2 = 3.$	97, df =	3 (P = 0)	.26); $I^2 =$	24%	
Test for overall effect:	Z = 1.86	5 (P = 0)	0.06)				
Total (95% CI)		5279		54256	100.0%	1.47 [1.28, 1.69]	•
Total events	282		2113				
Heterogeneity: Tau ² =	0.00; Cl	$ni^2 = 5.$	08, df =	5 (P = 0)	.41); $I^2 =$	2%	
Test for overall effect:	Z = 5.53	3 (P < 0	0.00001)				Favours [RA] Favours [control]
Test for subgroup diff	erences:	$Chi^2 =$	0.42, df	= 1 (P =	0.52), I ²	= 0%	. arous [ing rations [control]

Figure 3. Comparison between patients with rheumatoid arthritis (RA) and patients without RA (control). Outcome: surgical site infection. The odds ratios (ORs) were determined using the Mantel–Haenszel (M-H) method with a random effects model. Square data markers represent ORs, with marker size reflecting the statistical weight of the study. Horizontal lines represent 95% confidence intervals (CIs). Diamond represents subgroup and overall ORs, and 95% CIs for the outcome of interest.

Table 5. Details of Complications Following Surger	ils of Complications Followir	lications following	Surgery
--	-------------------------------	---------------------	---------

	Turne of		Patients with RA	Patients without RA			
Study	Surgery	n (%)	Details, n (%)	n (%)	Details, n (%)		
Gulati, 2016 ⁹	Laminectomy	5 (13.5)	Dural tear 3 (8.1); UTI 3 (8.1)	179 (12.8)	Dural tear 59 (4.2); nerve injury 1 (0.1); blood replacement 8 (0.6); cardiovascular complications 2 (0.1); anaphylactic reaction 2 (0.1); wrong level 1 (0.1); SSI 50 (3.6); UTI 58 (4.2); pneumonia 10 (0.7); PE 2 (0.1); DVT 1 (0.1)		
Ratnasamy, 2023 ²⁴	Discectomy	686 (32)	PE 29 (1.3); SSI 103 (4.8); DVT 84 (3.9); sepsis 93 (4.3); MI 40 (1.9); wound dehiscence 53 (2.5); hematoma 37 (1.7); AKI 107 (5); transfusion 38 (1.8); UTI 300 (14); pancreatitis 20 (0.9); pneumonia 170 (7.9)	1086 (12.8)	PE 80 (0.9); SSI 271 (3.2); DVT 117 (1.4); sepsis 100 (1.2); MI 32 (0.4); wound dehiscence 113 (1.3); hematoma 75 (0.9); AKI 155 (1.8); transfusion 47 (0.6); UTI 375 (4.4); pancreatitis 15 (0.2); pneumonia 134 (1.6)		
Crawford, 2008 ¹⁰	PLF	(57)	SSI 2; nonunion 2; implant complications 2; ASD 6	7 (36)	SSI I; nonunion 2; implant complications 2; ASD 6		
Kang, 2016 ¹¹	PLF	19 (47.5)	NA	23 (17.1)	NA		
Bell, 2021 ²¹	PLF	NA	Cerebrovascular accident 17 (0.6); sepsis 61 (2); death 3 (0.1); thromboembolic event 69 (2.3); pneumonia 59 (2); AKI 58 (1.9); SSI 171 (5.7)	`NA [´]	Cerebrovascular accident 209 (0.5); MI 167 (0.4); sepsis 382 (0.9); death 79 (0.2); thromboembolic event 726 (1.6); pneumonia 540 (1.2); AKI 656 (1.5); SSI 1787 (4.0)		
Thiebaut, 2021 ²²	LIF	NA	ASD 10 (76.9); pseudarthrosis 3 (23.1); mechanical failure 5 (38.5)	NA	ASD 17 (50); pseudarthrosis 3 (8.8); mechnical failure 19 (55.9)		
Park, 2022 ²³	Fusion surgery	NA	NA	NA	NA		

PLF, posterolateral fusion; LIF, lateral interbody fusion; UTI, urinary tract infection; SSI, surgical site infection; PE, pulmonary embolism; DVT, deep vein thrombosis; MI, myocardial infarction; AKI, acute kidney injury; ASD, adjacent segment degeneration; NA, not applicable.

follow-up and found that patients with RA consistently exhibited higher reoperation rates in the first 2 years, with the survival curve crossing at approximately 2 years. Another study concluded that reoperation-free survival was significantly lower in patients with RA at the 2-year follow-up. These results indicate that reoperation within 2 years of lumbar spine surgery is likely common among patients with RA.

