TRANSITING NERVE ROOTLET ABNORMALITIES ON MRI AFTER LUMBAR LAMINECTOMY: ASSOCIATIONS WITH PERSISTENT POSTOPERATIVE PAIN

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OBJECTIVE: To determine whether changes in the transiting nerve rootlet or its surroundings, as seen on MRI performed after lumbar hemilaminectomy, are associated with persistent postoperative pain (PPP), commonly known as the failed back surgery syndrome.

MATERIALS AND METHODS: Seventy-three patients (mean age, 61 years; 43 males and 30 females) who underwent single-level partial hemilaminectomy of the lumbar spine without postoperative complications or other level spinal abnormalities between January 2010 and December 2018 were enrolled. Two musculoskeletal radiologists evaluated transiting nerve rootlet abnormalities (thickening, signal alteration, distinction, and displacement), epidural fibrosis, and intrathecal arachnoiditis on MRI obtained one year after the operations. A spine surgeon blinded to the radiologic findings evaluated each patient for PPP. Univariate and multivariate analyses were used to evaluate the association between the MRI findings and PPP.

RESULTS: The presence of transiting nerve rootlet thickening, signal alteration, and ill-distinction was significantly different between the patients with PPP and those without, for both readers \((p \leq 0.020)\). Conversely, the presence of transiting nerve rootlet displacement, epidural fibrosis, and intrathecal arachnoiditis was not significantly different between the two groups \((p \geq 0.128)\). Among the above radiologic findings, transiting nerve rootlet thickening and signal alteration were the most significant findings in the multivariate analyses \((p \leq 0.009)\).

CONCLUSION: On MRI, PPP was associated with transiting nerve rootlet abnormalities, including thickening, signal alterations, and ill-distinction, but was not associated with epidural fibrosis or intrathecal arachnoiditis. The most relevant findings were the nerve rootlet thickening and signal alteration.

KEYWORDS: Persistent postoperative pain; Failed back surgery; Laminectomy; Spinal nerve roots; Epidural fibrosis
adjacent segment disease, the battered root syndrome, or the nerve root entrapment syndrome (5). Despite postoperative MRI showing successful nerve root decompression at the correct level without recurrent disc herniation, there have been patients who continued to complain of PPP. The possible causes of such PPP include iatrogenic nerve root irritation or injury during operation and nerve root entrapment or adhesion during epidural fibrosis (5).

Lee et al. (6) reported that residual or recurrent postoperative pain was associated with nerve root enhancement, displacement, and thickening but not with epidural fibrosis. However, this study was performed using contrast-enhanced MRI due to the low-magnetic field used (83 patients at 0.5T and 57 patients at 1.5T) and performed only univariate analyses, examining the relationship between each radiologic factor and each symptom, and not a comprehensive multivariate analysis. Furthermore, the American College of Radiology has recommended the use of MRI without contrast for routine evaluations of patients who present with spinal pain and a history of prior lumbar surgery (7). The relationship between epidural fibrosis and PPP after lumbar spine surgery is also controversial; some studies did report to have found a relationship between the two (8-10) but others did not (6, 11). High-magnetic field MRI, defined as having a field strength higher than 1T (12), enables the evaluation of detailed anatomic structures such as the ventral and dorsal transiting nerve rootlet as well as the nerve root. We hypothesized that transiting nerve rootlet abnormalities or epidural fibrosis seen on MRI might be connected to PPP in patients following successful decompression surgery without complications. Thus, we aimed to determine whether changes in the transiting nerve rootlet or its surroundings, as seen on MRI performed after lumbar hemilaminectomy, are associated with PPP.

MATERIALS AND METHODS

This retrospective study was approved by the Institutional Review Board and the requirement for informed consent was waived.

Study Group
We enrolled patients who underwent single-level partial hemilaminectomy of the lumbar spine and MRI one year after the operation, and who had no postoperative complications or other level spinal abnormalities.

