

# Change in Functional Impairment, Disability, and Quality of Life Following Operative **Treatment for Degenerative Cervical Myelopathy: A Systematic Review and Meta-Analysis**

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#### Abstract

Study Design: Systematic review.

Objectives: The primary objective of this systematic review was to define the change in impairment, disability, and pain following surgical intervention in patients with degenerative cervical myelopathy (DCM). Secondary objectives included to assess the impact of preoperative disease severity and duration of symptoms on outcomes and to summarize complications associated with surgery.

Methods: A systematic literature search was conducted to identify prospective studies evaluating the effectiveness and safety of operative treatment in patients with DCM. Outcomes of interest were functional status, disability, pain, and complications. The quality of each study was evaluated using the Newcastle-Ottawa Scale, and the strength of the overall body of evidence was rated using guidelines outlined by the Grading of Recommendation Assessment, Development and Evaluation (GRADE) Working Group.

**Results:** Of the 385 retrieved citations, 32 met inclusion criteria and are summarized in this review. Based on our results, pooled standard mean differences showed a large effect for improvement in Japanese Orthopaedic Association or modified Japanese Orthopaedic Association score from baseline at short-, medium-, and long-term follow-up: 6 to 12 months (1.92; 95% confidence interval [CI] = 1.41 to 2.43), 13 to 36 months (1.40; 95% CI = 1.12 to 1.67), and  $\geq$ 36 months (1.92; 95% CI = 1.14 to 2.69) (moderate evidence). Surgery also resulted in significant improvements in Nurick, Neck Disability Index, and Visual Analogue Scale scores (low to very low evidence). The cumulative incidence of complications was low (14.1%; 95% CI = 10.1% to 18.2%).

Conclusion: Surgical intervention for DCM results in significant improvements in functional impairment, disability, and pain and is associated with an acceptably low rate of complications.

#### **Keywords**

degenerative cervical myelopathy, surgery, outcomes, complications, severity

# Introduction

Degenerative disease of the cervical spine is the most common indication for cervical spine surgery.<sup>1,2</sup> Neurological consequences of such disease include radiculopathy and myelopathy, which result in progressive pain, disability, and diminished quality of life. The utility of nonoperative therapy for patients with degenerative cervical myelopathy (DCM) was previously investigated by Rhee et al<sup>3</sup>; this systematic review revealed a paucity of evidence supporting such management and

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recommended against nonoperative treatment in patients with moderate to severe disease.

When indicated, the goals of surgical intervention include decompressing the spinal cord, stabilizing the spinal column, and, if necessary, realigning the sagittal plane. The techniques required to accomplish these goals remain controversial and are often selected based on the nature of the underlying pathology, clinical presentation, the presence of sagittal plane deformity, and other patient and surgeon factors.

Despite the frequent application of surgery for this disease, there is a paucity of high-quality data supporting operative management in patients with DCM.<sup>4</sup> An underpowered randomized controlled trial provided evidence that outcomes in patients with mild to moderate myelopathy do not differ between conservative and surgical treatment arms.<sup>5,6</sup> Conversely, other observational cohort studies with large sample sizes reveal superior neurological recovery following surgical intervention compared to conservative management.<sup>7,8</sup>

The primary objective of this study was to define the change in impairment, disability, and pain following surgical intervention in patients with DCM. Secondary objectives included to evaluate the impact of preoperative disease severity and duration of symptoms on outcomes and to summarize treatment complications associated with surgery. Specifically, key questions for this systematic review were the following:

**Key Question 1:** What are the expected functional, disability, and pain outcomes following surgical intervention for DCM?

**Key Question 2:** Do these expected outcomes depend on preoperative disease severity or duration of symptoms?

**Key Question 3:** What are the complications associated with surgical intervention?

## **Materials and Methods**

# Electronic Literature Search

A systematic review of the literature was conducted to identify relevant articles published up to May 18, 2015. Electronic databases (PubMed, EMBASE, Cochrane Collaboration library) and reference lists of key articles were searched to identify prospective studies on adult patients with DCM (cervical spondylotic myelopathy or ossification of the posterior longitudinal ligament [OPLL]) that evaluated outcomes or safety of operative treatment. For Key Question 1, we included studies that assessed functional, disability, and quality of life outcomes following surgical intervention. For Key Question 2, we identified studies that evaluated the impact of preoperative disease severity and duration of symptoms on postoperative outcomes. For Key Question 3, we searched for studies that summarized complications associated with surgery. Table 1 displays the PICO table and highlights the inclusion and exclusion criteria for this review. Studies were excluded if fewer than 50 patients were included, the follow-up was less than 1

year, arthroplasty procedures were performed, and the outcome metrics did not include modified Japanese Orthopaedic Association (mJOA), Neck Disability Index (NDI), or Nurick scores. For studies comparing operative and nonoperative outcomes, we only extracted data from the surgical arm. For studies that compared different types of surgery (eg, anterior vs posterior), we combined the results from both approaches. Studies that did not report primary outcomes but did analyze complications were included for Key Question 3. Two investigators (EDB, JRD) reviewed the full texts of potential articles to obtain a final collection of studies.

#### Data Extraction

We extracted the following data from the included articles: study design, patient demographics, diagnosis, surgical approach, preoperative and postoperative neurological status (mJOA, NDI, Nurick, Visual Analog Scale [VAS] scores), and complications. Preoperative disease severity and duration of symptoms were recorded when available. Each citation was independently reviewed by 2 of the authors.

#### Study Quality and Overall Strength of Body of Literature

The risk of bias was appraised by 2 of the study authors using the modified Newcastle-Ottawa Scale.<sup>9</sup> The evaluation for risk of bias is included in the supplemental material (available in the online version of the article).

Next, the overall quality of evidence with respect to each outcome was determined based on methodology outlined by the Grades of Recommendation Assessment, Development and Evaluation (GRADE) Working Group<sup>10,11</sup> and recommendations from the Agency for Healthcare Research and Quality.<sup>12,13</sup>

The overall body of evidence is considered "Low" if all studies are observational. The quality of the body of evidence may be upgraded or downgraded depending on a number of factors. Criteria for downgrading 1 or 2 levels include (1) inconsistency of results, (2) indirectness of evidence, or (3) imprecision of the effect estimates (eg, wide variance). Alternately, the body of evidence may be upgraded 1 or 2 levels based on (1) large magnitude of effect or (2) doseresponse gradient.

A quality level of "High" indicates high confidence that the true effect lies close to the estimate of effect. A "Moderate" quality level reflects moderate confidence in the effect estimate; the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different. A "Low" quality level represents limited confidence in the effect estimate; the true effect may be substantially different from the estimate of the effect.<sup>11</sup> "Very Low" ratings indicate very little confidence in the effect estimate; the true effect is likely to be substantially different from the estimate of effect. This rating may be used if there is no evidence or if it is not possible to estimate an effect.

Table I. Inclusion and Exclusion Criteria.

	Inclusion	Exclusions
Patient	- Adult patients ( $\geq\!\!18$ years) with CSM or cervical myelopathy caused by OPLL	<ul> <li>Patients &lt;18 years of age</li> <li>Myelopathy due to infection, malignancy or inflammatory arthritis</li> </ul>
Intervention	<ul> <li>Surgical decompression with or without fusion (eg, ACDF, laminoplasty, laminectomy, and fusion)</li> </ul>	<ul> <li>≤50 patients analyzed (after loss-to-follow up)</li> <li>Cervical disc arthroplasty</li> </ul>
Comparison	Impairment at baseline	Competing interventions
Prognostic factors	<ul><li>Myelopathy severity</li><li>Symptom duration</li></ul>	<ul> <li>Factors other than myelopathy severity or symptom duration</li> </ul>
Outcomes	<ul> <li>Primary <ul> <li>JOA or mJOA</li> <li>NDI</li> <li>Nurick</li> </ul> </li> <li>Secondary <ul> <li>Pain scores (arm and neck pain)</li> <li>30-meter walk test</li> </ul> </li> <li>Complications/harms <ul> <li>If a study does not report any of the above primary outcomes but presents data on complications, the study will be included for the question on safety</li> </ul> </li> </ul>	<ul> <li>JOA/mJOA, Nurick, or NDI not reported for effectiveness</li> </ul>
Study design	• Prospective studies with baseline and follow-up outcome data or the proportion of patients that improved by a certain amount	<ul> <li>Retrospective studies</li> <li>Studies that do not present change scores from baseline or do not present data to calculate change scores</li> <li>Studies with <i follow="" li="" up<="" year=""> </i></li></ul>

Abbreviations: ACDF, anterior cervical discectomy and fusion; CSM, cervical spondylotic myelopathy; JOA, Japanese Orthopaedic Association score; mJOA, modified Japanese Orthopaedic Association score; NDI, Neck Disability Index; OPLL, ossification of the posterior longitudinal ligament.

