

# Endplate changes following discectomy: natural history and associations between imaging and clinical data

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## Abstract

**Purpose** Some patients will experience post-operative back pain following lumbar discectomy, and the potential sources for that pain are poorly understood. One potential source is the vertebral endplates. The goal of this study was to document the changes that occur in lumbar endplates following discectomies, and to assess associations between endplate changes and clinical outcomes.

**Methods** Changes in lumbar endplates and discs were assessed from X-rays, CT and MRI exams by comparing preoperative imaging with imaging obtained at yearly intervals up to 5 years. 260 endplates in 137 patients with single-level herniation and discectomy were analyzed. The geometry of osseous defects in the endplates was measured

from the CT exams, and marrow and disc changes adjacent to endplates were assessed from the MRI exams. Clinical outcome assessments were collected at each time point. Descriptive statistics were used to describe endplate defect sizes, and logistic regression and analysis of variance were used to identify potential associations between endplate and vertebral body changes and clinical outcomes.

**Results** Approximately 14 % of the endplates had osseous defects prior to surgery. After surgery, 24 % of inferior and 43 % of superior endplates had defects. Change occurred within the first year and remained relatively constant over the next few years. Disc signal intensity worsened and disc height decreased following surgery. New Modic changes were also observed. None of these changes were associated with having achieved a clinically significant improvement in outcome scores. The follow-up rates were low at the later time points and significant associations cannot be ruled out.

**Conclusions** This study documents lesion characteristics in detail and supports that osseous defects in the endplates at the level of a lumbar discectomy may be a relatively common finding following surgery, along with disc height loss, loss of disc signal intensity, and Modic changes. The clinical significance of these imaging findings could not be conclusively determined in this study.

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## Introduction

Nearly half a million lumbar discectomies are performed in the United States each year [1] and although the clinical outcomes are generally good, there are complications

associated with the procedure including re-herniations, accelerated disc degeneration, and post-operative low back pain. The latter—post-operative low back pain—is relatively common but remains poorly understood. There are several possible sources of this back pain. One such source is the vertebral endplates which are known to be extensively innervated [2, 3].

There is evidence of an association between lesion size and back pain [4]. Endplates may be injured as a result of the original disc herniation [5–10] and there may also be changes that occur in the endplate that are directly associated with the surgical procedure itself [11]. Fragments of endplates can be found in extruded disc material [6, 12]. Thus, it might prove useful to fully delineate the endplate changes that occur following discectomy to better understand this as a potential source of pain. Some of the post-discectomy changes that can occur in the marrow adjacent to endplates have been described [13–15], but the specific osseous changes that can occur involving the subchondral plate of the endplate have not been delineated in detail.

Accordingly, the aim of this study was to provide detailed information regarding the endplate changes that can occur following lumbar discectomy. The preoperative endplate condition was used as the baseline internal control for pre-existing defects. Knowledge regarding subsequent endplate changes can help us to better understand this possible source of post-discectomy symptoms and may help identify possible intervention strategies aimed at reducing their occurrence.

## Methods

Pre- and post-discectomy X-rays, CT, and MRI exams were obtained for patients who had a single-level lumbar discectomy procedure performed at one of five clinical sites in Europe. A total of 137 patients were enrolled in the study. All patients signed informed consents as approved by the respective ethics committees.

The CT exams were carefully assessed for apparent endplate defects. Axial, sagittal, and coronal plane slices were reviewed. Multi-planar reconstructions were also used to obtain sections parallel and perpendicular to the endplate. Endplate defects were defined as any localized morphologic feature of the osseous anatomy that could not be explained by the overall shape of the endplate, recognizing that endplates can have a variety of morphologies [16]. Defects present pre-operatively were identified and measured. Defects present post-operatively were also identified and measured, with particular attention to any changes that occurred between pre- and post-operative CT exams. The superior and inferior endplates of the intervertebral disc were assessed separately.