One study that examined outcomes after lumbar fusion surgery identified the reasons for reoperation.¹¹ Among patients with RA, six cases out of 40 reported implant failure. By contrast, implant failure occurred in three out of 134 patients without RA. Patients with RA are more susceptible to osteoporosis and sarcopenia,^{5,26} which may contribute to a higher incidence of reoperation.

SSI

The analysis also highlighted a lower incidence of SSI in patients without RA across all surgical types, demonstrating minimal heterogeneity ($I^2 = 2\%$) and indicating a consistent pattern across studies. Subgroup analysis of surgical techniques showed no statistically significant differences between decompression and fusion surgeries. The incidence of SSI was found to be high in patients with RA undergoing spinal surgery, including procedures on the cervical and thoracic spines.¹³ Similar results were obtained when the data and analysis were restricted to the lumbar spine. These findings could reflect the systemic effects of RA, such as chronic inflammation and the use of immunomodulatory medications.²⁷ These factors could potentially increase susceptibility to infections regardless of the type of surgery performed.

Other Complications

Complications are more prevalent in patients with RA, which is aligned with the systemic nature of RA and its potential to complicate postoperative recovery. The variance in complication rates between patients with RA and those without RA (13.5%-57% vs 12.8%-36%, respectively) also indicates a multifactorial risk profile in patients with RA undergoing spine surgery.

For example, the incidence of dural tears in patients with RA undergoing laminectomy is noteworthy. A study analyzing a database of spinal surgeries for non-cervical lesions also reported that dural tears were more common in patients with RA.²⁸ Although these complications were relatively less frequent in patients without RA, the increased percentage in patients with RA could be attributed to dural adhesions resulting from chronic inflammation. Interestingly, when examining more detailed complications, such as adjacent segment degeneration, the data revealed that these are prevalent postoperative issues, particularly in patients with RA. Magnetic resonance imaging of lumbar lesions in patients with RA indicates that erosion of the lumbar endplates and facets is prevalent.²⁹ Chronic inflammation associated with RA may

contribute to the increased incidence of adjacent segment degeneration. Although managing disease activity is a standard approach for treating RA, particular consideration may be required for patients who have undergone lumbar fusion surgery.

Table 3 also highlights the systemic nature of complications in patients with RA, with a broad spectrum of events ranging from cerebrovascular accidents to sepsis and myocardial infarction. This underscores the necessity for a multidisciplinary approach to perioperative care in patients with RA, encompassing not only surgical intervention but also the management of potential cardiovascular and thromboembolic complications.

Practical Considerations for Clinical Management

Based on our meta-analytic findings and previous reports, several practical considerations for managing patients with RA undergoing lumbar spine surgery are suggested. Preoperative medical optimization is essential: careful control of RA disease activity and comorbidities (e.g., chronic obstructive pulmonary disease, obesity, diabetes) can help reduce perioperative complications.³⁰ The decision to continue, modify, or temporarily discontinue biological disease-modifying antirheumatic drugs and/or glucocorticoids should be made in collaboration with rheumatologists, weighing the risk of RA flare against the risk of SSI. From a surgical standpoint, a meticulous technique to minimize tissue damage and operative time is crucial, especially given the predisposition of patients with RA to wound-healing problems. Extended postoperative antibiotic prophylaxis, although debated, may be considered in selected cases to mitigate infection risks. In prosthetic joint surgery, a 7-day extended course of antimicrobial therapy after surgery has been shown to reduce the incidence of periprosthetic joint infections in high-risk patients, including those with RA.³¹ Nutritional optimization addressing hypoalbuminemia also appears beneficial for this group. Finally, close postoperative monitoring, including regular imaging and early intervention when implant loosening or infection is suspected, is paramount to reducing the likelihood of reoperation. These recommendations, although not yet backed by high-level evidence specific to patients with RA, may help guide more tailored, multidisciplinary care aimed at improving surgical outcomes.

