Disc degeneration after disc herniation: are we accelerating the process?

Authors Josh E Schroeder¹, Joseph R Dettori², Erika D Brodt², Leon Kaplan¹

Institutions ¹ Orthopedic Department, Hadassah Hebrew University Medical Center Jerusalem, Israel ² Spectrum Research Inc, Tacoma, WA, USA

ABSTRACT

Study design: Systematic review.

- *Study rationale:* Disc degeneration is a common process starting early in life. Often disc herniation is an early step in disc degeneration, which may cause pain or stenosis. How quickly this subsequent disc degeneration occurs following a disc herniation and subsequent surgical treatment and whether certain spinal procedures increase the rate of degeneration remain unclear.
- *Objectives:* To investigate the risk of subsequent radiographic disc degeneration following discectomy, discography, and conservative care in patients with a first-time diagnosed herniated nucleus pulpous (HNP) and to ascertain whether this risk in these defined groups changes over time.
- *Methods:* A systematic review of pertinent articles published up to June 2012. Key articles were searched to identify studies evaluating the risk of subsequent radiographic disc degeneration following treatment for HNP. Studies that included patients undergoing secondary surgery for disc herniation or that did not use a validated classification system to measure the severity of disc degeneration were excluded. Two independent reviewers assessed the strength of evidence using the GRADE criteria and disagreements were resolved by consensus.
- **Results:** From a total of 147 possible citations, three cohort studies (class of evidence III) met our inclusion criteria and form the basis for this report. The risk of subsequent lumbar disc degeneration following standard discectomy was significantly greater compared with both microdiscectomy (48.7% vs 9.1%) and asymptomatic controls (90% vs 68%) in two studies with mean follow-ups of 5.5 and 25.3 years, respectively. Following conservative care for first-time HNP in the third study, the risk of progression of lumbar disc degeneration was 47.6% over the first 2 years of follow-up and 95.2% over the next 6 years of follow-up. In the same study, the risk of lumbar disc degeneration was shown to increase incrementally over the course of the 8-year follow-up, with all patients showing signs of degeneration at final examination.
- *Conclusion:* Standard discectomy in first-time lumbar HNP may increase the risk of subsequent same-level lumbar disc degeneration compared with microdiscectomy as seen in one low-quality study. However, disc degeneration is likely a natural, temporal consequence following HNP, as demonstrated in a second low-quality study. The overall strength of evidence for the conclusions is very low.

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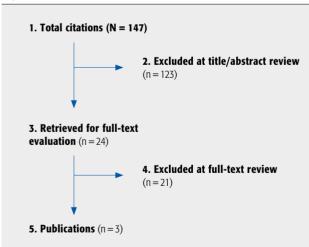
STUDY RATIONALE AND CONTEXT

Disc degeneration is a common process starting in the second decade of life [1, 2]. We commonly use imaging tests to quantify its severity [3]. In many cases, disc herniation is an early step in disc degeneration [4]. As the disc progressively degenerates, it may clinically manifest itself with pain or stenosis. How quickly this subsequent disc degeneration occurs following treatment for disc herniation and whether certain spinal procedures increase the rate of degeneration remain unclear. Of particular concern is if nonoperative care may be associated with a slower rate of progression than invasive diagnostic or surgical interventions, and if certain less invasive procedures offer advantages over more conventional extensile procedures.

OBJECTIVES

To investigate the risk of subsequent radiographic disc degeneration following discectomy, discography, or conservative care in patients with a first-time diagnosed herniated nucleus pulpous (HNP) and to ascertain whether this risk in these defined groups changes over time.

Fig 1 Results of literature search.



MATERIALS AND METHODS

Study design: Systematic review.

Search: PubMed; Cochrane collaboration database; National Guideline Clearinghouse database; bibliographies of key articles.

Dates searched: 1970 through June 4, 2012.

Inclusion criteria: (1) adults, (2) discectomy, discography or conservative treatment for first-time HNP.

Exclusion criteria: (1) secondary surgery for HNP, (2) fusion surgery, (3) tumor, (4) infection, (5) no disc severity classification system to measure disc degeneration, and (6) clinical outcomes only reported.

Outcomes: Severity classification system of disc degeneration (Kambin severity scale or Pfirrmann classification).

Analysis: Descriptive statistics of the cumulative incidence or prevalence.

Cumulative incidence: The proportion of people who had been degeneration-free at the time of the initial evaluation that had subsequent development of new degeneration at final follow-up.

Prevalence: The proportion of people with disc degeneration at any follow-up point.

Data were summarized in tables and figures but were not pooled due to the limited number of studies available and the heterogeneity of patient populations, outcome measures, and follow-up periods.