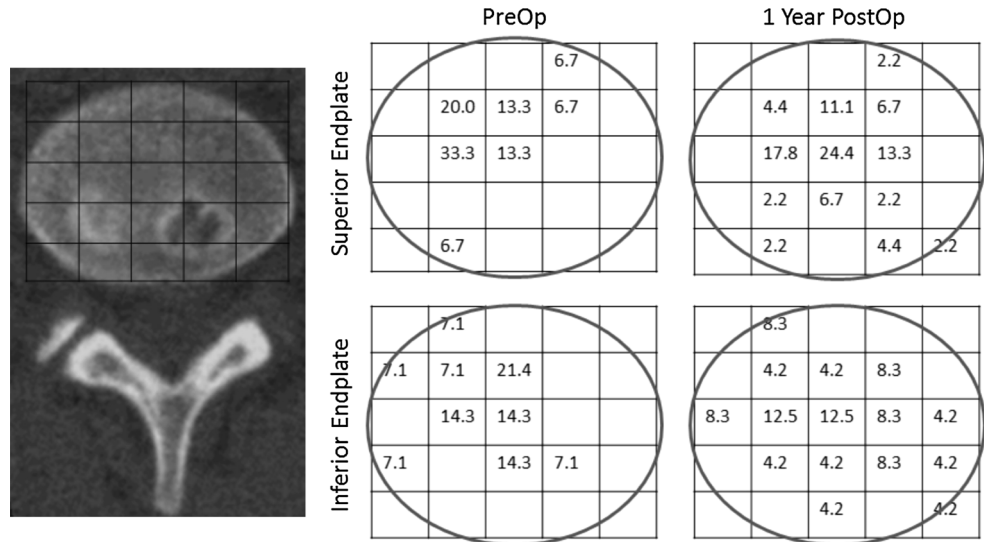
The total number of identifiable lesions was recorded. Quantitative measurements of lesion size were made for the two largest lesions in each endplate and these were labeled as the primary and secondary lesions. Defect size in the plane of the endplate was measured from sagittal and coronal plane slices and the depth of the defect was also measured perpendicular to the plane of the endplate in the sagittal and coronal planes. Each lesion was also approximated as an ellipse in the plane of the endplate using the axial, sagittal and coronal plane slices. The major and minor axis of each lesion were identified and measured. The depth was measured perpendicular to the plane of the endplate. Multi-planar reconstructions were used to assure that the measurement landmarks were consistent between sagittal, axial, and coronal views. In addition, the location of each defect was classified by the position on a  $5 \times 5$  grid virtually overlaid on to each endplate (Fig. 1). The grid assigned to each defect was the grid that contained the largest proportion of the defect.

In addition, MRI exams were used to assess disc signal intensity (T2-weighted images) using the Pfirrmann et al. [17] grading system (Table 1). Modic changes in the vertebral bodies adjacent to the treated disc were also assessed (Table 2) [18]. Disc height and spondylolisthesis were measured from neutral lateral X-rays. These measurements and the changes between time points were measured using previously validated computer-assisted software [19] (QMA<sup>®</sup>, Medical Metrics, Inc). All assessments (CT, MRI, X-ray) were done by radiologists at an independent core lab (Medical Metrics, Inc, Houston, TX, USA), with radiologists blinded to clinical outcomes.

In addition to the imaging assessments, the Oswestry Disability Index (ODI) and the Visual Analog Scale (VAS) for back pain were collected pre-operatively and at 1 and 2 years PostOp. The change in these scores between PreOp and PostOp was calculated for each patient. The patient was classified as having a good outcome if the ODI dropped by 12.8 points (with ODI scored on a scale where 100 = worst possible disability) or the VAS back pain score dropped by 12 points (scored on a scale where 100 = worst possible pain). These thresholds were based on a study of the Minimum Clinically Important Difference (MCID) in lumbar spine surgery patients [20].