Limitations

Although this meta-analysis offers important insights, it has several limitations. First, all the included studies were retrospective and with a moderate risk of bias. Most studies adjusted for confounders through patient matching, but they could not address all our pre-defined confounders. Future prospective cohort studies addressing these issues are necessary to confirm our findings. Second, the high heterogeneity observed in the data regarding reoperation and survival necessitates careful interpretation of the results. Nonetheless, the consistent direction of the effect sizes across studies suggests that RA may influence the primary outcome. Third, no studies have addressed disease activity in RA, and the effect of RA on postoperative outcomes has not been considered. A comparative study of patients with RA with and without well-controlled disease activity could clarify the underlying mechanisms.

Conclusion

This meta-analysis demonstrated an association between RA and increased postoperative complications, including reoperation and SSI, in patients undergoing lumbar spine surgery. These results underscore the importance of comprehensive perioperative management for improving clinical outcomes.

Acknowledgments

We would like to thank Editage (https://www.editage.jp) for English language editing.

Author Contributions

Shintaro Honda, Akihiro Shiroshita and Natsumi Saka contributed to the study conception and design. Material preparation and data collection were performed by Shintaro Honda, Masaki Sakamoto and Akihiro Shiroshita. The first draft of the manuscript was written by Shintaro Honda Akihiro Shiroshita and Natsumi Saka, and all authors commented on previous versions of the manuscript. All authors read and approved the final manuscript. All authors certify that they have no affiliations with or involvement in any organization or entity with any financial interest or non-financial interest in the subject matter or materials discussed in this manuscript.

Declaration of Conflicting Interests

The author(s) declared the following potential conflicts of interest with respect to the research, authorship, and/or publication of this article: The Department of Advanced Medicine for Rheumatic Diseases is supported by Nagahama City, Shiga, Japan; Toyooka City, Hyogo, Japan; and two pharmaceutical companies (Asahi Kasei Pharma Corp., and AYUMI Pharmaceutical Co.). The stated companies had no role in the study design, data collection or analysis, manuscript writing, or the decision to submit the manuscript for publication. KM received speaking and/or consulting fees from AbbVie GK, Eisai Co., Ltd., Pfizer Inc., Chugai Pharmaceutical Co., Ltd., Mitsubishi Tanabe Pharma Corporation, Pfizer Inc., Bristol-Myers Squibb, Daiichi Sankyo Co. Ltd., and Asahi Kasei Pharma Corp. SMatsuda received research grants and/or speaker fees from Astellas Pharma Inc., Daiichi Sankyo Co. Ltd., Mitsubishi Tanabe Pharma Corporation, Eisai Co., Ltd., Takeda Pharmaceutical Company Limited, Chugai Pharmaceutical Co. Ltd, Pfizer Inc., and Asahi Kasei Corporation.

Funding

The author(s) received no financial support for the research, authorship, and/or publication of this article.

Ethical Statement

Ethical Approval

This study did not involve any experiments on humans or animals requiring ethical approval, as it is a systematic review.

Informed Consent

The requirement for informed consent from the study participants was waived because of the study design.

ORCID iDs

Shintaro Honda b https://orcid.org/0000-0001-9387-7730 Takayoshi Shimizu b https://orcid.org/0000-0002-2683-0489 Soichiro Masuda b https://orcid.org/0000-0003-4770-0728

Supplemental Material

Supplemental material for this article is available online.

