Research Article

Numbness and Weakness Recovered at a Less Extent in Patients with Lumbar Disc Herniation after Percutaneous Endoscopic Lumbar Discectomy

Yuming Wang, Fuqiang Gao, and Haibo Zou

Department of Orthopedics, China-Japan Friendship Hospital, 100029 Beijing, China

Correspondence should be addressed to Fuqiang Gao; gaofuqiang@bjmu.edu.cn and Haibo Zou; drzouhaibo@163.com

Received 14 June 2019; Revised 23 September 2019; Accepted 8 October 2019

Guest Editor: Hai-Qiang Wang

Copyright © 2019 Yuming Wang et al. This is an open access article distributed under the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Background. Patients with lumbar disc herniation (LDH) may present with motor disorders and various sensory disorders, among which pain and numbness are the most common ones. Percutaneous endoscopic lumbar discectomy (PELD) is reported to be both safe and effective. However, most of the previous studies focused on the recovery of pain, and the relief extent of numbness and weakness has rarely been reported. The Sciatica Bohorsomeness Index (SBI) is a self-assessment tool for LDH patients. It has demonstrated acceptable reliability, construct validity, and responsiveness. Objectives. Our aim was to explore the curative effect of percutaneous endoscopic lumbar discectomy and to compare the various extent of relief among pain, numbness, and weakness.

Methods. The medical records of patients admitted for LDH from September 2016 to December 2018 were collected, and the patients were followed up for 3 months to evaluate the relief of their clinical symptoms. Preoperative and postoperative total scores and subitem scores of SBI were compared to evaluate the relief of pain, numbness, and weakness. Surgical outcomes of PELD were evaluated by the Nakai score, and patients were divided into two groups accordingly, which were the relief group (excellent and good in the Nakai score) and the less relief group (fair and poor in the Nakai score). Risk factors for PELD outcomes and preoperative presence of numbness and/or weakness were analyzed by the logistic model, and $p$ value less than 0.05 was considered significant.

Results. A total of 86 patients met the inclusion criteria and acquired 3 months follow-up. Relief extent of pain, numbness, and weakness, was 82%, 41%, and 21%, respectively. There were 71 cases in the relief group and 15 cases in the less relief group. Results of the logistic regression analysis showed that the preoperative pain score of SBI ($p = 0.002; OR: 1.647 (1.199–2.261)$) was a relatively independent risk factor for PELD outcomes, and multiplicity of duration of preoperative symptoms and imaging grade ($p = 0.004; OR: 1.015 (1.005–1.026)$) was a relatively independent risk factor for preoperative presence of numbness and/or weakness. Conclusions. PELD had a good curative effect in the treatment of LDH. Patients of LDH recovered best from pain, followed by numbness and weakness after PELD. Higher level of patients self-reported preoperative pain indicated a better surgical outcome for LDH patients, and preoperative long duration of symptoms together with a severe compression of nerve root significantly increased the risk of presenting numbness and/or weakness.

1. Introduction

Patients with lumbar disc herniation (LDH) usually present with symptoms like pain, numbness, and weakness, which have a negative influence on their social functions [1, 2]. Percutaneous endoscopic lumbar discectomy (PELD) first appeared in 1992 [3] and has been widely used in clinical practice for lumbar spine diseases due to less invasiveness and faster recovery compared with traditional surgery [4–6]. Many studies have reported that PELD could achieve comparative therapeutic effect as traditional surgery. However, most previous studies applied the Japanese Orthopaedic Association (JOA) score, the Visual Analog Scale for Pain (VAS Pain), the Oswestry Disability Index (ODI) score, or the 36-Item Short-Form Health Survey score (SF36) to evaluate surgery outcomes, which mainly provided evidence about patients’ recovery of either overall function of neurodeficit, pain, social dysfunction, or quality of life.
Pain Research and Management

[7–9], while variable relief extent of symptoms such as numbness and weakness has rarely been reported [10].