Data were analyzed using descriptive statistics, analysis of variance, and by logistic regression analyses (Stata Ver 11, College Station, TX, USA) to determine if each measure of the number and size of the defects was associated with whether the patient received a good outcome at 1 year PostOp (a 12.8-point improvement in the ODI or 12-point improvement in the VAS back pain score). Ordered logistic regression was used to analyze the number of lesions per endplate as an ordinal variable. Logistic regression does not assume or require normality, linearity, and

**Fig. 1** Illustration of the 5 × 5 grid that was used to define the location of the endplate defects in the axial plane, and the proportion of defects located in each *grid box*. Data for the superior and inferior endplates are provided, both at PreOp (113 disc spaces examined) and at 1 year PostOp (11 disc spaces examined). The lesion location was the *grid box* that contained the largest proportion of the lesions. Pre-operatively, most of lesions were toward the center of the endplate, with a higher proportion of lesions located along the posterior border of the endplates following discectomy



**Table 1** Pfirmann grading system used for disc signal intensity assessments from MRI<sup>14</sup>

Grade	Description
Grade I	Bright white disc, homogenous structure, clear distinction between nucleus and annulus, hyperintense signal intensity, isointense to cerebrospinal fluid
Grade II	White disc, inhomogenous structure with or without horizontal bands, clear distinction between nucleus and annulus, hyperintense signal intensity, isointense to cerebrospinal fluid
Grade III	Gray disc, inhomogenous structure, unclear distinction of nucleus and annulus, intermediate signal intensity
Grade IV	Gray to black (dark gray) disc, inhomogenous structure, no distinction between the nucleus and annulus, intermediate to hypointense signal intensity
Grade V	Black disc, inhomogenous structure, no distinction between the nucleus and annulus, hypointense signal intensity

**Table 2** The Modic grading system used to assess changes adjacent to the endplates at the treated level following discectomy

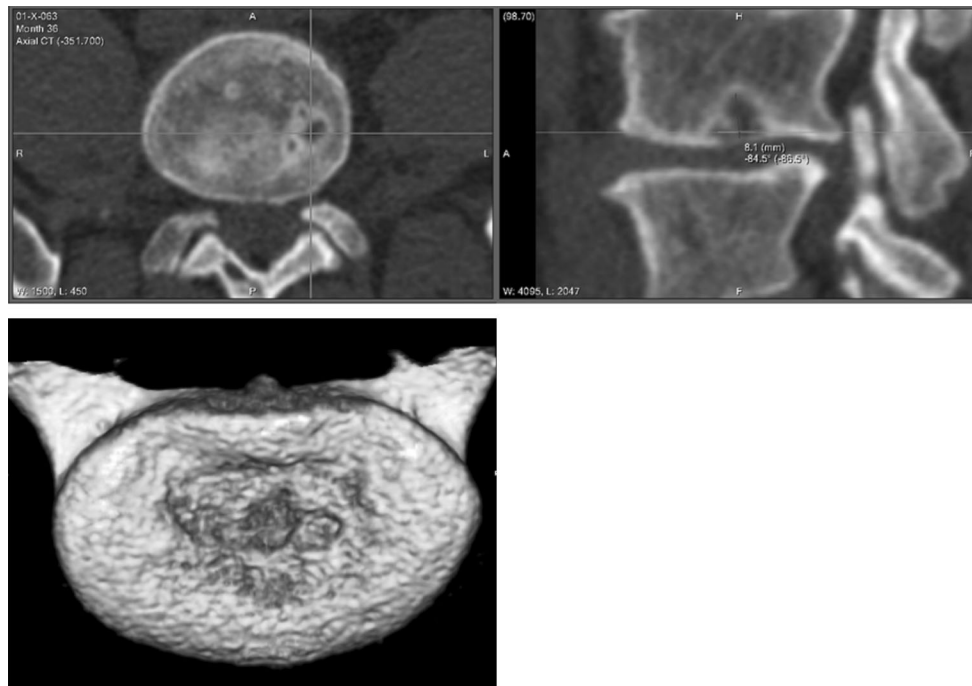
Grade	Description
None	No edematous reaction or vascular congestion induced in the adjacent bone marrow of the endplates
Type I	Hypointense reaction and vascular congestion in the adjacent marrows on T1-weighted MR imaging; hyperintense on T2-weighted images
Type II	Bone marrow converted to a predominantly fatty marrow. Hyperintense on T1 and isointense to hypointense on T2. The exact signal intensity depends on the degree of T2 weighting
Type III	Dense sclerosis of the vertebral endplates. Reflected on MR imaging as hypointensity on both T1 and T2-weighted images

homogeneity of variance for the independent variables, and can assess for the significance of relationships between the dependent variable and metric or dichotomous independent variables.