References

- Lawrence JS, Sharp J, Ball J, Bier F. Rheumatoid arthritis of the lumbar spine. *Ann Rheum Dis*. 1964;23(3):205. doi:10.1136/ard. 23.3.205
- Sims-Williams H, Jayson MI, Baddeley H. Rheumatoid involvement of the lumbar spine. *Ann Rheum Dis.* 1977;36(6): 524. doi:10.1136/ard.36.6.524
- Kawaguchi Y, Matsuno H, Kanamori M, Ishihara H, Ohmori K, Kimura T. Radiologic findings of the lumbar spine in patients with rheumatoid arthritis, and a review of pathologic mechanisms. J Spinal Disord Tech. 2003;16(1):38-43. doi:10.1097/ 00024720-200302000-00007
- Miura K, Morita O, Hirano T, et al. Prevalence of and factors associated with dysfunctional low back pain in patients with rheumatoid arthritis. *Eur Spine J.* 2019;28(5):976-982. doi:10. 1007/s00586-019-05938-x
- Senosi MR, Fathi HM, Baki NMA, Zaki O, Magdy AM, Gheita TA. Bone mineral density, vitamin D receptor (VDR) gene polymorphisms, fracture risk assessment (FRAX), and trabecular bone score (TBS) in rheumatoid arthritis patients: connecting pieces of the puzzle. *Clin Rheumatol.* 2022;41(5): 1333-1342. doi:10.1007/s10067-022-06048-8
- Masamoto K, Otsuki B, Fujibayashi S, et al. Factors influencing spinal sagittal balance, bone mineral density, and Oswestry Disability Index outcome measures in patients with rheumatoid arthritis. *Eur Spine J.* 2018;27(2):406-415. doi:10.1007/s00586-017-5401-3
- Honda S, Murata K, Fujibayashi S, et al. Effect of high disease activity on spinal sagittal malalignment in patients with rheumatoid arthritis. *J Neurosurg Spine* 2024;41(1):24-32. doi:10. 3171/2024.1.SPINE231199

- Joo P, Ge L, Mesfin A. Surgical management of the lumbar spine in rheumatoid arthritis. *Global Spine J.* 2020;10(6):767-774. doi:10.1177/2192568219886267
- Gulati A, Solberg T, Giannadakis C, et al. Surgery for lumbar spinal stenosis in patients with rheumatoid arthritis: a multicenter observational study. *European J Rheumatology*. 2016; 3(2):56-60. doi:10.5152/eurjrheum.2016.15070
- Crawford CH, Carreon LY, Djurasovic M, Glassman SD. Lumbar fusion outcomes in patients with rheumatoid arthritis. *Eur Spine J*. 2008;17(6):822-825. doi:10.1007/s00586-008-0610-4
- Kang CN, Kim CW, Moon JK. The outcomes of instrumented posterolateral lumbar fusion in patients with rheumatoid arthritis. *Bone Jt J.* 2016;98-B(1):102-108. doi:10.1302/0301-620x.98b1.36247
- Streufert BD, Onyedimma C, Yolcu YU, et al. Rheumatoid arthritis in spine surgery: a systematic review and meta-analysis. *Glob Spine* J. 2022;12(7):1583-1595. doi:10.1177/21925682211057543
- Zhang S, Wang L, Bao L, et al. Does rheumatoid arthritis affect the infection and complications rates of spinal surgery? A systematic review and meta-analysis. *World Neurosurg*. 2021; 145:260-266. doi:10.1016/j.wneu.2020.09.039
- Page MJ, McKenzie JE, Bossuyt PM, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *BMJ*. 2021;372:n71. doi:10.1136/bmj.n71
- Hayden JA, van dervWindt DA, Cartwright JL, Côté P, Bombardier C. Assessing bias in studies of prognostic factors. *Ann Intern Med.* 2013;158(4):280. doi:10.7326/0003-4819-158-4-201302190-00009
- Marquez-Lara A, Nandyala SV, Sankaranarayanan S, Noureldin M, Singh K. Body mass index as a predictor of complications and mortality after lumbar spine surgery. *Spine*. 2014;39(10): 798-804. doi:10.1097/brs.00000000000232
- Soroceanu A, Burton DC, Oren JH, et al. Medical complications after adult spinal deformity surgery. SPINE. 2016;41(22): 1718-1723. doi:10.1097/brs.00000000001636
- Hikata T, Ishii K, Matsumoto M, et al. Risk factor for poor patient satisfaction after lumbar spine surgery in elderly patients aged over 80 years. *Clin Spine Surg.* 2021;34(4):E223-E228. doi:10.1097/bsd.00000000001101
- Higgins JPT, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses. *BMJ*. 2003;327(7414):557. doi: 10.1136/bmj.327.7414.557
- Tierney JF, Stewart LA, Ghersi D, Burdett S, Sydes MR. Practical methods for incorporating summary time-to-event data into meta-analysis. *Trials*. 2007;8(1):16. doi:10.1186/1745-6215-8-16