Five hundred and three lumbar MRI examinations were identified between January 2010 and December 2018 through a computerized search of radiology reports using the following keywords: report of “laminectomy,” modality of “MRI,” and anatomy of “lumbar spine.” Of the 503 examinations, 238 were of patients who had undergone single-level partial hemilaminectomy of the lumbar spine. Additionally, 150 of these 238 patients underwent follow-up MRI at one year after the operation. Of the 150 patients, 73 patients (mean age, 61 years; age range, 28–87 years; 43 males [mean age, 60 years; range, 28–87 years] and 30 females [mean age, 63 years; range, 30–87 years]) who had no postoperative complications or other level spinal abnormalities were included in the final analysis (Fig. 1).

Clinical Outcome
Information on age, sex, preoperative symptoms, operative date and level, MRI examination date, and spinal symptoms at the time of the MRI examination was obtained from the medical records. A spine surgeon, blinded to the radiologic data, retrospectively evaluated each patient’s medical records for the presence of PPP at the operative level. PPP was defined as no significant symptom relief when compared to the preoperative symptoms. Significant symptom relief was defined as all of the following: 1) no or minimal remaining pain, 2) ability to work not adversely affected, 3) no use of analgesic medications, and 4) patient satisfaction with the procedure (13). Ambiguous low back or buttock pain unrelated to the preoperative symptoms or operative site was not defined as PPP.

MRI Protocol
MRI examinations were performed using 1.5T units (MAGNETOM Vision and Avanto, Siemens Healthineers), in 52 patients, or 3T units (MAGNETOM Skyra and Trio, Siemens Healthineers; Achieva TX, Philips Healthcare), in 21 patients. All MRI examinations included an axial T1-weighted fast spin-echo (FSE) sequence (repetition time ms/echo time ms: 500–680/10–16) and T2-weighted FSE sequence (3000–6710/81–120), with a 4-mm section thickness and a 140 x 140–160 x 160 mm field of view, and a sagittal T1-weighted FSE sequence (358–767/10–15) and T2-weighted FSE sequence (2980–5082/81–134), with a 4-mm section thickness and a 250 x 250–320 x 320 mm field of view.

Image Analysis
Two radiologists (readers 1 and 2 with 15 and 3 years...
Check for Transiting Nerve Rootlet Abnormalities on MRI

Transiting Nerve Rootlet Abnormalities

The transiting nerve rootlet abnormalities were evaluated by looking at the ventral and dorsal nerve rootlets between the dural sac and the foraminal zone (Fig. 2). The transiting nerve rootlet abnormalities were divided into four subcategories: thickening, signal alteration, distinction, and displacement or compression (6, 14) (Fig. 2). Transiting nerve rootlet thickening was defined as a more than 50% increase in the diameter of the nerve rootlet at the operative site when compared with the contralateral nerve rootlet at the same level (15). A presence of a nerve rootlet signal alteration was defined as hyperintensity of the nerve rootlet when compared with the contralateral nerve rootlet at the same level on the T2-weighted sequence. Nerve rootlet distinction was defined as follows: 1) well-distinction if the margin of the nerve rootlet was well-circumscribed, sharply bordered, and well-distinct from the surrounding cerebrospinal fluid; 2) partially ill-distinction if part of the nerve rootlet margin was indistinct; and 3) completely ill-distinction if the whole nerve rootlet margin was indistinct and indistinguishable from the surrounding structures. Nerve rootlet displacement or compression was defined as present if the position of the nerve rootlet was different from what was expected or compressed when a more than 50% decrease in diameter was observed when compared with the position of the contralateral nerve rootlet at the same level (6).

Epidural Fibrosis

Epidural fibrosis was diagnosed when there was an area of intermediate or low signal intensity on a T1-weighted sequence in the epidural space that obscured or obliterated the normal high signal of the epidural fat. The amount of epidural fibrosis was graded on a scale of 1–4 for each quadrant of each imaging slice encompassing the operative level, as follows: 1 was assigned if the quadrant area showed no or minimal fibrosis, i.e., < 25% of the quadrant was filled with fibrosis, 2 if it showed ≥ 25% and < 50% fibrosis, 3 if ≥ 50% and < 75%, and 4 if ≥ 75% (8-10).
Intrathecal Arachnoiditis

Intrathecal arachnoiditis was defined as present if multiple nerve roots were clumped together or adhered to the margins of the thecal sac (16, 17). This study evaluated the intrathecal nerve roots, excluding the transiting nerve rootlets or dorsal root ganglia.