# Data Analysis

Outcomes were stratified by follow-up timing as short term (less than 12 months), medium term (13-36 months), and long term (>36 months). Effect measures were first defined as mean differences (MD) with missing standard deviations imputed using the highest value from included studies. Standardized mean differences (SMDs) were computed to reflect the heterogeneity of outcome measures for JOA and mJOA scores that are correlated but not identical.<sup>14</sup> Using Cohen's criteria,<sup>15</sup> an SMD of 0.2 to 0.49 represents a "small" effect, 0.5 to 0.79 a "moderate" effect, and 0.8 or greater a "large" effect. When it was necessary to merge multiple surgical cohorts from a single article, the weighted average scores were calculated and Cohen's approximation<sup>16</sup> for combining standard deviations was used. Results were further stratified by study quality based on risk of bias (high or low) to determine if there were qualitative and quantitative differences.

### Results

#### Study Selection

The search strategy yielded 385 relevant citations, of which 312 were excluded based on their title and/or abstract. Seventy-seven studies were selected for full-text review. Forty-five of these were excluded for the following reasons: retrospective study design, sample included both radiculopathy and myelopathy, desired outcomes or preoperative and

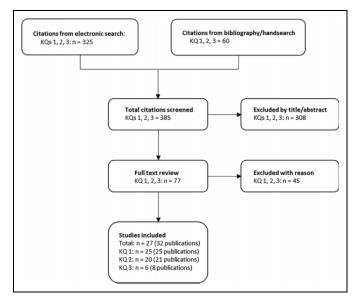


Figure 1. Overview of search strategy.

postoperative comparisons were not available, fewer than 50 patients, duplicated results from a separate study, or follow-up period less than 1 year. An overview of our selection process is summarized in Figure 1. Thirty-two studies ultimately met the inclusion criteria and are summarized in this review.<sup>17-48</sup> Study characteristics are provided in Table 2; sample sizes ranged from 52 to 479 patients, mean ages ranged from 46 to 65 years

Author (Year); Study Design	Operative I reatment	Patient Characteristics <sup>4</sup>	Baseline Severity (Mean)	symptom Duration (Mean)	Follow-up, % Followed	Outcome Measures
Bapat (2008); Prospective case series	Anterior or posterior decompression	N = 129 Mean age: 49.3 ± 11.4 years Male: 84.5% Diørnosis of OPIL: 0%	mJOA: 10.4 ± 3.33	5.1 $\pm$ 5.7 months	Mean 33.6 months (range 24-60) 77.7% (129/166)	<ul><li>mJOA</li><li>Nurick</li><li>Safety</li></ul>
Chen (2013); RCT	ACDF with titanium (n = 29) or PEEK cages (n = 31)		NR	NR	Mean 99.7 months (range 86-116) 75% (60/80)	<ul><li>JOA score</li><li>NDI score</li></ul>
Cheung (2008); Prospective case series	Cheung (2008); Prospective ACDF using autologous ICBG (n = 27); case series laminoplasty (n = 28)	Diagnosis of OFLE. 0% N = 55 Mean age: 64 ± 11 years Male: 63.6% Diamocia of OP11 undowurs	JOA: 10.0 ± 3.5	$17 \pm 30$ months	Mean 54 $\pm$ 26 months %NR	• JOA
Chibbaro (2009); Prospective case series	Multilevel oblique corpectomy without fusion	Diagnosis of OFLL, unknown N = 268 Mean age: 58 years (range 29-83) Male: 60% Diaanosis of OPLI - unknown	mJOA: 8.1 NDI: 55.2	9.6 months (range 4-33)	96 months %NR	<ul><li>mJOA</li><li>NDI</li><li>Safety</li></ul>
Fehlings (2012) <sup>c</sup> ; Prospective case series	ACDF or ACCF with instrumentation ( $n = 176$ ); laminectomy and fusion or laminoplasty ( $n = 107$ ); circumferential ( $n = 19$ )	N = 302 Mean age: 57 years (range 29-86) Male: 58 9% Diannosti of OPI I i unknown	mJOA: 12.8 <sup>b</sup>	R	24 months 91.1% (275/302)	<ul> <li>Safety (perioperative and delayed complications)</li> </ul>
Fehlings (2013) <sup>c</sup> ; Prospective case series	ACDF or ACCF with instrumentation (n = 169); laminectomy and fusion or laminoplasty (n = 95); circumferential (n = 14)	N = 278 Mean age: 56.3 ± 11.7 years Male: 59.4% Diamosis of OPLL: unknown	mJOA: 12.9 (95% Cl 12.5- 13.2) Nurrick: 3.1 ± 0.97 NDI: 41.8 + 20.8	25.8 $\pm$ 45.5 months	12 months 93.5% (260/278)	mJOA     Nurick     NDI
Fehlings (2015); Prospective case series	Anterior (ACDF, corpectomy) (n = 276); posterior (laminectomy with or without instrumented fusion, aminoplasty) (n = 191); circumferential (n =	N = 479 Mean age: 56.4 ± 11.9 years Male: 64.7% Diagnosis of OPLL: 28.2%	mJOA: 12.5 (95% CI 12.2- 12.8) Nurrick: 3.3 (95% CI 3.2-3.4) NDI: 36.4 (95% CI 34.3-38.4)	27.0 ± 34.7 months	12 months 84.3% (404/479) 24 months 81.6% (391/479)	<ul> <li>mJOA</li> <li>Nurick</li> <li>NDI</li> <li>30-meter walk test</li> </ul>
Hirai (2011); Prospective comparative	L1) ACCF (n = 39); laminoplasty using Miyazaki and Kirita method (n = 47)		JOA: 9.8 <sup>b</sup>	10.8 months <sup>b</sup>	60 months 91.5% (86/95)	<ul> <li>JOA</li> <li>Safety</li> </ul>
Hu (2014); Prospective comparative	Open-door laminoplasty with suture anchor (n $=$ 30) or titanium miniplate (n $=$ 25) fixation	Diagnosis of Or LL. 0% N = 55 Mean age: 55.6 ± 4.1 years Male: 54.5% Discretic of 0011.0%	JOA: 8.3 <sup>b</sup>	17.6 $\pm$ 3.6 months	Mean 20.9 months <sup>b</sup> 82.1% (55/67)	<ul><li>JOA</li><li>Safety</li></ul>
Kaneyama (2010); Prospective comparative	Open-door laminoplasty, $(n = 73)$ or double-door laminoplasty $(n = 73)$	Diagnosis of OLLEL 0% N = 146 Mean age: 64.1 years <sup>b</sup> Male: 73.9% Discrosis of OLL 1.1.1.2%	JOA: 9.95 <sup>b</sup>	R	2 weeks 100% (146/146)	C5 palsy
Karpova (2013)°; Prospective case series	ACDF or ACCF (n = 42); laminectomy and fusion or laminoplasty (n = 17); and circumferential (n = 2)	Diagnosis of Or LL 21.2% N = 61 Mean age: 56.2 ± 11.9 years Male: 69% OBI 1. 6.6%	mJOA: 12.8 ± 2.7	21.1 ± 18.2 months	12 months 94% (61/85)	<ul><li>mJOA</li><li>Safety</li></ul>
Kimura (2014); Prospective case series	Double-door laminoplasty	Diagnosis of Or LL 0.0% N = 116 Mean age: 63.3 ± 12.7 years Male: 67.2% Diagnosis of OPLL: 33.6%	JOA: II	$34.3\pm47.2$ months	Mean 35 ± 13 months 94.3% (116/123)	• JOA • Safety

Table 2. Characteristic of Included Studies Evaluating Operative Treatment for Degenerative Cervical Myelopathy.