- Bell J, Sequeira S, Kamalapathy P, Puvanesarajah V, Hassanzadeh H. Rheumatoid arthritis increases risk of medical complications following posterior lumbar fusion. *World Neurosurg*. 2021;149: e729-e736. doi:10.1016/j.wneu.2021.01.110
- Thiébaut B, Bastard C, Eymard F, Bouthors C, Lachaniette CHF, Dubory A. Long-term results of anterior-only lumbar interbody fusions in rheumatoid arthritis patients: comparative retrospective cohort study. *World Neurosurg*. 2021;154:e109-e117. doi:10.1016/j.wneu.2021.06.128
- Park JS, Park SJ, Park J, Shin G, Hong JY. The influence of rheumatoid arthritis on higher reoperation rates over time following lumbar spinal fusion—a nationwide cohort study. *J Clin Medicine*. 2022;11(10):2788. doi:10.3390/jcm11102788
- Ratnasamy PP, Rudisill KE, Gouzoulis MJ, Kammien AJ, Grauer JN. Rheumatoid arthritis patients are at increased risk for adverse events following lumbar discectomy. *Spine J* 2023; 23(7):990-996. doi:10.1016/j.spinee.2023.03.012
- Simmonds MC, Tierney J, Bowden J, Higgins JPT, et al. Metaanalysis of time-to-event data: a comparison of two-stage methods. *Res Synth Methods*. 2011;2(3):139-149. doi:10.1002/jrsm.44
- Torii M, Hashimoto M, Hanai A, et al. Prevalence and factors associated with sarcopenia in patients with rheumatoid arthritis. *Mod Rheumatol.* 2019;29(4):589-595. doi:10.1080/14397595. 2018.1510565
- Smolen JS, Aletaha D, McInnes IB. Rheumatoid arthritis. *Lancet*. 2016;388(10055):2023-2038. doi:10.1016/s0140-6736(16)30173-8
- Bernstein DN, Kurucan E, Menga EN, Molinari RW, Rubery PT, Mesfin A. Comparison of adult spinal deformity patients with and without rheumatoid arthritis undergoing primary non-cervical spinal fusion surgery: a nationwide analysis of 52,818 patients. *Spine J.* 2018;18(10):1861-1866. doi:10.1016/j.spinee.2018.03.020
- Yamada K, Suzuki A, Takahashi S, et al. MRI evaluation of lumbar endplate and facet erosion in rheumatoid arthritis. *J Spinal Disord Tech*. 2014;27(4):E128-E135. doi:10.1097/bsd. 0b013e3182a22a34
- Ito H, Murata K, Sobue Y, et al. Comprehensive risk analysis of postoperative complications in patients with rheumatoid arthritis for the 2020 update of the Japan College of Rheumatology clinical practice guidelines for the management of rheumatoid arthritis. *Mod Rheumatol.* 2021;32(2):296-306. doi:10.1080/ 14397595.2021.1913824
- Kheir MM, Dilley JE, Ziemba-Davis M, Meneghini RM. The AAHKS clinical research award: extended oral antibiotics prevent periprosthetic joint infection in high-risk cases: 3855 patients with 1-year follow-up. J Arthroplast. 2021;36(7): S18-S25. doi:10.1016/j.arth.2021.01.051