**Results**

Some of the endplates could not be reliably assessed because multi-planar reconstructions were not possible due

to variable slice thickness or gantry tilt. Reliable CT-based assessments could be made on endplates of 113 disc spaces at PreOp, 111 disc spaces at 1 year, 68 disc spaces at 2 and 3 years, and 38 disc spaces at 4 years. Using MRI, disc signal intensities and Modic changes could be assessed in 52 patients at PreOp, 86 patients at 1 year, 44 at 2 years, 26 at 3 years, 16 at 4 years, and 7 at 5 years. The change in disc height relative to PreOp could be calculated for 130 patients at 1 year and 78, 66, 34, and 10 patients at 2, 3, 4, and 5 years. Clinical outcome scores were available for



**Fig. 2** Examples of endplate defects that were observed following discectomy. In all of these three patients, there was no defect in the endplate pre-operatively. The *top two* images are axial and sagittal plane slices through a defect in one patient at 3 years PostOp. The

*bottom figures* shows a three-dimensional reconstructions of vertebral endplates from an additional patient, created from a thin-slice CT scan taken 3 years after discectomy

136 patients at PreOp, 94 patients at 1 year, and 38 patients at 2 years. The statistical tests involving outcomes data should be interpreted with an appreciation of the low follow-up. It is possible that patients who did not return for follow-up may differ from those that did.

#### Endplate bony defects

The defects tended to be located near the center of the endplate at PreOp (Fig. 1). Following discectomy, however, new endplate defects were also commonly observed in the posterior regions of the endplate. The numbers of defects in each zone were too small for meaningful statistics comparing distributions within zones, so the observations should be viewed as trends. A range of osseous endplate defect morphologies was observed. In general, these defects were similar to the morphologies described in the literature on Schmorl's nodes [21–25]. Examples of endplate lesions observed pre- and post-operatively are shown in Fig. 2. There were no observations that would suggest that the observed lesions were morphologically different at PostOp compared to PreOp or that the defects were morphologically different than what has been described as a typical Schmorl's node [21–25].

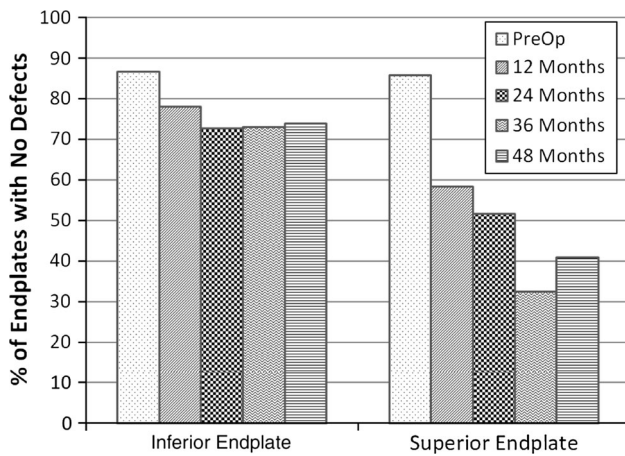
The proportion of superior and inferior endplates that were free of endplate lesions at each time point decreased

in the first year after surgery but did not appear to change in the subsequent years (Fig. 3). This early change was more pronounced in the superior endplate of the treated disc space, where there was almost 50 % fewer superior endplates that were free of defects following surgery. Based on ordered logistic regression, the change from PreOp to 1 year in number of endplate lesions was nearly significant at the inferior endplates ( $P = 0.079$ ) and was highly significant at the superior endplates ( $P < 0.0001$ ).