Author (Year); Study Design	Operative Treatment	Patient Characteristics <sup>a</sup>	Baseline Severity (Mean)	Symptom Duration (Mean)	Follow-up, % Followed	Outcome Measures
Kommu (2014); Prospective case series	ACCF with plating (n = 14); laminectomy with (n = 5) and without (n = 40) instrumented fusion; laminoplasty (n = 3); circumferential (n = 1)	N = 63 Mean age: 51.1 years (range 30- 80) Male: 77.8% Diannsis of OPI1 : 0%	Nurick: 2.82	Motor symptoms: 7.1 months Sensory symptoms: 9.6 months	24 months %NR	<ul> <li>Nurick</li> <li>Safety</li> </ul>
Lian (2010); RCT	Noncontiguous ACDF plus discectomy using PEEK cages or titanium mesh (n = 55); continuous ACDF using titanium mesh (n = 50)	ange 38-	JOA: 8.9 <sup>b</sup>	R	Mean 31.5 months (range 24-48) 95.5% (105/110)	<ul> <li>JOA</li> <li>VAS (pain)</li> <li>Safety</li> </ul>
Machino (2013); Prospective case series	Double-door laminoplasty	Diagnosis of Or LL <i>V&amp;</i> N = 520 Mean age: 62.2 years Male: 63.7%	JOA: 10.4 $\pm$ 2.8	20.1 $\pm$ 32.0 months	Mean 33.3 months (range 12-90) 91.1% (520/571)	<ul> <li>JOA</li> <li>Safety</li> </ul>
Moussellard (2014); Prospective case series	ACCF ( $n = 7$ ); ACDF ( $n = 4$ ): laminectomy and fusion ( $n = 27$ ): laminoplasty and fusion ( $n = 28$ ): circumferential ( $n = 1$ )		JOA: 11.5 ± 2.6	23.5 $\pm$ 25.3 months	24 months %NR	<ul> <li>JOA</li> <li>30-meter walk test</li> </ul>
Nakashima (2014); RCT	Open-door (n = 44) or French-door (n = 46) laminoplasty	N = 90 Mean age: 63.0 years <sup>b</sup> Male: 63.3% Diagnosts of OPI + 73.3%	JOA: 9.0 <sup>b</sup>	NR	Mean 28.9 months <sup>b</sup> 95.7% (90/92)	<ul> <li>JOA</li> <li>Infection</li> <li>C5 palsy</li> </ul>
Ohashi (2014)/Katsumi (2012) <sup>d</sup> ; Prospective comparative	En bloc laminoplasty with prophylactic bilateral C4-C5 foraminotomy		JOA: 11.6 ± 2.7	R	24 months 85.8% (121/141)	<ul> <li>JOA</li> <li>VAS (pain)</li> <li>Safety</li> </ul>
Riew (2008); Prospective case series	ACDF (Prestige ST trial)	N = 52 Mean age: 46.0 ± 8.5 years Male: 40.4% Diamosis of ODI + 0%	NDI: 53.5 ± 16.9	NR	24 months 71.2% (37/52)	<ul> <li>NDI</li> <li>VAS (pain)</li> <li>Safety</li> </ul>
Sasai (2003); Prospective comparative	Midsagittal laminoplasty with $(n = 11)$ and without $(n = 63)$ microcervical foraminotomy; en bloc laminoplasty with microcervical foraminotomy $(n = 37)$	Dugnosis of Orlic Un N = 111 Mean age: 63 years Male: 61.3% Diagnosis of OPLL: 25.3%	JOA: 10.7 <sup>b</sup>	R	Postoperative (overall mean 39 months) 100% (111/111)	C5 palsy
Setzer (2009); Prospective case series	ACDF using titanium cages	N = 60 Mean age: 61.5 ± 14.6 years Male: 66.7%	mJOA: 10.32 ± 3.6	$22.0 \pm 30.5$ months	Mean 18.8 $\pm$ 4.6 months %NR	• mJOA
Suri (2003); Prospective case series	ACDF/ACCF or laminectomy/laminoplasty	Diagnosis of OFLL uniknown N = 146 Mean age: 47 years (range 17-76) Male: 79.5% Diamotis of OBL Lunknown	Nurick: 3.19 <sup>b</sup>	11.7 months (range 1.5-120)	24 months %NR	• Nurick
Suzuki (2009); Prospective comparative	Double-door laminoplasty	N = 98 Mean age: 59.6 years Male: 70.4% Diagnosts of OPIL: 70.4%	JOA: II.I ± 0.4	17.1 months <sup>b</sup>	Mean 81.6 months (range 60-144) 84.5% (98/116)	• JOA
Tanaka (2006); Prospective case series	Open-door laminoplasty: with additional foraminotomy ( $n = 18$ )	N = 62 Mean age: 64 years (range 32-89) Male: 75.8% Diagnosis of OPLL: 17.7%	JOA: 9.9 (range 5-15.5)	Я	14 months (range 12- 28) %NR	• JOA • Safety

(continued)

Author (Year); Study Design Operative Treatment	Operative Treatment	Patient Characteristics <sup>4</sup>	Baseline Severity (Mean)	утрот Оцганов (Глеан)	rollow-up, & rollowed	Outcome Measures
Tetreault (2013) <sup>c</sup> ; Prospective case series	Anterior (ACDF or ACCF with instrumentation) (n = 169); posterior (laminectomy and fusion or laminoplasty) (n = 951; circumérential (n = 14)	N = 272 Mean age: 56.5 ± 11.5 years Male: 58.8% Diaencis of OPI 1 - 8.8%	mJOA: 12.7 ± 2.6	23.8 $\pm$ 34.6 months	12 months 79.8% (217/272)	• mJOA
Thakar (2009, 2012); Prospective case series	Uninstrumented ACCF using iliac or fibular bone graft	N = 70 Mean age: 51.9 $\pm$ 9.9 years Male: 94.3% Diamosis of OPLL: unknown	Nurick: 3.13 ± 0.91	$15.2 \pm 4.5$ months	18 months (range 12- 22) %NR	<ul> <li>Nurick</li> <li>VAS (pain)</li> <li>Safety</li> </ul>
Wang (2015); Prospective comparative	Double-door laminoplasty using Kurokawa method (n = 64) or modified Kurokawa method (n = 88)	N = 152 Mean age: 57.2 ± 19 years Male: 74.3% Diagnosis of OPLL: 0%	JOA: 8.91 ± 2.79 NDI: 25.52 ± 4.13	N	Mean 52 ± 33 months 100% (152/152) 36-60 months 53.3% (81/152) ≥60 months 46.7% (71/152)	<ul> <li>JOA</li> <li>NDI</li> <li>VAS (pain)</li> </ul>
Ying (2007): RCT	ACCF with preserved posterior vertebral wall with ICBG or cages (n = 89); conventional ACCF with ICBG or cages (n = 89)	N = 178 Mean age: 49.0 years <sup>b</sup> Male: 65.7% Diaenosis of OPIL: 0%	JOA: 12.92 <sup>b</sup>	NR	78.7% (140/178)	<ul> <li>JOA</li> <li>Safety</li> </ul>
Yoshida (2013); Prospective Expansive laminoplasty case series	Expansive laminoplasty	N = 369 Mean age: 65 years (range 27-87) Male: 61.2% Diamosis of OPLL: 0%	JOA: 9.2 <sup>b</sup>	NR	Mean 54.1 months (range 24-97) 81.8% (369/451)	• JOA
Zong (2014); Prospective comparative	Posterior decompression (not specified)	N = 396 Mean age: 60.2 years <sup>b</sup> Male: 33.6% Diagnosis of OPLL: unknown	mJOA: 7.44 <sup>b</sup> NDI: 42.97 <sup>b</sup>	12.8 months <sup>b</sup>	24 months 64.9% (396/610)	<ul> <li>mJOA</li> <li>NDI</li> <li>VAS (pain)</li> </ul>

<sup>c</sup>Same study population (multicenter AOSpine North American Study): Fehlings (2012) was included for perioperative and delayed complications; Fehlings (2013) included primary outcome results and provided data regarding the association of preoperative disease severity on change in impairment; Karpova (2013) analyzed data from a single center and provided data regarding the impact of preoperative symptom duration on change <sup>a</sup>With the exception of Fehlings (2012, 2013, 2015), Riew (2008), Tetreault (2013), Wang (2015), and Ying (2007), demographics are reported only for patients that were analyzed (ie, those after loss-to-follow-up). Orthopaedic Association score; NDI, Neck Disability Index; NR, not reported; OPLL, ossification of the posterior longitudinal ligament; RCT, randomized controlled trial; VAS, visual analog scale for pain. <sup>b</sup>For studies that only provided demographic data stratified by treatment or other comparison groups, weighted means were calculated to provide an estimate for the entire study population. in impairment; and Tetreault (2013) provided supplemental modeling data using preoperative disease severity and symptom duration as predictors of achieving a mJOA  $\geq$  16.

<sup>d</sup>This was a comparative study using a historical control. Only the treatment group enrolled prospectively is included in our analysis.

Table 2. (continued)

and males comprised between 34% and 94% of the study populations. Disease pathoanatomy was also variable, with 9 studies including patients with OPLL (range = 6.6% to 33.6% of patients); of the remaining studies, 12 specifically excluded OPLL patients and 6 did not specify whether or not OPLL patients were included. Surgical approaches expectedly varied across studies: 7 included anterior decompression and fusion techniques only<sup>18,20,30,35,37,42,43,45</sup>; 12 included posterior decompression techniques that were either motionpreserving (laminectomy or laminoplasty) or used fusion<sup>24,25,27,28,31,33,34,36,39,40,44,46,47</sup>; and 8 studies incorporated both techniques.<sup>17,19,21-23,26,29,32,38,41,48</sup>

# Effect of Surgery on Functional Impairment, Disability, and Pain

Japanese Orthopaedic Association Score. Twenty-one studies reported change in JOA or mJOA scores in patients with moderate to severe DCM (mean baseline JOA = 7.4 to 12.9); follow-up duration ranged from 12 to 98 months.<sup>17-20,22-24,28,30-34,37,39,40,44-48</sup> The risk of bias was low or moderately low in 14 studies<sup>17,18,22,23,28,30,31,33,34,39,44-46,48</sup> and moderately high in 7 because of undefined or low follow-up rates.<sup>19,20,24,32,37,40,47</sup> A strong effect for improvement in function was observed at each time point (P < .001): the pooled SMD across 10 studies was 1.92 (95% confidence interval [CI] = 1.41 to 2.43) at 12-months follow-up (Figure 2a)<sup>20,22-24,28,30,32,34,45,48</sup>; 1.40 (95% CI = 1.12 to 1.67) across 12 studies at 13- to 36-months follow-up (Figure 2b)<sup>17,23,30-34,37,40,44,47,48</sup>; and 1.92 (95% CI = 1.41 to 2.69) across 7 studies at 36-months follow-up (Figure 2c).<sup>18-20,23,39,44,46</sup>