Sciatica Bothersomeness Index (SBI) is a self-assessment tool to assess the severity of sciatica. It contains four items, which can evaluate the main symptoms of LDH patients quantitatively, such as pain, numbness, and weakness [11]. It has been applied in previous studies about lumbar disc herniation and has demonstrated acceptable reliability, construct validity, and responsiveness [12, 13]. The purpose of this study was to investigate the curative effect of endoscopic treatment of lumbar disc herniation. The SBI score was used both preoperatively and postoperatively to evaluate the variable extent of relief among pain, numbness, and weakness. We hypothesized that numbness and weakness recovered at a less extent than pain in patients with lumbar disc herniation after percutaneous endoscopic lumbar discectomy.

2. Methods

2.1. Participants. LDH patients admitted for PELD from September 2016 to December 2018 were enrolled in this study. The criteria for inclusion were as follows: (1) complained of unilateral lower limb radiating pain, numbness, weakness, or other symptoms caused by single nerve root compression; (2) with symptoms consistent with preoperative imaging; and (3) with no significant relief of symptoms after 6 weeks of regular conservative treatment. The exclusion criteria were as follows: (1) with lumbar spinal stenosis; (2) and/or with lumbar instability; and (3) with spondylolisthesis or deformity.

2.2. Surgical Procedure. Two approaches for PELD were selected according to disk segments of herniation, which were transfemoral approaches for L45 and interlaminar approaches for L5S1. Surgical procedure was performed by one doctor under local anesthesia with 1% lidocaine, and the patients were laid in the prone position. A cannula was inserted into the spinal canal under the guidance of the C-arm X-ray machine. Free disk fragments were removed as much as possible. Microscopically, free movement of the nerve root and dural sac with the change of abdominal cavity pressure during breathing indicated that the goal of nerve decompression had been achieved.

2.3. Postoperative Management. All patients were treated with neurotrophic drugs routinely after operation and were able to get out of bed with a brace 24 hours after operation. After 3 weeks postoperation, the patients were instructed to return to everyday activities gradually.

2.4. Outcome Measurement. (1) Demographic data included gender, age, and body mass index (BMI); (2) clinical data included duration of preoperative symptoms, the SBI scores for evaluating preoperative symptoms, and the ODI score for evaluating social dysfunction. The SBI score is a patient self-rated instrument usually applied to evaluate the severity of sciatica, which contains 4 items and includes the most common symptoms of LDH patients, such as pain, numbness, and weakness. Each item has a range of score from 0 to 6. The index has labels at the categories 0 (not bothersome), 3 (somewhat bothersome), and 6 (extremely bothersome), which provides a total score from 0 to 24 when summing up the ratings across the 4 items [11]. (3) Imaging data were graded by the Pfirrmann grading system in which nerve root compression was graded into three categories based on preoperative MR images (A: normal or contact; B: deviation; and C: compression) [14]. Evaluation of the image data was completed independently by two attending doctors who had been specializing in spine surgery for at least five years. Any discordance in the evaluation of the image results was discussed, and final agreement was reached before they were recorded for analysis. (4) Follow-up data: all patients were followed up at 3 months after operation. The relief of symptoms was evaluated by the change of the total SBI score and each subitem score. The extent of symptom relief was calculated by the formula (preoperative scores−postoperative scores)/preoperative scores. The curative effect of PELD was evaluated by the Nakai score (Table 1) [15].

2.5. Statistical Analysis. All data were analyzed using the Statistical Package for the Social Sciences (SPSS) version 22 (IBM, USA). For all qualitative variables, the total number of patients and the percentage were provided. For quantitative variables, the mean and standard deviation were provided. To reveal the relationship between social dysfunction and different symptoms like pain, numbness, and weakness, correlations between preoperative ODI and the SBI score and its subitem scores were analyzed by the Pearson correlation analysis. The risk factors for curative effect and preoperative presence of numbness and/or weakness were analyzed by the logistic regression model. Factors according to its clinical significance were first assessed individually as predictive variables by the logistic regression analysis, and the final model included all the predictive variables with p values less than 0.2 as covariates, which were further assessed together by the logistic regression. p value less than 0.05 was taken as the criterion for covariates being significant risk factors.

3. Results

A total of 87 patients were enrolled in this study, and one case was lost to follow-up. The 86 enrolled patients included 44 males and 42 females, aged from 21–80, with an average age of 49. Demographic and clinical characteristics of the study patients are listed in Table 2.