The average size of the primary and secondary lesions is summarized in Table 3. With data for superior and inferior endplates pooled, there was no difference in the size of the lesions measured from PostOp versus PreOp CT scans ( $P = 0.51$ ). Analysis of variance tests indicated no significant difference in the size of the lesions in the superior versus inferior endplates at PreOp ( $P = 0.36$ ); however, the defects tended to be larger in the inferior versus superior endplates at 1 year PostOp ( $6.3 \pm 2.9$  mm versus  $4.9 \pm 1.6$  mm,  $P = 0.047$ ).

#### Clinical outcomes

Pre-operatively, the ODI score averaged  $49.9 \pm 17.8$  for the 136 patients who completed the questionnaire. At 1 year, ODI averaged  $19.3 \pm 16.6$  for the 94 patients who completed the questionnaire. The average change in ODI



**Fig. 3** The percent of endplates with no defects decreased following discectomy, but did not appear to change with time following surgery. Data based on assessing 113 disc spaces at PreOp, 111 disc spaces at 1 year, 68 disc spaces at 2 and 3 years, and 38 disc spaces at 4 years. Note that there is relatively low follow-up at later time points

was  $-30.4 \pm 21.6$  at the 1-year follow-up and  $-32.6 \pm 23.0$  at the 2-year follow-up. At the 1- and 2-year follow-up, 84.0 and 81.6 % of patients had achieved at least a 12.8-point improvement in the ODI score, 54.3 and 65.8 % had achieved at least a 12-point improvement in the VAS back pain score, and 88.3 and 86.8 % had achieved a MCID in either the ODI or the VAS back scores.

Table 4 provides the statistical significance value for logistic regression tests of whether each specific measurement of lesion number or size was independently associated with whether a patient received at least a 12.8 point improvement in the ODI score 1 year PostOp. There were no variables that were found to be close to significant. Similar tests were run using an improvement in VAS back

of  $\geq 12$  points as the measure of whether a patient achieved a good clinical outcome. No measure of the number or size of endplate defects was associated with this definition of a good outcome ( $P > 0.50$  for all tests).

**MRI assessment of degeneration**

The MRI exams document evidence of progressive disc degeneration following discectomy. The intervertebral disc signal intensity data show a trend toward loss of disc hydration over time (Fig. 4). Based on analysis of variance tests, however, there was no statistically significant evidence that disc signal intensity was associated with the change in clinical outcomes following discectomy ( $P > 0.11$  for all outcome scores). Similarly, logistic regression analysis indicated no association between disc signal intensity and whether a patient had at least a 12.8 improvement in the ODI score 1 year PostOp.

Disc height measurements also support a progressive degeneration of the intervertebral disc following discectomy. Disc height significantly decreased with time following discectomy, based on the change in disc height relative to Preop, that was calculated on a per-patient basis and then averaged (Fig. 5,  $P = 0.014$ , oneway analysis of variance). Using  $>1$  mm as the threshold to define a significant loss in disc height relative to PreOp, the data suggest that approximately 1/2 of the patients have this loss within the first year, with an additional 25 % of patients developing this loss over subsequent years (Fig. 6). There was no evidence of a significant change in spondylolisthesis, on average, in the first 5 years following discectomy ( $P = 0.19$ ). Two patients developed a change in spondylolisthesis  $>10$  % of the endplate during the course of the study.