Neck Disability Index and Nurick Grade. Seven studies assessed change in NDI scores (follow-up duration ranged from 12 to 99.7 months) $^{18,20,22,35,44,47,48}$  and 5 studies evaluated change in Nurick grade (follow-up duration ranged from 12 to 24 months).<sup>22,29,38,42,48</sup> The risk of bias was low or moderately low in 4 studies reporting NDI<sup>18,22,44,48</sup> and in 2 studies reporting Nurick grade.<sup>22,48</sup> The remaining studies were classified as high or moderately high risk of bias because of undefined or low follow-up rates.<sup>20,29,35,38,42,47</sup> A strong effect for improvement was observed at each time point (P < .001) for both NDI and Nurick scores. For up to 12-months follow-up, the pooled MD for NDI was 18.02 (95% CI = 11.02 to 25.02) across 5 studies (Figure  $3a)^{20,22,35,44,48}$  and 1.42 (95% CI = 1.11 to 1.74) for Nurick across 2 studies (Figure 4a).<sup>22,48</sup> In the medium term (13-36 months), the pooled MD for NDI was 19.71 (14.01 to 25.42) across 5 studies (Figure 3b)<sup>20,35,44,47,48</sup> and 1.06 (95% CI = 0.69 to 1.43) for Nurick across 4 studies (Figure 4b).<sup>29,38,42,48</sup> In the long term (over 36 months), the pooled MD for NDI was 23.21 (95% CI = 11.84 to 34.58) across 2 studies (Figure 3c).<sup>18,20</sup>

*Visual Analog Scale for Pain.* Five studies reported change in VAS scores with follow-up duration ranging from 6 to 72 months.<sup>20,30,34,44,47</sup> The risk of bias was low or moderately low in 3 studies,<sup>30,34,44</sup> and moderately high in 2 studies because of

undefined or low follow-up rates<sup>20,47</sup> and/or concerns regarding selection bias.<sup>47</sup> A strong effect was seen for improvement in pain at each time point (P < .001): the pooled MD was 32.74 (95% CI = 18.39 to 47.10) across 4 studies at 12-months follow-up (Figure 5a)<sup>20,30,34,44</sup> and 32.55 (95% CI = 21.37 to 43.72) across 6 studies at 13- to 36-months follow-up (Figure 5b).<sup>20,30,34,35,44,47</sup> The MD from one study was 40.00 (95% CI = 37.01 to 42.99) at greater than 36-months follow-up (Figure 5c).<sup>20</sup>

# Effect of Preoperative Duration of Symptoms and Myelopathy Severity on Surgical Outcomes

Four studies stratified their sample based on preoperative duration of symptoms (<6, 6-12, >12 months<sup>19</sup>; <12 or >12 months<sup>26,39</sup>; or <12, 12-24, and >24 months<sup>38</sup>) (Table 3). Patients in these subgroups exhibited similar functional improvements as assessed by JOA or mJOA scores.<sup>19,26,39</sup> A fifth study by Tetreault et al evaluated whether duration of symptoms is a significant predictor of a postoperative mJOA score  $\geq 16$ .<sup>41</sup> Based on their results, the odds of achieving a mJOA  $\geq$  16 decreased by 22% when a patient moved from a shorter to a longer duration of symptoms group ( $\leq 3$ ; >3,  $\leq 6$ ; >6,  $\leq 12$ ; >12,  $\leq 24$ ; >24 months) (adjusted odds ratio: 0.78; 95% CI = 0.61 to 0.997; *P* = .048).

Two studies compared postoperative outcome in cohorts of varying preoperative myelopathy severities (Table 4).<sup>20,22</sup> Two other studies examined the association between baseline severity scores and surgical outcomes.<sup>37,41</sup> With respect to change in mJOA, Fehlings et al<sup>22</sup> reported improvements across all levels of impairment; patients with severe myelopathy (mJOA < 12), however, demonstrated the greatest functional change (4.91, 95% CI = 4.34 to 5.49), whereas those with mild disease (mJOA > 15) achieved the least amount of improvement (1.29, 95% CI = 0.70 to 1.87; Table 4). Conversely, Chibbaro et al<sup>20</sup> reported that a similar percentage of patients with either moderate (mJOA = 10-13) or severe (mJOA = 5-9) myelopathy exhibited improvement on the mJOA scale at long-term followup (96 months). In a study by Tetreault et al,<sup>41</sup> the odds of achieving a mJOA  $\geq$  16 at 1-year following surgery was 1.22 times greater for every 1-point increase in preoperative mJOA score. A complementary result was observed in a study by Setzer et al,<sup>37</sup> who reported that patients with more severe preoperative impairment were less likely to exhibit a postoperative improvement of 2 or more points on the mJOA scale at 18-months follow-up (odds ratio = 0.72; 95% CI = 0.66 to 0.92).<sup>37</sup>

#### Complications

Pooled estimates of complications across included studies are presented in Table 5 in order of decreasing frequency. The cumulative incidence of complications is low (14.1%; 95% CI = 10.1% to 18.2%). Mortality was reported in 0.3% of patients (95% CI = 0.0% to 0.5%). Neurological complications included worsening of myelopathy (1.3%; 95% CI = 0.5% to 2.1%), laryngeal nerve injury/dysphagia (2.2%; 95% CI = 1.4% to 3.0%), and C5 radiculopathy or palsy (1.9%; 95% CI = 1.4% to 2.4%). Infection was

	Pos	t Sco	re	Pre	Scor	e		Std. Mean Difference		Std. Me	an Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% C	1	IV, Ran	ndom, 95% CI
Chibbaro 2009	14.9	1.4	268	8.1	2.2	268	10.2%	3.68 [3.40, 3.96	)		-
Fehlings 2013	15.7	1.4	260	12.9	2.2	260	10.3%	1.52 [1.32, 1.71	)		-
Fehlings 2015	14.9	1.4	479	12.5	2.2	479	10.4%	1.30 [1.16, 1.44	]		-
Hirai 2011	13.6	2.5	86	9.8	3	86	10.0%	1.37 [1.04, 1.70	]		-
Hu 2014	14.7	1	55	8.3	1.6	55	8.6%	4.76 [4.02, 5.51	]		
Kimura 2014	14	1.4	116	11.1	2.2	116	10.1%	1.57 [1.27, 1.86	]		-
Lian 2010	13.3	2	105	9	2.6	105	10.1%	1.85 [1.52, 2.17	]		-
Moussellard 2014	13.4	2.2	67	11.5	2.6	67	10.0%	0.78 [0.43, 1.14	]		-
Ohashi 2014	14.7	1.9	121	11.6	2.7	121	10.2%	1.32 [1.05, 1.60	]		-
Ying 2007	16.2	0.7	140	12.9	3.1	140	10.2%	1.46 [1.20, 1.73	1		-
Total (95% CI)			1697			1697	100.0%	1.92 [1.41, 2.43	1		•
Heterogeneity: Tau <sup>2</sup> =	= 0.64; 0	Chi <sup>2</sup> =	332.2	4, df =	9 (P	< 0.00	001); I <sup>2</sup> =	= 97%	-+	5	
Test for overall effect:	Z = 7.3	39 (P	< 0.00	001)							· -
). 13-36 month	IS										
		t Sco	re	Pre	Scor	P		Std. Mean Difference		Std. Me	ean Difference
Study or Subgroup						-	Weight	IV, Random, 95% C			ndom, 95% Cl
Bapat 2008	15.8		129	10.4		129	8.3%	2.10 [1.80, 2.41			-
Fehlings 2015	14.9		479	12.5	3.5	479	9.0%	0.91 [0.78, 1.04			
Hirai 2011	14.2	_	86	9.8	3	86	8.1%	1.59 [1.24, 1.93			-
Lian 2010		2.2	105	9.0	3	105	8.2%	1.89 [1.57, 2.22			-
Machino 2013	13.6			10.4	-	520	9.0%	1.20 [1.07, 1.34			
Moussellard 2014	12.7		67	11.5		67	8.1%	0.46 [0.12, 0.80			-
Nakashima 2014	12.7		90		2.7	90	8.2%	1.39 [1.06, 1.72			-
Ohashi 2014	14.8			11.6		121	8.4%	1.37 [1.09, 1.65			-
Setzer 2009	11.5			10.3		60	7.9%				L
Tanaka 2006	14.2		62	9.9	3	62	7.7%	1.62 [1.22, 2.03			-
Wang 2015	14.9		152		3.1	152	8.4%				-
Zong 2014	12.3		396		3.4	396	8.9%	1.59 [1.43, 1.75			-
			2267								
Total (95% CI)							100.0%	1.40 [1.12, 1.67	· .		•
Heterogeneity: Tau <sup>2</sup> =					11 ()	< 0.0	0001); F	= 94%	-4	-2	0 2
Test for overall effect:	2 = 9.8	59 (P	< 0.00	001)							
c. >36 months											
	Pos	t Sco	re	Pre	Scor	e		Std. Mean Difference		Std. Me	an Difference
Study or Subgroup						-	Weight	IV, Random, 95% C			ndom, 95% Cl
Chen 2013	13.5		60		1.4	60	13.9%	2.34 [1.87, 2.81			-
Cheung 2008	12.4	3.2	55	10	3.5	55	14.2%	0.71 [0.32, 1.10	1		-
Chibbaro 2009	16.2	1.9	268	8.1	2.2	268	14.4%	3.94 [3.64, 4.23			
Hirai 2011	13.9	2.6	86	9.8	3	86	14.3%	1.45 [1.12, 1.79			-
Suzuki 2009	14.3	2	98	11.1	4	98	14.4%	1.01 [0.71, 1.31			+
Wang 2015	14.9		71		2.9	152	14.3%	2.26 [1.91, 2.61			-
Yoshida 2013	12.7	1.9	369	9.2	2.2	369	14.6%	1.70 [1.53, 1.87			-
1031104 2015											