Proportions of pain, numbness, and weakness are shown in Figure 1. Pain was the most common symptom, followed by numbness and weakness. Before surgery, 75 (87.2%) out of 86 patients had pain, 61 patients (70.9%) had numbness, and only 37 patients (43%) had weakness. Only 23 patients (26.7%) had all these 3 symptoms at baseline.

Results of the Pearson correlation analysis showed that the ODI score displayed a positive correlation with the
Table 1: Nakai score.

| Scoring used in the study                                      | Excellent | Good | Fair | Poor |
|---------------------------------------------------------------|-----------|------|------|------|
| The patient has resumed work-related and other activities     |           |      |      |      |
| with slight or no symptoms                                    |           |      |      |      |
| The patient has resumed work-related and other activities     |           |      |      |      |
| but occasionally feels pain in the back or lower limbs        |           |      |      |      |
| The patient has reduced work-related and other activities     |           |      |      |      |
| because of residual pain in the back or lower limbs           |           |      |      |      |
| The patient cannot work or carry out activities of daily     |           |      |      |      |
| living and is considered to be disabled                        |           |      |      |      |

Table 2: Demographic, clinical, and imaging characteristics of enrolled patients.

| Demographic data                                      |        |
|-------------------------------------------------------|--------|
| Age (years)                                           | 49 (21–80) |
| Gender (n, %)                                         |        |
| Male                                                  | 44 (51%) |
| Female                                                | 42 (49%) |
| BMI (kg/m²)                                           | 25.2 (3.80) |
| Clinical data                                         |        |
| Duration of symptoms (weeks)                          | 118 (1–1440) |
| Preop ODI                                             | 49 (22) |
| Preop SBI total                                       | 9.4 (4.35) |
| Preop SBI pain                                        | 3.7 (1.74) |
| Preop SBI numbness                                    | 2.9 (2.08) |
| Preop SBI weakness                                    | 0.7 (1.11) |
| Imaging grade                                         |        |
| A                                                     | 32 (37.2%) |
| B                                                     | 19 (22%) |
| C                                                     | 35 (40.6%) |

Figure 1: Proportions of various symptoms.

preoperative SBI total score, pain score, and weakness score, while no significant correlations were found between the ODI score and the numbness score (Table 3).

At 3 months after operation, the mean ODI score of the study population was decreased to 16.2 (11.33), p < 0.001. The mean SBI total score was decreased to 3.2 (2.32), p < 0.001, with a symptom relief rate of 65%. The mean pain score was decreased to 0.6 (0.82), p < 0.001, with a symptom relief rate of 82%. The mean numbness score was decreased to 1.6 (1.34), p < 0.001, with a symptom relief rate of 41%. The mean weakness score was decreased to 0.5 (0.85), p < 0.001, with a symptom relief rate of 21% (Figure 2).

According to the Nakai score, 11 (12.7%) cases had achieved excellent surgical outcomes, while good, fair, and poor were 60 (69.8%), 12 (14.0%), and 3 (3.5%), respectively. Patients were divided into two groups based on the Nakai score, which were the relief group with excellent or good in the Nakai score and the less relief group with fair or poor in the Nakai score. There were 71 cases in the relief group and 15 cases in the less relief group.

Demographics, preoperative clinical data, and imaging grade stratified by surgical outcomes are shown in Table 4. All the predictive variables were first assessed individually, and the significance levels are also listed in Table 4. Gender (p = 0.193), BMI (p = 0.184), preoperative SBI pain score (p = 0.002), preoperative SBI numbness score (p = 0.08), preoperative SBI weakness score (p = 0.186), preoperative SBI total score (p = 0.05), and preoperative ODI score (p = 0.007) were included as covariates in the final model of surgical outcomes. Among them, preoperative SBI pain score (odds ratio [OR] = 1.647; 95% confidence interval [CI], 1.199–2.261; p = 0.002) significantly increased the probability of relatively good surgical outcomes, while gender (p = 0.242), BMI (p = 0.175), preoperative SBI numbness score (p = 0.302), preoperative SBI weakness score (p = 0.126), preoperative SBI total score (p = 0.525), and preoperative ODI score (p = 0.059) were found to be nonsignificant predictors of the relatively good surgical outcomes and were excluded from the model.