Overall strength of evidence: Risk of bias determined using CoE rating scheme developed by the Oxford Centre for Evidence-based Medicine by *The Journal of Bone and Joint Surgery American Volume* [Wright] (see Web Appendix at *www.aospine.org/ebsj*). The overall body of evidence with respect to each key question was determined based on modified precepts outlined by the Grades of Recommendation Assessment, Development and Evaluation (GRADE) system.

35

RESULTS

- From 147 citations, 24 articles underwent full-text review; three cohort studies, all graded CoE III, met our inclusion criteria and form the basis for this report (**Fig 1**). Characteristics of each study are outlined in **Table 1**. See Web Appendix at *www.aospine.org/ebsj* for critical appraisal and a list of excluded articles.
- We found no studies reporting the risk of radiographic disc degeneration following discography for HNP.
- Two studies investigated the risk of disc degeneration following standard discectomy; one compared with posterolateral microdiscectomy [5] and one compared with asymptomatic controls [6]. The third study was conducted in patients who had undergone conservative care [7].

Key question 1: What is the risk of subsequent radiographic disc degeneration following discectomy, discography, and conservative care in patients with a first-time diagnosed HNP?

Standard discectomy vs posterolateral microdiscectomy (Fig 2, Table 2)

- The discectomy (n = 39) and microdiscectomy (n = 44) groups differed regarding gender (males: 69.2% vs 39.5%) and percentage follow-up (78.0% vs 88.0%), respectively.
- Patients who underwent standard discectomy showed a fivefold greater cumulative incidence of moderate to

severe (grade \geq 2, Kambin severity scale) lumbar disc degeneration at a mean of 5.5 years' follow-up than those who underwent microdiscectomy: 48.7% vs 9.1% (relative risk [RR] = 5.4; 95% confidence interval [CI], 2.0–14.4; *P*=.0001).

• In the standard discectomy group, only 20.5% of patients showed no signs of subsequent disc degeneration (grade 0) whereas in the microdiscectomy group, most patients (61.4%) showed no subsequent degeneration.

Standard discectomy vs asymptomatic controls (Fig 3, Table 2)

- The two groups (n = 50 each) were matched for age and gender with most patients being men (66.0%) with a mean age of 64 years.
- The prevalence of moderate to severe (grade \geq 2, Kambin severity scale) lumbar disc degeneration at a mean follow-up of 25.3 years was significantly greater in the discectomy group (90%) compared with the asymptomatic control group (68%) (RR=1.3; 95% CI, 1.1–1.6; *P*=.012).

Conservative treatment

- Of 21 patients treated conservatively, 38.1% were men with a mean age of 49 years.
- The prevalence of progression of disc degeneration, defined as an increase of 1–2 grades, was 47.6% from initial presentation to 2 years, and 95.2% from 2 years to final follow-up. When defined as an increase of two grades, the prevalence of progression was 4.8% and 47.6%, respectively.

Fig 2 Cumulative incidence of lumbar disc degeneration (Kambin severity scale) at a mean follow-up of 5.5 years following standard discectomy and microdiscectomy.

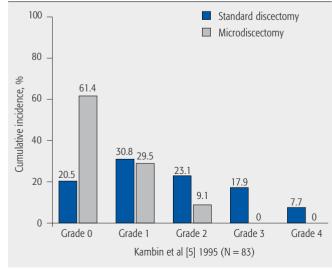
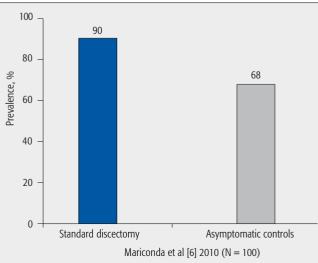


Fig 3 Prevalence of moderate to severe lumbar disc degeneration (grade ≥ 2 on the Kambin severity scale) at a mean follow-up of 25.3 years following standard discectomy and among asymptomatic controls.