Statistical Analysis

For the analysis, the patients were divided into two groups, i.e., patients with and without PPP. The clinical information was compared between the two groups using Fisher’s exact test or the t test. Univariate and multivariate stepwise logistic regression models were used to evaluate the associations between the radiological findings and PPP. For the analysis, nerve rootlet distinction and epidural fibrosis were each classified into two categories because the numbers were too small for several subcategories (“normal” vs. “abnormal” defined as “well-defined” vs. “partially + completely ill-defined” for nerve rootlet distinction; “grade 1” vs. “grade 2 + 3 + 4” for epidural fibrosis). Inter-reader agreement was assessed using kappa (κ) statistics for each radiologic finding with the following scale: 0–0.20, poor; 0.21–0.40, fair; 0.41–0.60, moderate; 0.61–0.80, good; and 0.81–1.00, excellent (18).

The analyses were performed using the SPSS for Windows software (version 21.0, IBM Corp.) and differences with p values < 0.05 were considered statistically significant.

### Table 1 Patient Demographics

<table>
<thead>
<tr>
<th>Category</th>
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<th>P</th>
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</thead>
<tbody>
<tr>
<td>Mean age (CI), years</td>
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<td>21</td>
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<tr>
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<td>11</td>
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<td>Operative level*</td>
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<tr>
<td>L3–4</td>
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<td>1</td>
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<tr>
<td>L4–5</td>
<td>16</td>
<td>13</td>
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<tr>
<td>L5–S1</td>
<td>22</td>
<td>18</td>
<td></td>
</tr>
<tr>
<td>Mean time interval between operation and MRI (CI), month</td>
<td>19 (13, 26)</td>
<td>20 (14, 26)</td>
<td>0.367</td>
</tr>
<tr>
<td>Total number of patients</td>
<td>41</td>
<td>32</td>
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</tr>
</tbody>
</table>

*The analysis of the operation level was only a comparison between L4–5 and L5–S1, excluding L3–4 due to small sample size. 95% CI = 95% confidence interval, PPP = persistent postoperative pain

### RESULTS

Among the 73 patients, 41 had no PPP and 32 had PPP. The clinical information, including the patient age, sex, operative level, and mean time interval between the operation and the MRI, was not significantly different between the patients with and those without PPP (p ≥ 0.345) (Table 1). The preoperative diagnosis was either herniated disc (n = 56) or spinal stenosis (n = 17). Most patients underwent follow-up MRI for symptoms such as radiculopathy, leg weakness, or back or buttock pain (94.5%,
Check for Transiting Nerve Rootlet Abnormalities on MRI

n = 69) and four patients underwent MRI to screen for possible postoperative complications.