**Figure 2.** Forest plots and meta-analysis of postoperative Japanese Orthopaedic Association (JOA) scores among patients with degenerative cervical myelopathy. Significant improvements in JOA scores are observed following surgery at short (Panel A, 6-12 months), medium (Panel B, 13-36 months), and long (Panel C, >36 months) term follow-up.

infrequent and reported in 1.5% of patients (95% CI = 1.0% to 2.1%). Instrumentation or graft complication occurred in 2.0% of cases (95% CI = 1.3% to 2.7%).

# **Evidence Summary**

The evidence summary is presented in Table 6.

Effect of Surgery on Functional Impairment, Disability and Pain. There is moderate strength of evidence that surgical intervention for DCM results in significant improvements in function at short-, medium-, and long-term follow-up as assessed by JOA or mJOA scores. Clinically meaningful improvements in both the NDI and Nurick scores were also observed following surgery. However, the strength of evidence for these findings was rated as very low for NDI and low for Nurick score. Finally, based on moderate evidence, surgical intervention results in significant improvements in pain scores at 6- to 12-months follow-up.

	P	e Score		Po	st Score	2		Mean Difference		Mean	Difference	
Study or Subgroup	Mean		Total				Weight	IV, Random, 95% C			dom, 95% CI	
Chibbaro 2009	55.2	14.08	268	28.4	14.78	268	20.8%	26.80 [24.36, 29.24]	1			
Fehlings 2013	42.01	21.27	260	30.73	23.61	260		11.28 [7.42, 15.14			-	
ehlings 2015	36.38	22.89	479	23.44	20.94	479		12.94 [10.16, 15.72]			+	
Riew 2008	53.5	16.9	52	24.1	21.9	52	17.1%	29.40 [21.88, 36.92]				-
Wang 2015	26.36	5.16	152	14.8	3.68	152	21.3%	11.56 [10.55, 12.57]	1			
Total (95% CI)			1211			1211	100.0%	18.02 [11.02, 25.02]	í.		•	
Heterogeneity: Tau <sup>2</sup> =	59.67:	$Chi^2 = 1$	46.22	df = 4	(P < 0.0	00001)	$ ^2 = 97\%$	5	<b>—</b>	-		
Test for overall effect:									-50	-25	Ó 2'5	50
. 13-36 mont		e Score		Po	st Score			Mean Difference		Mean	Difference	
Study or Subgroup	Mean			Mean			Weight	IV, Random, 95% C			dom, 95% Cl	
Chibbaro 2009		12.05	268		12.56	268	-	27.70 [25.62, 29.78]		, nam	-	
Fehlings 2015		22.89	479		22.89	479		13.18 [10.28, 16.08]				
Riew 2008	53.5	16.9	52	22.4	22.2	52		31.10 [23.52, 38.68]				_
Wang 2015	26.35	5.16		13.12	3.6	152		13.23 [12.23, 14.23]				
	43	11.1	396	26.7	10.7	396		16.30 [14.78, 17.82			•	
ong 2014						0.000						
			1347			1347	100.0%	19.71 [14.01, 25.42]			•	
Total (95% CI) Heterogeneity: Tau <sup>2</sup> =			69.57		(P < 0.0				-50	-25	0 25	50
Zong 2014 Total (95% Cl) Heterogeneity: Tau <sup>2</sup> = Test for overall effect: c. >36 months	Z = 6.7		69.57		(P < 0.0				·	-25	0 25	50
Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect:	Z = 6.7		169.57	1)	(P < 0.0				·		0 25	50
Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect:	Z = 6.7	8 (P < 0	169.57	l) Pos	t Score	00001)		5	·	Mean		50
Fotal (95% CI) Heterogeneity: Tau <sup>2</sup> = Fest for overall effect: . >36 months	Z = 6.7 Pr Mean	8 (P < 0	169.57	l) Pos	t Score SD T	00001)	;   <sup>2</sup> = 98% Veight	s Mean Difference	·	Mean	Difference	50
Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Fest for overall effect: . >36 months Study or Subgroup	Z = 6.7 Pr Mean 35.8	8 (P < ( e Score SD	Total	Pos Mean	t Score SD T 2.45	00001) Fotal V 60	; I <sup>2</sup> = 98% <u>Veight</u> 49.9% 1	Mean Difference IV, Random, 95% CI	·	Mean	Difference	5(

Figure 3. Forest plots and meta-analysis of postoperative Neck Disability Index (NDI) scores among patients with degenerative cervical myelopathy. Significant improvements in NDI scores are observed following surgery at short (Panel A, 6-12 months), medium (Panel B, 13-36 months), and long (Panel C, >36 months) term follow-up.

Effect of Preoperative Duration of Symptoms and Myelopathy Severity on Surgical Outcomes. There is very low strength of evidence that preoperative duration of symptoms is not associated with surgical outcomes, evaluated by change in JOA or mJOA scores between patients with a duration of symptoms greater than or less than 12 months. There is low strength of evidence that the odds of achieving a postoperative mJOA  $\geq 16$ decrease as duration of symptoms increases.

There is moderate strength of evidence that improvements on the mJOA are less for patients with mild myelopathy than those with moderate or severe disease. There is low strength of evidence that the odds of achieving a postoperative mJOA  $\geq 16$ decrease as preoperative myelopathy severity increases. Based on very low strength of evidence, improvement in NDI and Nurick scores do not depend on preoperative disease severity.

*Complications.* Surgical complications occur infrequently in patients treated surgically for DCM. C5 radiculopathy or palsy and surgical site infection were reported in fewer than 2% of

patients. The strength of evidence for these estimates is low. The cumulative incidence of reoperation, dural tears, worsening of myelopathy, and death was less than 1.5% and was 2% for instrumentation failure. The strength of evidence to support these estimates is very low.

#### Discussion

The management of DCM remains controversial with limited evidence available to the clinician and patient to make informed decisions. The only randomized controlled trial<sup>5,6</sup> concluded that there is no difference in mJOA scores between patients with "milder" DCM (mJOA  $\geq$  12) that receive operative versus nonoperative care. This is in contrast to results reported in other observational studies,<sup>7,8</sup> which suggest that surgery leads to superior functional recovery. These conflicting results highlight the importance of patient selection and identifying ideal surgical candidates. Further controversy arises because there are a variety of surgical techniques available to

#### a. 6-12 months

	Pr	e Score	2	Pos	st Scor	e		Mean Difference		Mean	Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI		IV, Ran	dom, 95% CI	
Fehlings 2013	3.11	1.23	260	1.52	1.4	260	47.9%	1.59 [1.36, 1.82]				
Fehlings 2015	3.3	1.23	479	2.03	1.67	479	52.1%	1.27 [1.08, 1.46]				
Total (95% CI)			739			739	100.0%	1.42 [1.11, 1.74]			•	
Heterogeneity: Tau <sup>2</sup> =	0.04; 0	Chi <sup>2</sup> =	4.58, d	f = 1 (F)	9 = 0.0	)3); l <sup>2</sup> =	78%			-		+
Test for overall effect:	Z = 8.9	90 (P <	0.000	01)					-4	-2	0 2	4
). 13-36 month	s											
o. 13-36 month		e Score	2	Pos	st Scor	'e		Mean Difference		Mean	Difference	
		e Score SD	e Total	Pos Mean	st Scor SD		Weight	Mean Difference IV, Random, 95% CI			Difference dom, 95% CI	
Study or Subgroup	Pro Mean		1 - C - C - C - C - C - C - C - C - C -	Mean			Weight 28.3%					
5. 13-36 month Study or Subgroup Fehlings 2015 Thakar 2009	Pro Mean 3.3	SD	Total	Mean 1.96	SD	Total		IV, Random, 95% CI				
Study or Subgroup Fehlings 2015	Pro Mean 3.3 3.13	SD 1.23	Total 479	Mean 1.96 1.76	SD 0.12	Total 479	28.3%	IV, Random, 95% CI 1.34 [1.23, 1.45]				
Study or Subgroup Fehlings 2015 Thakar 2009	Pro Mean 3.3 3.13	SD 1.23 0.91	Total 479 70	Mean 1.96 1.76	SD 0.12 1.13	Total 479 70	28.3% 23.3%	IV, Random, 95% CI 1.34 [1.23, 1.45] 1.37 [1.03, 1.71]				
Study or Subgroup Fehlings 2015 Thakar 2009 Kommu 2014	Pro Mean 3.3 3.13 2.85	SD 1.23 0.91 0.96	Total 479 70 63	Mean 1.96 1.76 2.05	SD 0.12 1.13 1.12	Total 479 70 63 146	28.3% 23.3% 22.6%	IV, Random, 95% CI 1.34 [1.23, 1.45] 1.37 [1.03, 1.71] 0.80 [0.44, 1.16]				