Author (y)	Study design	Treatment	Population	Follow-up (% followed up)	Definition of radiographic spinal degeneration
Kambin et al [5] (1995)	Cohort	Posterior laminotomy and discectomy (group 1) Posterolateral	Group 1 - N=39 - Male: 69.2% - Mean age: 40.3 ± 2.0 y	Group 1: mean 5.4 y (78%; 39/50)	Kambin severity scale (radiograph): Grade 0: absence of degenerative changes Grade 1:
		microdiscectomy (group 2)	Group 2 - N=44 - Male: 39.5%	Group 2: mean 5.6 y (88%; 44/50)	marginal osteophytes, absence of disc space narrowing and normal facet joints Grade 2:
			– Mean age: 41.9±1.8 y	83% (83/100)	narrowing of the intervertebral disc space $\leq 33\%$ associated with marginal osteophytes Grade 3: narrowing of the intervertebral disc space $> 33\%$ in the presence of large marginal osteophytosis and hypertrophic changes of facet joints Grade 4: complete collapse of the disc space and/or the
Mariconda et al	Cohort	Standard discectomy	Group 1	Mean 25.3 ± 3.0 y	development of degenerative spondylolisthesis Kambin severity scale (radiograph)
[6] (2010)		(group 1)	- N=50 - Male: 66.0%	% Not reported	,
		Matched, asymptomatic nonoperated	- Mean age: 63.4±8.4 y Group 2		
		subjects (group 2)	 N=50 Male: 66.0% Mean age: 64.7±9.6 y 		
Masui et al [7] (2005)	Cohort	Conservative treatment	N=21 Male: 38.1% Mean age: 49	Mean 8.1 (range, 7.3–10.0) y	Pfirrmann classification (MRI): Grade I
			(range, 24–68) y	52.5% (21/40)	 Structure: homogeneous, bright white Distinction of nucleus and anulus: clear Signal intensity: hyperintense, isointense to cerebrospinal fluid Disc height: normal Grade II Structure: inhomogeneous with or without horizontal bands Distinction of nucleus and anulus: clear Signal intensity: hyperintense, isointense to cerebrospinal fluid Disc height: normal Grade III Structure: inhomogeneous, gray Distinction of nucleus and anulus: unclear Signal intensity: intermediate Disc height: normal to slightly decreased Grade IV Structure: in homogeneous, gray to black Distinction of nucleus and anulus: lost Signal intensity: intermediate to hypointense Disc height: normal to moderately decreased Grade V Structure: inhomogeneous, black Disc height: normal to moderately decreased Grade V Structure: inhomogeneous, black Disc height: normal to moderately decreased Grade V Structure: inhomogeneous, black Distinction of nucleus and anulus: lost Signal intensity: hypointense Disc height: normal to moderately decreased Grade V Structure: inhomogeneous, black Distinction of nucleus and anulus: lost Signal intensity: hypointense Disc height: collapsed disc space

Key question 2: Does this risk of subsequent radiographic disc degeneration in these defined groups change over time?

Conservative treatment (Fig 4, Table 2)

• The prevalence of moderate to severe (grade≥III, Pfirmann classification) lumbar disc degeneration increased incrementally over the study period from 28.6% at initial presentation to 66.6% at 2 years, and finally to 100% at the 8-year follow-up.

Fig 4 Prevalence of moderate to severe lumbar disc degeneration (grade ≥ III using the Pfirrmann classification) at initial presentation, 2 years, and 8 years following conservative treatment.

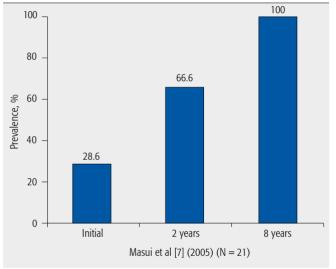


 Table 2
 Summary of reported risks of subsequent disc degeneration following various treatments for lumbar herniated nucleus pulpous.

Author (y)	Outcome	Definition of	Population	Grade, %					Follow-up,
		degeneration		0	1	2 3		4	mean
Kambin et al [5] (1995)	Cumulative incidence	Kambin severity scale	le Standard discectomy		30.8	23.1	17.9	7.7	5.5 y
			Microdiscectomy	61.4	29.5	9.1	0	0	
Mariconda et al [6] (2010)	Prevalence	Kambin severity scale	Standard discectomy	4.0	6.0	26.0	58.0	6.0	25.3 y
			Controls	10.0	22.0	26.0	26.0	16.0	
				I	П	ш	IV	v	
Masui et al [7] (2005)	Prevalence	Pfirrmann classification	Conservative care	9.5	61.9	23.8	4.8	0	Initial
				0	33.3	57.1	9.5	0	2 у
				0	0	9.5	61.9	28.6	8.1 y

CLINICAL GUIDELINES

No clinical guidelines were found addressing this topic.