**Table 3** Descriptive statistics for primary endplate defect dimensions

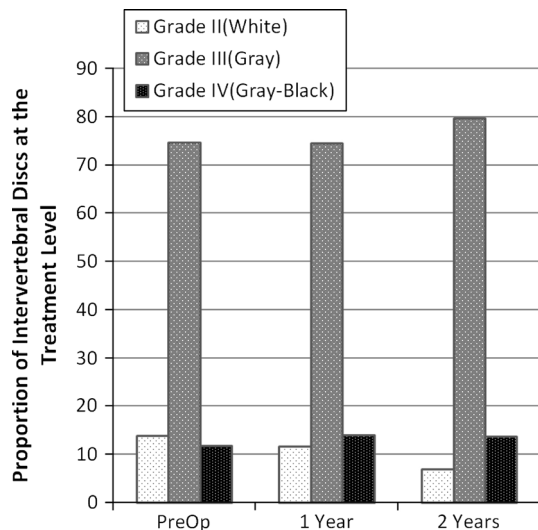
Dimension	Preop	1 year
Measured from sagittal plane slices		
Maximum width in the anterior–posterior direction	5.5 (1.9) range 2.6–8.6	4.8 (1.6) range 2–9.4
Depth perpendicular to the plane of the endplate	3.3 (1) range 2 to 5.1	3.1 (1) range 1.4–5.6
Measured from coronal plane slices		
Maximum width in the anterior–posterior direction	4.7 (2) range 2.5–11.1	4.8 (2.2) range 1.8–14.3
Depth perpendicular to the plane of the endplate	2.9 (0.9) range 2–5.5	3 (1.1) range 1.1–5.6
Major and minor axis		
Major axis	5.8 (2.3) range 2.6–11.1	5.4 (2.2) range 2–14.3
Minor axis	4.5 (1.4) range 2.5–6.9	4.2 (1.4) range 1.8–8.8
Depth perpendicular to the plane of the endplate	2.7 (0.7) range 2–4.2	2.8 (0.9) range 1.1–4.9

The measured dimensions of the endplate defects are summarized below (based on 9 defects measured at PreOp and 69 measured at 1 year). The mean value is provided along with standard deviation (in brackets) and the range. Data for the superior and inferior endplates are combined as no significant differences between endplates could be identified with the available sample size. All measurements are in units of millimeters

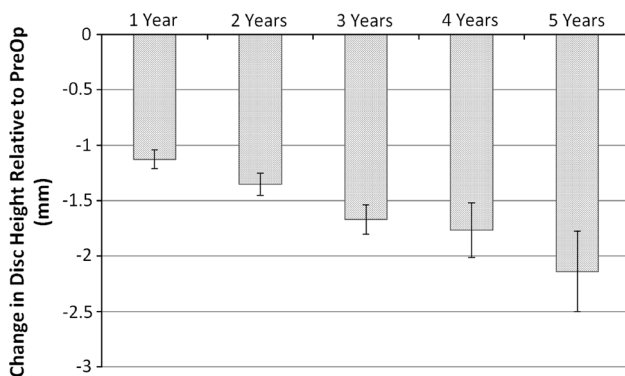


**Table 4** Significance levels for logistic regression tests to determine if a measurement of lesion size was significantly associated with whether the patient achieved at least a 12.8-point improvement in the ODI score

Slice plane	Measurement	Value at 1 year
All	Number of lesions	0.61
Sagittal	Maximum AP size	0.51
Sagittal	Depth perpendicular to plane of endplate	0.67
Coronal	Maximum width	0.92
Coronal	Depth perpendicular to plane of endplate	0.67
Whichever shows the greatest size	Major axis	0.72
Whichever shows the greatest size	Minor axis	0.67
Whichever shows the greatest size	Depth perpendicular to plane of endplate	0.54



**Fig. 4** Based on the T2-weighted MRI exams, an increasing proportion of discs showed evidence of loss of disc signal intensity with time following discectomy



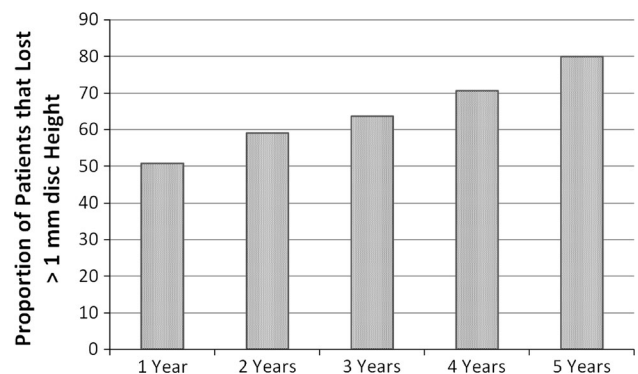
**Fig. 5** The change in intervertebral disc height, relative to disc height at PreOp. The error bars show  $\pm 1$  standard error. Data based on measurements for 130 discs at 1 year, 78 at 2 years, 66 at 3 years, 34 at 4 years, and 10 at 5 years. Note the low follow-up at later time points