**Figure 4.** Forest plots and meta-analysis of postoperative Nurick scores among patients with degenerative cervical myelopathy. Significant improvements in Nurick scores are observed following surgery at short (Panel A, 6-12 months) and medium (Panel B, 13-36 months) term follow-up.

the surgeon to accomplish the objectives of spinal cord decompression, spinal column stabilization, and, if necessary, spinal realignment. Based on a study by Fehlings et al, patients treated anteriorly or posteriorly exhibit similar improvements<sup>22</sup>; however, selection criteria between approaches remain controversial and is the topic of a current randomized controlled trial of ventral versus dorsal surgery.<sup>49</sup>

There is a lack of evidence on the comparative effectiveness of surgery and nonoperative treatment in patients with DCM. As a result, this systematic review aims to summarize data from prospective studies that report changes in outcomes and rates of complications associated with surgical intervention.

The first objective of this review was to determine the change in functional impairment, disability, and pain following surgical management. Based on our conclusions, there is moderate evidence to suggest that surgery results in significant gains in mJOA or JOA scores at short-, medium-, and longterm follow-up. These reported improvements exceed the minimum clinically important difference for this metric, 50,51 suggesting that patients undergoing surgery are expected to exhibit functional gains in addition to disease stabilization. Furthermore, based on low evidence, surgery also results in significant improvements in Nurick scores at short- and medium-term follow-up. The mJOA has been adopted as the standard for evaluating functional impairment in patients with DCM; this scale is responsive to change, especially in patients with moderate and severe myelopathy.<sup>52</sup> Reported improvements on the Nurick Grade were less than those observed on the mJOA; however, this is likely because this scale only has 6 categories and because it is inherently less sensitive.

The second objective of this review was to evaluate whether duration of symptoms or preoperative myelopathy severity were associated with surgical outcomes. Based on our findings, operative management results in significant functional improvements in patients with both a "short" and "long" duration of symptoms and in patients with mild, moderate, and severe disease. The clinical prediction rule proposed by Tetreault et al,<sup>41</sup> however, suggested that the likelihood of achieving a postoperative mJOA  $\geq$  16 decreases with a longer duration of symptoms and a lower baseline severity score. The authors of this study acknowledged that patients with severe myelopathy are, in part, less likely to attain a postoperative mJOA  $\geq$  16 as this would require a  $\geq$ 5-point improvement; to address this limitation, a second model was developed to predict a postoperative mJOA > 12 and also included baseline severity score as a relevant factor.<sup>48</sup> Future studies are required to validate these associations and should use outcome measures that are better suited to assess patients with mild myelopathy. In a recent review, Kalsi-Ryan et al<sup>53</sup> suggested that ancillary measures be used in combination with the mJOA or JOA scale to assess impairment in patients with DCM; these include tools that focus on upper extremity function (Graded Redefined Assessment of Strength Sensibility and Prehension and grip dynamometer), gait (GAITRite analysis and 30-meter walk test), and balance (Berg Balance Score).

The third objective of this review was to evaluate the incidence of surgical complications. The quality of evidence for our conclusions was low to very low, due to risk of bias and imprecision. The imprecision of our estimates likely results from the variability of definitions across studies; this highlights

	Pr	e Score	2	Pos	t Scol	re		Mean Difference		Mean	Differ	ence	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI		IV, Ran	dom, 9	5% CI	
Chibbaro 2009	65	19.7	268	13	4.9	268	25.5%	52.00 [49.57, 54.43]					
Lian 2010	49.7	13.5	105	10.75	6.5	105	25.4%	38.95 [36.08, 41.82]				-	-
Ohashi 2014	32	30	121	31	_	121		1.00 [-6.31, 8.31]			+		
Wang 2015	71.55	21.1	152	34.2	16	152	25.1%	37.35 [33.14, 41.56]				-	
Total (95% CI)			646			646	100.0%	32.74 [18.39, 47.10]				-	•
Heterogeneity: Tau <sup>2</sup> =	= 209.06	; Chi <sup>2</sup>	= 193.	96, df =	= 3 (P	< 0.00	001); I <sup>2</sup> =	98%		-25	<u> </u>	1	-
Test for overall effect:	Z = 4.4	7 (P <	0.000	01)					-50	-25	0	25	50
b. 13-36 mont		Score		Pos	t Scor	ne -		Mean Difference		Mean	Differ	ence	
Study or Subgroup	Mean		Total			-	Weight	IV, Random, 95% CI		IV, Ran			
Chibbaro 2009		20.8	268		13.8	268		45.00 [42.01, 47.99]		,	1		+
Lian 2010	49.7		105	11.9	7	105		37.80 [34.89, 40.71]					-
Ohashi 2014	32	30	121	32	27	121		0.00 [-7.19, 7.19]			-		
Riew 2008	67.5		55		13.8	52		52.50 [45.85, 59.15]					-
Wang 2015	71.6	20.7	152	33.5	17.5	152		38.10 [33.79, 42.41]				-	-
			396	13.2	9.6	396							
Zong 2014	34.3	26.2	290	15.2	9.0	390	17.0%	21.10 [18.35, 23.85]				•	
	34.3	26.2	1097	15.2	9.0			21.10 [18.35, 23.85] 32.55 [21.37, 43.72]				•	•
Zong 2014			1097			1094	100.0%	32.55 [21.37, 43.72]	-	-1-		* •	-
Zong 2014 Total (95% CI)	= 189.17	; Chi²	1097 = 262.	25, df =		1094	100.0%	32.55 [21.37, 43.72]	-50	-25	0	25	50
Zong 2014 Total (95% CI) Heterogeneity: Tau <sup>2</sup> =	= 189.17 : Z = 5.7	; Chi²	1097 = 262.	25, df =		1094	100.0%	32.55 [21.37, 43.72]	-50	-25	0	25	50
Zong 2014 Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect: c. >36 months	= 189.17 : Z = 5.7 Pre	'; Chi <sup>2</sup> '1 (Ρ < • Score	1097 = 262. 0.000	25, df = 01) Pos	= 5 (P t Scor	1094 < 0.00	100.0% 1001); l <sup>2</sup> =	32.55 [21.37, 43.72] 98% Mean Difference	-50	Mean	Differ	ence	50
Zong 2014 Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect: c. >36 months	= 189.17 : Z = 5.7	'; Chi <sup>2</sup> '1 (Ρ < • Score	1097 = 262. 0.000	25, df = 01) Pos	= 5 (P t Scor	1094 < 0.00 e Total	100.0% 1001);   <sup>2</sup> = Weight	32.55 [21.37, 43.72] 98% Mean Difference IV, Random, 95% CI	-50		Differ	ence	50
Zong 2014 Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect: c. >36 months Study or Subgroup	= 189.17 : Z = 5.7 Pre Mean	'; Chi <sup>2</sup> '1 (Ρ < • Score	1097 = 262. 0.000	25, df = 01) Pos Mean	= 5 (P t Scor	1094 < 0.00 e Total	100.0% 1001);   <sup>2</sup> = Weight	32.55 [21.37, 43.72] 98% Mean Difference	-50	Mean	Differ	ence	50
Zong 2014 Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect	= 189.17 : Z = 5.7 Pre Mean	2; Chi <sup>2</sup> 1 (P < Score	1097 = 262. 0.000	25, df = 01) Pos Mean	= 5 (P t Scor SD	1094 < 0.00 e <u>Total</u> 268	100.0% 1001); l <sup>2</sup> = <u>Weight</u> 100.0%	32.55 [21.37, 43.72] 98% Mean Difference IV, Random, 95% CI	-50	Mean	Differ	ence	50
Zong 2014 Total (95% CI) Heterogeneity: Tau <sup>2</sup> = Test for overall effect: c. >36 months <u>Study or Subgroup</u> Chibbaro 2009	= 189.17 : Z = 5.7 Pre <u>Mean</u> 65	2; Chi <sup>2</sup> 1 (P < Score	1097 = 262. 0.000 Total 268	25, df = 01) Pos Mean	= 5 (P t Scor SD	1094 < 0.00 e <u>Total</u> 268	100.0% 1001); l <sup>2</sup> = <u>Weight</u> 100.0%	32.55 [21.37, 43.72] 98% Mean Difference IV, Random, 95% CI 40.00 [37.01, 42.99]	-50	Mean	Differ	ence	50