To further explore the risk factors for preoperative presence of numbness and/or weakness, we divided the patients into the numbness and/or weakness group and the pain alone group based on the SBI subitem scores. Patients with SBI numbness and/or weakness scores greater than 2 points were grouped into the numbness and/or weakness group while the rest were in the pain alone group. Demographics, preoperative ODI score, duration of preoperative symptoms, and imaging grade stratified by presence of numbness and/or weakness are shown in Table 5. All the above predictive variables were first assessed individually, and the significance levels are also listed in Table 5. The reason for analyzing the multiplicativity of duration of preoperative symptoms and imaging grade was that they had complex interactions between them which might have influence on each other. Duration of preoperative symptoms (p = 0.006), imaging grade (p = 0.123), and joint effect of duration of preoperative symptoms and imaging...
grade (multiplicativity) \((p = 0.004)\) were included as covariates in the final model of presence of numbness and/or weakness. Among them, joint effect of duration of preoperative symptoms and imaging grade (odds ratio \([OR] = 1.015\); 95% confidence interval [CI], 1.005–1.026; \(p = 0.004\)) significantly increased the probability of presence of numbness and/or weakness, while duration of preoperative symptoms \((p = 0.633)\) and imaging grade \((p = 0.341)\) individually were found to be nonsignificant predictors and were excluded from the model.

### 4. Discussion

Patients with lumbar disc herniation usually present symptoms like pain, numbness, and weakness, which have a negative influence on their social functions [1, 16]. Our results showed that 75 (87.2%) out of 86 patients had pain, 61 patients (70.9%) had numbness, and only 37 patients (43%) had weakness. Preoperatively, pain was the most common symptom, followed by numbness and weakness, which was consistent with the results of the previous literature [10].
Huang and Sengupta conducted a retrospective study in which they followed up 85 patients who had surgical decompression of the nerve root due to lumbar disease and found that most of the patients had pain and numbness before surgery. Our results also showed that the preoperative pain score and weakness score correlated with the ODI score, while numbness did not, which indicated that although commonly presented in LDH patients, numbness did not have so much negative influence on their social function as pain and weakness did. This was consistent with our previous result that pain relieved by the largest extent, while numbness recovered at a slower pace [10]. By analyzing the risk factors for pain and weakness before surgery, we found that preoperative numbness and/or weakness before surgery is still meaningful in clinical application because persistent numbness and/or weakness after satisfactory pain relief is more difficult to recovery, resulting in a worse surgical outcome. However, given the fact that most LDH patients presented pain simultaneously with other symptoms, it was reasonable that their effects on the surgical outcomes were confounded by pain. While numbness and weakness were not independent risk factors, we believed the results were still meaningful in clinical application because persistent numbness and/or weakness after satisfactory pain relief is commonly seen in patients during the postoperative follow-up [10].

The SBI scores have been used in the previous studies about lumbar disc herniation and have demonstrated acceptable reliability, construct validity, and responsiveness [11, 13]. Three of them were also incorporated in the North American Spine Society questionnaire [12]. Our results showed that the preoperative SBI pain score significantly increased the probability of relatively good surgical outcomes, which indicated that pain was the most significant outcome criteria for patients during the early stage after surgery. Patients with more preoperative pain tended to have more relief of symptoms postoperatively. This was consistent with our common clinical observation that pain relief contributed more to the overall symptoms relief than other symptoms like numbness and weakness, which was also consistent with our previous results that pain relieved by the largest extent. Theoretically, as caused mainly by larger myelinated nerve fibers, numbness and weakness tended to be more difficult to recovery, resulting in a worse surgical outcome. However, given the fact that most LDH patients presented pain simultaneously with other symptoms, it was reasonable that their effects on the surgical outcomes were confounded by pain. While numbness and weakness were not independent risk factors, we believed the results were still meaningful in clinical application because persistent numbness and/or weakness after satisfactory pain relief is commonly seen in patients during the postoperative follow-up [10].

For qualitative variables, the total number of patients and the percentage are provided. For quantitative variables, the mean and standard deviation are provided. These predictive variables are first assessed individually; the significance levels are also listed. *Indicates predictive variables included in the final model.