Table 3 Strength of	evidence summa	ary.					
	Strength of evidence				Conclusions/comments		
Key question 1: What with a first-time diagr				ition followi	ng discectomy, discography, and conservative care in patients		
Discectomy vs microdiscectomy	Very low	Low	Moderate	High	Subsequent lumbar disc degeneration occurred five times more often following standard discectomy compared with microdiscectomy (48.7% vs 9.1%; mean follow-up, 5.5 years) in one low-quality study		
Discectomy vs asymptomatic controls	Very low	Low	Moderate	High	A significantly greater risk of subsequent lumbar disc degeneration was seen in the discectomy group compared with the control group (90% vs 68%; mean follow-up, 25.3 years) in one low-quality study		
Conservative treatment	Very low	Low	Moderate	High	The risk of progression of disc degeneration, defined as an increase of 1–2 grades, was 47.6% from initial presentation to 2 years, and 95.2% from 2 years to final follow-up at 8 years in one low-quality study		
Key question 2: Does	risk of subsequen	t radiographic	disc degeneration in	1 these defin	ed groups changes over time?		
Discectomy, microdiscectomy	No evidence				None		
Conservative treatment	Very low	Low	Moderate	High	In one low-quality study, the risk of lumbar disc degeneration increased incrementally during follow-up: initial presentation, 28.6%; 2 years, 66.6%; and 8 years, 100%		

EVIDENCE SUMMARY

ILLUSTRATIVE CASE

The overall strength of evidence evaluating the risk of subsequent radiographic degeneration in the lumbar spine following standard discectomy compared with microdiscectomy, standard discectomy compared with asymptomatic controls, and conservative treatment for first-time HNP is very low, that is, any effect estimate is uncertain (**Table 3**). The overall strength of evidence evaluating whether this risk of subsequent radiographic disc degeneration changes over time is also very low, as addressed by only one study in patients undergoing conservative care. A 53-year-old healthy woman suffered from extreme radicular pain in her right leg in the S1 distribution. On magnetic resonance imaging (MRI), a central disc herniation was noted (**Fig 5**). She was treated with oral analgesics and physical therapy; however, the pain did not remit. Six months later she was referred for a discectomy. On surgery, a standard discectomy was performed and a large disc fragment was removed from the canal (**Fig 6**). Her pain resolved and she went back to her daily life. Nine months later, she returned to the clinic with a new onset of lower back pain and pain radiating down the left leg. A new MRI was performed showing loss of disc height, severe degeneration, and recurrent herniation of the disc (**Fig 7**).

Fig 5 Preoperative sagittal and axial magnetic resonance images of the spine. A L5-S1 central disc herniation is noted.



Fig 6 Intraoperative image of the disc tissue removed in surgery.



Fig 7 Sagittal magnetic resonance image taken 9 months postoperatively showing advancement to severe disc degeneration with positive Modic end plate changes and reherniation of the L5-S1 disc.



DISCUSSION

- Strengths:
 - We included only studies that used a severity classification system to measure disc degeneration.

• Limitations:

- Few studies were available to address the issue, all with small sample sizes (n=83, n=100, n=21).
- All studies were class of evidence III.
- Heterogeneity of patient populations, outcome measures, and follow-up periods made comparisons across studies difficult.
- No studies were found that met our criteria in patients treated with discography.
- Not all cases of HNP will lead to advanced disc degeneration in 5 to 10 years; however, close to two-thirds will show some signs of degeneration.
- Surgical intervention may increase the risk of disc degeneration.
- Standard discectomy may cause more degeneration than microdiscectomy.
- The biologics behind disc nutrition and degeneration and the effects of interventions on these processes need to be better understood.
- As discectomy is the most common spine procedure in the world, better data needs to be gathered on the long-term effects of surgical procedures.

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EDITORIAL PERSPECTIVE

The reviewers were positive about the idea for and the execution of this systematic review, noting that this seemingly important question had received surprisingly scant attention in the peerreviewed literature to date.

Again, this is less of a question for Schroeder and colleagues and more of an issue for the original authors of the pertinent studies. The question remains, whether a 'sequestrectomy' or 'fragmentectomy' (removing only the offending fragment) is different from a 'formal discectomy,' in which the intervertebral segment is entered with instruments and further loose nucleus and annular materials are removed, or if the choice of magnification system (operating microscope vs surgeon loupes) has an effect on the rate and severity of disc degeneration. The data presented seems to imply that 'less is more' from a disc degeneration perspective as long as the offending disc fragment is removed; clearly, aggressive intervertebral maneuvers such as end-plate curettage seem not supported by these studies.

In the big picture, this systematic review clearly outlines the inexorable progression of lumbar disc degeneration over time — with a stated 68% incidence over 25 years in asymptomatic controls. Sadly, it seems clear that disc herniation is a common, first major step into the degenerative vortex of disc degeneration.

The increasing availability of large-scale digital imaging repositories will, in the future, hopefully allow for more longitudinal assessment of the variables that surround disc degeneration.