Figure 5. Forest plots and meta-analysis of postoperative Visual Analog Scale (VAS) scores among patients with degenerative cervical myelopathy. Significant improvements in VAS scores are observed following surgery at short (Panel A, 6-12 months), medium (Panel B, 13-36 months), and long (Panel C, >36 months) term follow-up.

a major knowledge gap and a need to develop a classification system for surgical complications. Nevertheless, reported cumulative incidences were low for perioperative mortality (0.3%), recurrent laryngeal nerve injury or dysphagia (2.2%), and C5 radiculopathy or palsy (1.9%). Nationwide estimates of perioperative morbidity have previously been presented by Shamji et al<sup>54,55</sup>; their results indicated that, among 8548 patients with diffuse cervical spondylosis, rates of mortality were 0.33% and 0.69% in patients approached anteriorly and posteriorly, respectively. Dysphagia was reported in 3.63% of patients approached anteriorly and 1.87% of patients approached posteriorly. Posterior surgery was associated with a more frequent need for transfusion (1.38% anteriorly, 7.20% posteriorly) and a higher rate of clinically significant hematoma (0.80% anteriorly, 2.12% posteriorly).

The cost-effectiveness of surgery must also be considered when developing treatment protocols. Two studies were identified that evaluated the cost-utility of surgery in Canadian patients enrolled in the AOSpine North America and/or International studies. The first study (based on the North America study) estimated the cost of surgery to be \$21066.44, with an incremental cost-utility ratio (ICUR) of \$32916 per quality adjusted life year (QALY).<sup>56</sup> A second study by Witiw et al (based on the North America and International studies) conducted a more rigorous cost-utility analysis using a 2-arm, Markov State Transition model where values for subjects undergoing surgery were compared with estimated counterfactual outcomes of initial conservative management.<sup>57</sup> In a primary model, the lifetime ICUR was determined to be \$11496/ OALY gained for surgical intervention, an estimate considered very cost-effective according to criteria defined by the World Health Organization. Further testing using a Monte Carlo probabilistic sensitivity analysis revealed that 97.9% of estimates fell within the WHO threshold, suggesting robustness to variability in the parameter estimates. To supplement this testing, a highly conservative assumption that individuals undergoing initial nonoperative management would not experience any neurologic decline over their lifetime was added to the model. In this scenario, the ICUR was calculated as \$20548/QALY gained with 94.7% of estimates falling within the WHO

Author and Symptom Duration	n	Preoperative JOA Score (95% CI or $\pm$ SD)	JOA Change Score (95% CI)	Nurick Improved (%) <sup>a</sup>
Short-term F/U (12 months)				
Karpova (2013)				
<12 months	NR	13.5 <sup>b</sup> (12.6, 14.1)	3.1 <sup>b</sup> (2.5, 3.7)	
>12 months	NR	11.9 <sup>b</sup>	2.6 <sup>b</sup> (2.0, 3.2)	
Suzuki (2009) <sup>c</sup>				
<12 months	65	11.2 $\pm$ 3.0	3.4 (2.1, 4.7)	
>12 months	33	11.1 ± 3.0	2.8 (1.4, 4.2)	
Medium-term F/U (24-36 months	)			
Suzuki (2009)				
<12 months	65	11.2 $\pm$ 3.0	3.4 (2.4, 4.6)	
>12 months	33	11.3 ± 3.0	2.7 (1.2, 3.8)	
Suri (2003)				
<12 months	86			58.1%
12-24 months	31			71.0%
>24 months	29			51.7%
Long-term F/U (54-60 months)				
Suzuki (2009)				
<12 months	65	11.2 $\pm$ 3.0	3.4 (2.4, 4.6)	
>12 months	33	11.2 $\pm$ 3.0	2.4 (1.2, 3.8)	
Cheung (2008)				
<6 months	NR	8.6 ± 3.4	2.9	
6-12 months	NR	10.3 ± 3.9	3.1	
>12 months	NR	10.6 ± 3.6	1.0	

**Table 3.** The Impact of Preoperative Duration of Symptoms on Japanese Orthopaedic Association (JOA) and Nurick Outcomes at 3 Different Follow-up Periods.

Abbreviations: CI, confidence interval; F/U, follow-up; JOA, Japanese Orthopaedic Association Score; NR, not reported; SD, standard deviation. <sup>a</sup>Proportion of patients that improved by I or more Nurick grades at follow-up.

<sup>b</sup>mJOA scores.

<sup>c</sup>Values estimated from Figure 4 in the Suzuki 2009 article; 95% confidence intervals calculated from estimate.

Table 4. Modified Japanese Orthopedic Association, Neck Disability Index, and Nurick Outcomes Based on Preoperat	ative Myelopathy S	everity.
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Author	n	mJOA Change Score (95% CI)	Improved mJOA (%)	NDI Change Score (95% CI)	Nurick Change Score (95% CI)
Short-term F/U (12 months)					
Fehlings (2013)					
Mild (mJOA $\geq$ 15)	78	1.29 (0.70, 1.87)		12.05 (7.76, 16.34)	1.54 (1.22, 1.86)
Moderate (mJOA 12-14)	105	2.58 (2.07, 3.09)		9.79 (5.90, 13.68)	1.51 (1.22, 1.81)
Severe (mJOA <12)	77	4.91 (4.34, 5.49)		12.53 (8.05, 17.02)	1.74 (1.41, 2.08)
Long-term F/U (96 months)					
Chibbaro (2009)					
Moderate (mJOA 10-13)	90		86.7		
Severe (mJOA 5-9)	178		86.5		

Abbreviations: CI, confidence interval; F/U, follow-up; mJOA, modified Japanese Orthopaedic Association Score; NDI, Neck Disability Index; NR, not reported; SD, standard deviation.

threshold; this finding further supports the cost-effectiveness of surgical intervention. Unfortunately, these analyses only explored the cost-effectiveness of surgery in Canada and did not stratify their sample based on preoperative myelopathy severity.

#### Limitations

A substantial limitation of this systematic review is that the impact of surgical approach on recovery was not studied. Surgical interventions vary in both approach (anterior or posterior) and objectives (decompression, stabilization, realignment); as a result, data may be too heterogeneous to conduct a meta-analysis. Among anterior surgical approaches, a previous systematic review<sup>58</sup> demonstrated heterogeneous neurological, pain, and alignment outcomes among multiple discectomy, corpectomy, and hybrid procedures. Among posterior surgical approaches, a previous systematic review by Yoon et al<sup>59</sup> reported that both laminoplasty and laminectomy and fusion procedures result in functional improvement. Unfortunately, the comparative effectiveness and safety of these and other approaches have not been rigorously addressed.

Complications	Number of Studies and References	n/N	Cumulative Incidence %	95% CI
Axial pain	3 [21, 30, 45]	33/585	5.6	3.8, 7.5
Laryngeal nerve injury/dysphagia	8 [17, 20, 21, 23, 29, 30, 43, 45]	26/1182	2.2	1.4, 3.0
Instrumentation/graft complication	9 [17, 18, 21, 24, 30, 35, 43, 45, 48]	28/1411	2.0	1.3, 2.7
C-5 radiculopathy/palsy	15 [17, 20, 21, 23-25, 27, 28, 30, 31, 33, 36, 40, 43, 48]	50/2661	1.9	1.4, 2.4
Pseudarthrosis	7 [17, 18, 21, 23, 30, 34, 45]	17/954	1.8	0.9, 2.6
Infection (deep and superficial)	10 [20, 21, 24, 27, 29-31, 33, 43]	32/2074	1.5	1.0, 2.1
Adjacent segment disease (symptomatic)	2 [17, 21]	6/404	1.5	0.3, 2.7
Reoperation/revision	7 [20, 21, 23, 29, 30, 32, 35]	13/943	1.4	0.6, 2.1
Dural tear/CSF leak	11 [20, 21, 24, 27, 28, 30-33, 43, 45]	26/1893	1.4	0.8, 1.9
Worsening of myelopathy	2 [21, 48]	10/781	1.3	0.5, 2.1
Hematoma	7 [21, 24, 30, 32, 43, 45, 48]	11/1237	0.9	0.4, 1.4
Radiculopathy/palsy (not C5)	3 [21, 36, 43]	4/464	0.9	0.0, 1.7
Neurologic deterioration/new deficit	3 [17, 21, 31]	9/969	0.9	0.3, 1.5
Delayed wound healing/dehiscence	2 [21, 32]	3/369	0.8	0.0, 1.7
Dysphonia	3 [21, 23, 48]	6/867	0.7	0.1, 1.2
Postoperative deformity	2 [17, 21]	2/404	0.5	0.0, 1.2
Death	6 [17, 18, 21, 30, 32, 48]	3/1162	0.3	0.0, 0.5
Stroke/transient ischemic attack	3 [21, 31, 43]	3/873	0.3	0.0, 0.7
Esophageal injury	2 [23, 30]	0/191	0.0	0.0, 2.9
Other	7 [17, 20, 21, 23, 32, 43, 48]	51/1382	3.7	2.7, 4.7
Cardiopulmonary	1 [21]	10/302	3.3	1.3, 5.3
Fracture	I [27]	3/141	2.1	0.0, 4.5
Bed sore	ı [17]	1/129	0.8	0.0, 2.3
Spinal cord injury	I [24]	0/55	0.0	0.0, 0.0
Any complication <sup>a</sup>	I [21]	40/283	14.1	10.1, 18.2

Abbreviations: CI, confidence interval; CSF, cerebrospinal fluid; n, number of complications; N, total number of patients.