### Table 5: Demographics, preoperative ODI score, duration of preoperative symptoms, and imaging grade stratified by presence of numbness and/or weakness.

| Variables                                      | Numbness and/or weakness group (57) | Pain alone group (29) | p value |
|-----------------------------------------------|-------------------------------------|-----------------------|---------|
| Age, mean (SD)                                | 50 (16.0)                           | 48 (17.9)             | 0.486   |
| Male, n (%)                                   | 31 (54.3)                           | 13 (44.8)             | 0.403   |
| BMI, mean (SD)                                | 25.3 (4.08)                         | 24.9 (3.24)           | 0.690   |
| Duration of symptoms, mean (SD)               | 171 (290.3)                         | 16 (16.0)             | 0.006*  |
| Preop ODI, mean (SD)                          | 51.3 (23.0)                         | 45.6 (20.0)           | 0.266   |
| Imaging grade, n (%)                          | 18 (31.5%)                          | 14 (48.2%)            | 0.123#  |
| Joint effect of duration of symptoms and imaging grade | —                                   | —                     | 0.004*  |

For qualitative variables, the total number of patients and the percentage are provided. For quantitative variables, the mean and standard deviation are provided. These predictive variables are first assessed individually; the significance levels are also listed. *Indicates predictive variables included in the final model.
nerve root compression may induce deformation and demyelination of nerve fibers distally resulting in numbness and weakness.

This study has some limitations. The reason for following up patients at 3 months after operation was that most symptoms of LDH recovered the fastest at the first 6 weeks and plateaued at 3 months postoperatively according to the previous literature [10]. The results of our study might uncover the extent of symptoms relief at the early stage after operation. Yet, there was still slow improvement of these symptoms even until 1 year postoperatively. Further studies might be needed with a longer follow-up to reveal the overall relief extent of various symptoms. And the other limitation is that this study is a retrospective analysis of prospectively collected data, which may have potential biases of a retrospective study.

5. Conclusion

In patients with LDH, pain was the most common symptom, followed by numbness and weakness. PELD had a good curative effect in the treatment of LDH. Patients recovered best from pain, followed by numbness and weakness. Higher level of patients’ self-reported preoperative pain indicated a better surgical outcome for LDH patients, and preoperative long duration of symptoms together with a severe compression of nerve root significantly increased the risk of presenting numbness and/or weakness.

Data Availability

The data used to support the findings of this study are available from the corresponding authors upon request.

Conflicts of Interest

The authors declare that they have no conflicts of interest.

Acknowledgments

The authors would like to acknowledge all the other doctors and nurses of the Department of Orthopedic of China-Japan Friendship Hospital who gave constructive suggestions in writing assistance.