The assessments of Modic changes adjacent to the endplates at the treated level support several observations (Fig. 7), including: (1) most endplates having no Modic

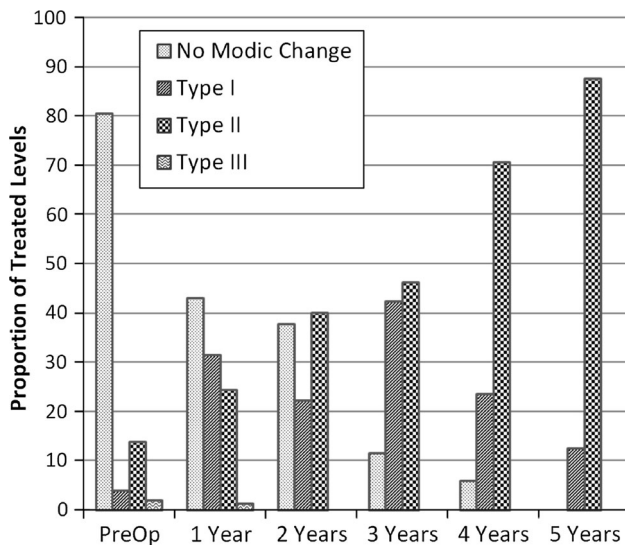
changes at PreOp; (2) 30–40 % developing type I changes during the first 3 years, with this proportion declining during the 4th and 5th years; (3) a progressive increase to where nearly 80 % of patients have type II changes by 5 years PostOp. There was no evidence for a significant relationship between the Modic change and clinical outcome scores. Interestingly, there was a significant association between loss of PreOp disc height and the Modic grade 1 year following discectomy ( $P = 0.02$ , Fig. 8). Having multiple measurable defects in the endplates was a significant ( $P < 0.0001$ ) risk factor for having Type I Modic changes.

**Discussion**

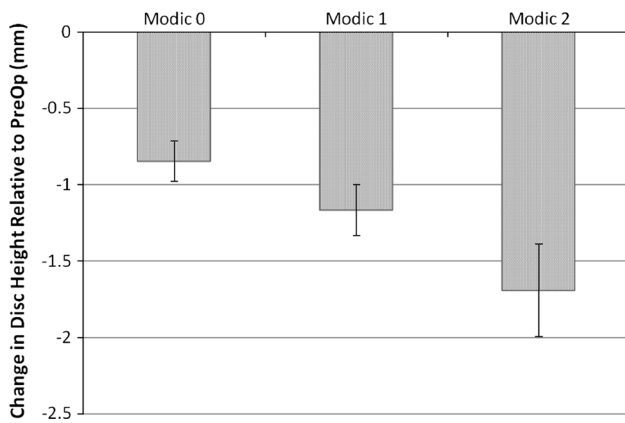
New endplate defects were common following limited lumbar discectomy involving removal of free and loose fragments outside and within the disc space with no direct intentional damage to the endplates. It is not known how many of these patients would have had new lesions if they had not had a discectomy. However, the natural history data for Schmorl’s nodes [26, 27] and our understanding of the rate of endplate marrow changes [28] would suggest that endplate changes generally progress slowly over time.



**Fig. 6** The proportion of patients that lost at least 1 mm of disc height at the treated level following discectomy. Note the low follow-up at later time points



**Fig. 7** The proportion of treated levels with MRI evidence of a Modic change adjacent to the endplates. Note that there is relatively low follow-up at later time points



**Fig. 8** The average change in intervertebral disc relative to PreOp for the different grades of Modic changes, 1 year following discectomy. The error bars show  $\pm 1$  standard error

A study by Wu et al. was the only previous study found reporting significant endplate changes over the course of a few years and they found this in only a small proportion of patients studied [27]. Barth et al. [14] reported that endplate degeneration was observed in 47 % of microdiscectomy patients and 14 % of microscopic sequestrectomy patients 2 years after surgery. These proportions span the proportion found in the current study. The results of the Barth et al. suggest an influence of surgical technique on endplate changes.