<sup>a</sup>Only reported for anterior and posterior surgery; 19 patients who had circumferential decompression were not included.

#### Table 6. Evidence Summary.

	Studies; N [References]	Strength of Evidence	Conclusions/Comments
Key Question 1: What are JOA/mJOA	the expected functional, disabilit 6-12 months 10 studies [20, 22-24, 28, 30, 32, 34, 45, 48] (N = 1697) 13-36 months 12 studies [17, 23, 30-34, 37, 40, 44, 47, 48] (N = 2267) $\geq$ 36 months 7 studies [18-20, 23, 39, 44, 46]	y and þain outcom Moderate <sup>a</sup>	tes following surgical intervention for DCM? Surgical intervention resulted in improved JOA/mJOA scores at all time points assessed. Pooled standardized mean differences showed a large effect for improvement in function from baseline at short term (10 studies, SMD 1.92), medium term (12 studies, SMD 1.40), and long term (7 studies, SMD 1.92) follow-up.
NDI	$\begin{array}{l} (N = 1088) \\ 6-12 \ months \\ 5 \ studies [20, 22, 35, 44, 48] \\ (N = 1211) \\ 13-36 \ months \\ 5 \ studies [20, 35, 44, 47, 48] \\ (N = 1347) \\ \geq 36 \ months \\ 2 \ studies [18, 20] \ (N = 328) \end{array}$	Very Low <sup>b</sup>	Surgical intervention resulted in improved NDI scores at all time points assessed. Pooled mean differences showed clinically meaningful improvement in disability from baseline at short term (5 studies, MD 18.02), medium term (5 studies, MD 19.71), and long term (2 studies, MD 23.21) follow-up.

	Studies; N [References]	Strength of Evidence	Conclusions/Comments
Nurick	6-12 months 2 studies [22, 48] (N = 739) 13-36 months 4 studies [29, 38, 43, 48] (N = 758)	Low	Surgical intervention resulted in improved Nurick scores at short-term and medium-term assessments. Pooled mean differences showed clinically meaningful improvement in disability from baseline at short term (2 studies, MD 1.42) and medium term (4 studies, MD 1.06) follow-up.
Pain (VAS 100-point scale)	$\begin{array}{l} (14 - 730) \\ 6-12 \ months \\ 4 \ studies \ [20, \ 30, \ 34, \ 44] \\ (N = 646) \\ 13-36 \ months \\ 6 \ studies \ [20, \ 30, \ 34, \ 35, \\ 44, \ 47] \\ (N = 1097) \\ \geq 36 \ months \\ 1 \ study \ [20] \ (N = 268) \end{array}$	Moderate <sup>a</sup> 6-12 months Very Low <sup>b</sup> 13-36 and ≥36 months	
Key Question 2: Do these e	expected outcomes depend on p	reoperative disease	severity or duration of symptoms?
Preoperative Duration of Sy		· ·	
JOA improvement	12 months 2 studies [26, 39] (N = NC) 24-36 months 1 study [39] (N = 98) 54-60 months 2 studies [19, 39] (N = NC)	Very Low <sup>c</sup>	Change in neurological status following surgical intervention does not depend on preoperative duration of symptoms. Patients with a longer duration of symptoms ( $\geq$ 12 months) had similar neurological improvement as those with a shorter duration of symptoms (<12 months) at all follow-up periods.
Success (mJOA ≥16)	I study [41] (N = 272) F/U 12 months	Low	The odds of a "successful" outcome following surgical intervention depend on preoperative duration of symptoms. The odds of achieving a mJOA $\geq 16$ decreased by 22% when a patient moves from a shorter to a longer duration of symptoms group ( $\leq 3$ ; >3, $\leq 6$ ; >6, $\leq 12$ ; >12, $\leq 24$ ; >24 months).
Preoperative Myelopathy Se			
JOA improvement	I study [22] (N = 260) F/U 12 months	Moderate <sup>d</sup>	Change in neurological status following surgical intervention depends on preoperative disease severity. Less improvement on the mJOA was observed among patients with milder symptoms at presentation (preoperative mJOA ≥15: 1.29 [95% CI 0.70, 1.87]; preoperative mJOA 12-14: 2.58 [2.07, 3.09]; preoperative mJOA <12: 4.91 [4.34, 5.49]).
	2 studies [37, 41] (N = 332) F/U 12 months	Low	The odds of a "successful" outcome following surgical intervention depends on preoperative disease severity. The odds of achieving a mJOA $\geq$ 16 were 1.22 times greater for every 1-point increase in preoperative mJOA score.
% improved mJOA	I study [20] (N = 268) F/U 96 months	Very low <sup>b</sup>	Likelihood of neurological improvement following surgical intervention does not depend on preoperative disease severity. Similar fractions of patients improved by 1 mJOA point when comparing patients with moderate (mJOA 10-13, 86.7%) and severe (mJOA 5-9, 86.5%) disease.
NDI improvement	I study [22] (N = 260) F/U 12 months	Very low <sup>c</sup>	Similar improvements in NDI across preoperative disease severities following surgical intervention. At 12-months follow-up, NDI changes were 12.1 (mild mJOA $\geq$ 15), 9.8 (moderate mJOA 12-14), and 12.5 (severe mJOA <12).
Nurick improvement	I study [22] (N = 260) F/U 12 months	Very low <sup>c</sup>	Similar improvements in Nurick scores across preoperative disease severities following surgical intervention. At 12-months follow-up, the Nurick score changes were 1.6 (mild mJOA ≥15), 1.5 (moderate mJOA 12-14), and 1.7 (severe mJOA <12).
Key Question 3: What are	the complications associated wit	h surgical interventi	ion?
C5 radiculopathy or palsy	15 studies [17, 20, 21, 23- 25, 27, 28, 30-32, 36, 40, 42, 48] (N = 2661)	Low	Pooled cumulative incidence of C5 radiculopathy or palsy is 1.9% (95% CI 1.4, 2.4).

# Table 6. (continued)

#### Table 6. (continued)

	Studies; N [References]	Strength of Evidence	Conclusions/Comments
Infection	10 studies [20, 21, 24, 27, 29-31, 33, 42, 48] (N = 2074)	Low	Pooled cumulative incidence of infection is 1.5% (95% CI 1.0, 2.1)
Reoperation	7 studies [20, 21, 23, 29, 30, 32, 35] (N = 943)	Very low <sup>e</sup>	Pooled cumulative incidence of reoperation 1.4% (95% CI 0.6, 2.1)
Dural tear/CSF leak	I studies [20, 21, 24, 27, 28, 30-34, 43, 45] (N = 1893)	Very low <sup>c</sup>	Pooled cumulative incidence of dural tear is 1.4% (95% CI 0.8, 1.9)
Worsening of myelopathy	2 studies [21, 48] $(N = 781)$	Very low <sup>c</sup>	Pooled cumulative incidence of worsening of myelopathy is 1.3% (95% CI 0.5, 2.1)
Death	6 studies [17, 18, 21, 30, 31, 48] (N = 1162)	Very low <sup>c</sup>	Pooled cumulative incidence of mortality is 0.3% (95% CI 0.0, 0.5)
Fracture	1  study [27]  (N = 141)	Very low <sup>c</sup>	Pooled cumulative incidence of fracture is 2.1% (95% CI 0.0, 4.5)

Abbreviations: CI, confidence interval; CSF, cerebrospinal fluid; F/U, follow-up; JOA, Japanese Orthopaedic Association score; mJOA, modified Japanese Orthopaedic Association score; NDI, Neck Disability Index.

<sup>a</sup>Upgraded I for magnitude of effect.

<sup>b</sup>Downgraded I for serious risk of bias.

<sup>c</sup>Downgraded I for serious risk of imprecision.

<sup>d</sup>Upgraded I for dose response.

<sup>e</sup>Downgraded I for serious risk of bias and imprecision.

# Conclusions

Based on our conclusions, in patients with DCM, surgery prevents further disease progression and also results in significant gains in functional impairment, disability, and pain. A shorter duration of symptoms and less severe myelopathy preoperatively are both important predictors of achieving a postoperative mJOA  $\geq 16$ . Finally, surgery for DCM is a relatively safe treatment option, with a cumulative incidence of complications estimated at 14.1%.

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#### Supplemental Material

The supplemental material is available in the online version of the article.

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