The results of the associations between the MRI findings and PPP are summarized in Table 2. Among the radiologic findings, the presence of nerve rootlet thickening, signal alteration, and ill-distinction were significantly different between the patients with and those without PPP for both readers \((p \leq 0.020)\) (Fig. 3). Nerve rootlet thickening was more common in the PPP group than in the non-PPP group \(\{\text{non-PPP vs. PPP: 16/41 (39.0%) vs. 24/32 (75.0%) for reader 1 \((p = 0.003; \text{odds ratio [95% confidence interval, 95% CI], 4.687 [1.696–12.959]}\) and 13/41 (31.7%) vs. 19/32 (59.4%) for reader 2 \((p = 0.020; \text{odds ratio [95% CI], 3.148 [1.200–8.259]}}\)\}). The presence of signal alteration was significantly different between the groups \(\{\text{non-PPP vs. PPP: 19/41 (46.3%) vs. 24/32 (75.0%) for reader 1 \((p = 0.016; \text{odds ratio [95% CI], 3.474 [1.267–9.523]}\) and 14/41 (34.1%) vs. 21/32 (65.6%) for reader 2 \((p = 0.009; \text{odds ratio [95% CI], 3.682 [1.390–9.752]}}\)\}). The frequency of “ill-distinction” was higher than that of “well-distinction” in the PPP group \(\{\text{non-PPP vs. PPP: 13/41 (31.7%) vs. 20/32 (62.5%) for reader 1 \((p = 0.010; \text{odds ratio [95% CI], 3.590 [1.358–9.489]}\) and 14/41 (34.1%) vs. 20/32 (62.5%) for reader 2 \((p = 0.018; \text{odds ratio [95% CI], 3.214 [1.226–8.426]}}\)\}). Conversely, the presence of nerve rootlet displacement or compression \((p = 0.128 \text{ for reader 1 and 0.999 for reader 2})\), epidural fibrosis \((p = 0.202 \text{ for reader 1 and 0.890 for reader 2})\), and intrathecal arachnoiditis \((p = 0.964 \text{ for reader 1 and 0.564 for reader 2})\) was not significantly different between the two groups for both readers. The most common epidural fibrosis was grade 1. Inter-reader agreement was moderate-to-good for nerve rootlet thickening, signal alteration, distinction, displacement or compression, and epidural fibrosis \((\kappa = 0.483–0.685)\) and fair for arachnoiditis \((\kappa = 0.316)\). In the multivariate regression analysis, the transiting nerve rootlet thickening \((p = 0.002; \text{odds ratio [95% CI], 4.687 [1.696–9.489]}\) and signal alteration \((p = 0.009; \text{odds ratio [95% CI], 3.682 [1.390–9.752]}}\).

### Table 2. Association between Radiologic Findings and PPP

<table>
<thead>
<tr>
<th>MRI Findings</th>
<th>Reader 1</th>
<th></th>
<th>Reader 2</th>
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<tr>
<td></td>
<td>No PPP</td>
<td>PPP</td>
<td>P</td>
<td>Odds Ratio</td>
<td>No PPP</td>
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<tr>
<td>NR abnormalities</td>
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<tr>
<td>Thickening</td>
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<tr>
<td>Signal alteration</td>
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<td></td>
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</tr>
<tr>
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<td>22</td>
<td>8</td>
<td>0.016</td>
<td>3.474</td>
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<tr>
<td>Present</td>
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<td>24</td>
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<td>Distinction*</td>
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<td>Well</td>
<td>28</td>
<td>12</td>
<td>0.010</td>
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<td>Partially ill</td>
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<td>Epidural fibrosis*</td>
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<td>Grade 1</td>
<td>29</td>
<td>18</td>
<td>0.202</td>
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<td>Grade 2</td>
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<tr>
<td>Grade 3</td>
<td>4</td>
<td>5</td>
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<tr>
<td>Grade 4</td>
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<td>Arachnoiditis</td>
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<td>22</td>
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Data are number of patients. P values and odds ratios were calculated with logistic regression analysis. *For analysis, nerve rootlet distinction and epidural fibrosis were each classified into two categories (“normal” vs. “abnormal”: “well-defined” vs. “partially + completely ill-defined” for distinction; “grade 1” vs. “grade 2 + 3 + 4” for epidural fibrosis). NA = not applicable, NR = transiting nerve rootlet.
CI), 3.682 [1.390–9.752]) were the findings most strongly associated with PPP.

**DISCUSSION**

PPP is an adverse surgical outcome of lumbar disc surgery that may result from many different possible mechanisms and, therefore, present with many different radiologic findings. Patients often complain of persistent back pain or sciatica, regardless of the presence or absence of neural compression. Our study suggests that the thickening, signal alteration, and ill-distinction of the transiting nerve rootlet, observed on MRI, are significantly associated with PPP. Among the radiologic findings, the nerve rootlet thickening and signal alteration were the findings associated the most with PPP. However, we found no association between epidural fibrosis or intrathecal arachnoiditis and PPP.