Despite the apparent spike in the proportion of endplates with lesions, there was no evidence that the presence of defect or any measurement of defect size was associated with clinical outcomes. This observation is limited by the low follow-up at later time points. A wide range of defect

size measurements were explored to assure that a critical size measurement was not missed. Size of defect does not appear to determine patient satisfaction. Jensen et al. found that preoperative Modic lesions can significantly affect clinical outcomes [29], suggesting the possibility that the marrow changes may be clinically more important than osseous defects in the subchondral plate.

Strengths of this study include a thorough quantitative assessment of osseous defects. To assure that the true dimensions were measured as accurately as possible, the lesion dimensions were made from sagittal plane and coronal plane slices from the CT exams, with axial-plane slices used when they provided the best visualization of defect size. (This also resulted in lesser quality CT studies being excluded). Consistency of some results with previously published studies was also noted. The proportions of patients with Modic changes at the different time points are similar to those reported by Rahme et al. [30]. At PreOp, Rahme et al. reported a higher proportion of patients with any type of Modic change (46 versus 20 %), while at their median follow-up period of 41 months, they reported type II changes in 63 % of patients whereas we found Type II changes in 50 % of patients at 3 years and 66 % at 4 years. Rahme et al. reported that all of their discectomy patients had some evidence of disc height loss at a median follow-up of 41 months, consistent with our observation of 80 % having >1 mm height loss by 5 years. Rahme et al. also found no significant correlation between Modic changes and symptoms. Ohtori et al. also reported no association between Modic changes and clinical outcomes following discectomy surgery [31].

This study has several limitations. While all patients had limited discectomies, the discectomies were performed at five different sites and there may have been differences in techniques used to perform the discectomy. The follow-up rate was less than ideal at later time points, but good early on and the PreOp CT and MRI images afforded a solid internal control. That said, there was no non-operative control group so it is not known how many would have had significant changes without surgery. It is also not known whether defects may have been a consequence of a specific surgical technique. This was a physician-sponsored study and dedicated resources were not available for encouraging patients to return for follow-up. It is not known why some patients agreed to return for follow-up while others did not, or how inclusion of only those patients who volunteered for follow-up may have biased the results. In addition, it would have been of value to also have assessments of the adjacent levels to better understand the natural history of endplate changes. Resource limitations did not allow for those assessments. The sites did not provide details on patient sex and age, since it was not expected that the study would have the

statistical power to analyze for sex- or age-related differences. It is assumed that the patients represent the typical population of patients undergoing discectomy surgery. Potential interactions between variables were not addressed in the current study, and it is possible that pre-existing Schmorl's nodes or PreOp degenerative changes might have affected the occurrence of new defects or endplate changes. A much larger sample size would be required to assess such interactions. And finally, due to sample size, the statistical power does not allow us to state definitively that there is truly no association between the presence or size of a defect or endplate change and PostOp symptoms. With a much larger sample size it is possible that a subgroup of patients could be identified where the endplate lesion is associated with symptoms. Nevertheless, this study provides new observations on the characteristics of lesions that may form following discectomy. The observations need to be interpreted with respect to the limitations of the study.

## Conclusion

Using the preoperative condition of each endplate as its own control, the proportion of endplates with osseous defects increased in the first year following surgery with no changes detected over the next 3 years, based on the lesions detected from CT scans obtained pre-operatively and following lumbar discectomy. The change is much more pronounced in the superior endplate of the treated disc space. New post-discectomy defects tended to be slightly larger but otherwise morphologically similar to lesions observed at PreOp. Consistent with prior studies, the new endplate changes were not significantly correlated with clinical outcomes. However, due to the unknown outcome of patients lost to follow-up, the association between new endplate changes and clinical outcomes remains uncertain.

**Conflict of interest** Patient follow-up and data analysis were supported by Intrinsic Therapeutics, Inc.

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