To our knowledge, this approach, of evaluating the association between the characteristics of the transiting nerve rootlet, but not the nerve root, and PPP, is a novel one. One study investigated the association between nerve root abnormalities and PPP, showing an association between nerve root enhancement, thickening, and displacement and PPP (6). However, that study investigated the correlation between the symptoms and nerve root changes shown on contrast-enhanced T1-weighted images, due to the low magnetic field used. Nevertheless, recently developed high-field MR that reveals detailed anatomical features can be used to evaluate the size, signal intensity, and margin of nerve rootlets. Furthermore, the American College of Radiology has recommended MRI without contrast for routine evaluations of patients who present with spinal pain and a history of prior lumbar surgery (7). Although we included several patients who underwent contrast-enhanced MRI, we only evaluated the non-contrast-enhanced images for the purposes of this study. The multivariate analyses in our study showed that transiting nerve rootlet thickening and signal alteration were the most relevant factors associated with PPP. We suggest naming these findings the “nerve rootlet swelling sign.” We speculate that despite the successful decompression of the nerve root, the nerve rootlet abnormalities noted on MRI might be due to intraoperative dural sleeve tears, nerve rootlet irritation, or the sequelae of previous nerve compression and that these might be associated with PPP.

Epidural fibrosis following lumbar surgery has been reported to occur in 10% to 75% of cases (8, 13) and PPP was associated with 10% to 24% of these epidural fibrosis cases (13). Although the exact mechanism of fibrosis formation has not been established, fibrosis-induced pain can be caused by the inflammation, compression, and traction of the fibrotic tissue on adjacent nerve roots (19, 20). Our results show that neither epidural fibrosis nor nerve rootlet displacement or compression was associated with PPP. Some authors have described epidural fibrosis as a cause of PPP (8-10) but other authors have opposed...
this opinion (6, 11). The contrasting conclusions in these studies may have resulted from their different definitions of epidural fibrosis, depending on the extent of epidural fibrotic tissue or compression of the thecal sac. Considering our results, transiting nerve rootlet swelling appears to be associated with PPP regardless of the existence of epidural fibrosis or nerve rootlet displacement. Therefore, it is important to focus on the transiting nerve rootlet itself in axial T2-weighted sequence images when evaluating the MRI images of patients with PPP.

Arachnoiditis is a form of leptomeningeal and neural inflammation within the spinal canal. Arachnoiditis may be caused by varying triggers, including intradural surgery or dural tears, meningitis, subarachnoid hemorrhage, or injections; however, in many cases of arachnoiditis, the cause remains idiopathic (16, 21). Sterile arachnoiditis has been reported to cause persistent pain in 6–16% of postoperative cases (22). The inter-reader agreement in our study was fair for the evaluation of arachnoiditis and moderate-to-good for other radiologic factors. The relatively low inter-reader agreement in our study occurred because the evaluation of adhering nerve roots was ambiguous in some mild cases. However, in both of the readers’ assessments, the presence of intrathecal arachnoiditis was not associated with PPP.

Our study has several limitations. First, we did not analyze the preoperative MRI findings. Comparing pre- and post-operative MRI findings could result in a more accurate radiologic evaluation. Second, although a spine surgeon independently determined the presence of PPP, quantitative pain scores such as the Oswestry Disability Index or a functional score were not included in the analysis. Third, the asymptomatic patients were much less likely to perform MRI than the patients with PPP. There is, therefore, a possibility of selection bias. Fourth, although we evaluated the nerve rootlet changes and epidural fibrosis with reference to the contralateral side, some cases were ambiguous to judge. If an MR neurography would have been added, evaluation could have been more accurate (23).

In conclusion PPP was associated with transiting nerve rootlet abnormalities, including thickening, signal alterations, and ill-distinction, but not with epidural fibrosis or intrathecal arachnoiditis. The most relevant findings were the nerve rootlet thickening and signal alteration. If pain persists in patients after lumbar spine surgery, it is recommended to check for transiting nerve rootlet abnormalities on MRI.

Conflicts of Interest
The authors have no potential conflicts of interest to disclose.

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