References

[1] O. H. Jensen, “The level-diagnosis of a lower lumbar disc herniation: the value of sensibility and motor testing,” Clinical Rheumatology, vol. 6, no. 4, pp. 564–569, 1987.
[2] I. Caby, N. Olivier, F. Mendelek, R. B. Kheir, J. Vanvelzenaher, and P. Pelayo, “Functional restoration of the spine: effect of initial pain level on the performance of subjects with chronic low back pain,” Pain Research and Management, vol. 19, no. 5, pp. e133–e138, 2014.
[3] P. Kambin, “Arthroscopic microdiscectomy,” Arthroscopy: The Journal of Arthroscopic & Related Surgery, vol. 8, no. 3, pp. 287–295, 1992.
[4] L. Pan, P. Zhang, and Q. Yin, “Comparison of tissue damages caused by endoscopic lumbar discectomy and traditional lumbar discectomy: a randomised controlled trial,” International Journal of Surgery, vol. 12, no. 5, pp. 534–537, 2014.
[5] F. U. Hermantin, T. Peters, L. Quartararo, and P. Kambin, “A prospective, randomized study comparing the results of open discectomy with those of video-assisted arthroscopic microdiscectomy,” The Journal of Bone & Joint Surgery, vol. 81, no. 7, pp. 958–965, 1999.
[6] B. Wu, G. Zhan, X. Tian et al., “Comparison of transfuraminal percutaneous endoscopic lumbar discectomy with and without foraminoplasty for lumbar disc herniation: a 2-year follow-up,” Pain Research and Management, vol. 2019, Article ID 6924941, 12 pages, 2019.
[7] Y.-M. Ryang, M. F. Oertel, L. Mayfrank, J. M. Gilibach, and V. Rohde, “Standard open microdiscectomy versus minimal access trocar microdiscectomy: results of a prospective randomized study,” Neurosurgery, vol. 62, no. 1, pp. 174–182, 2008.
[8] Y. Katayama, Y. Matsuyma, H. Yoshisahara et al., “Comparison of surgical outcomes between macro discectomy and micro discectomy for lumbar disc herniation: a prospective randomized study with surgery performed by the same spine surgeon,” Journal of Spinal Disorders & Techniques, vol. 19, no. 5, pp. 344–347, 2006.
[9] B. J. Moon, S. Yi, Y. Ha, K. N. Kim, D. H. Yoon, and D. A. Shin, “Clinical efficacy and safety of trans-sacral epiduroscopic laser decompression compared to percutaneous epidural neurolasty,” Pain Research and Management, vol. 2019, Article ID 5293460, 7 pages, 2019.
[10] P. Huang and D. K. Sengupta, “How fast pain, numbness, and paresthesia resolves after lumbar nerve root decompression: a retrospective study of patient’s self-reported computerized pain drawing,” Spine, vol. 39, pp. E529–E536, 2014.
[11] D. L. Patrick, R. A. Deyo, S. J. Atlas, D. E. Singer, A. Chapin, and R. B. Keller, “Assessing health-related quality of life in patients with sciatica,” Spine, vol. 20, no. 17, pp. 1899–1908, 1995.
[12] E. Aghaye, A. Elfering, C. Schizas, and A. F. Mannion, “Factor analysis of the North American Spine Society outcome assessment instrument: a study based on a spine registry of patients treated with lumbar and cervical disc arthroplasty,” The Spine Journal, vol. 14, no. 6, pp. 916–924, 2014.
[13] J. A. Rihn, M. Kurd, A. S. Hilibrand et al., “The influence of obesity on the outcome of treatment of lumbar disc herniation: analysis of the spine patient outcomes research trial (SPORT),” The Journal of Bone and Joint Surgery-American Volume, vol. 95, no. 1, pp. 1–8, 2013.
[14] C. W. A. Pfirrmann, C. Dora, M. R. Schmid, M. Zanetti, J. Hodler, and N. Boos, “MR image-based grading of lumbar nerve root compromise due to disk herniation: reliability study with surgical correlation,” Radiology, vol. 230, no. 2, pp. 583–588, 2004.
[15] O. Nakai, A. Oookawa, and I. Yamura, “Long-term roentgenographic and functional changes in patients who were treated with wide fenestration for central lumbar stenosis,” The Journal of Bone & Joint Surgery, vol. 73, no. 8, pp. 1184–1191, 1991.
[16] S. Orita, M. Yamashita, Y. Eguchi et al., “Pregabalin for refractory radicular leg pain due to lumbar spinal stenosis: a preliminary prospective study,” Pain Research and Management, vol. 2016, Article ID 5079675, 10 pages, 2016.
[17] O. Righesso, A. Falavigna, and O. Avanzi, “Comparison of open discectomy with microendoscopic discectomy in lumbar disc herniations: results of a randomized controlled trial,” Neurosurgery, vol. 61, no. 3, pp. 545–549, 2007.
[18] J. I. Brox, O. Reikerås, Y. Nygaard et al., "Lumbar instrumented fusion compared with cognitive intervention and exercises in patients with chronic back pain after previous surgery for disc herniation: a prospective randomized controlled study," *Pain*, vol. 122, no. 1, pp. 145–155, 2006.

[19] J. N. Weinstein, J. D. Lurie, T. D. Tosteson et al., "Surgical versus nonsurgical treatment for lumbar degenerative spondylolisthesis," *New England Journal of Medicine*, vol. 356, no. 22, pp. 2257–2270, 2007.

[20] J. Fairbank, H. Frost, J. Wilson-MacDonald, L.-M. Yu, K. Barker, and R. Collins, "Randomised controlled trial to compare surgical stabilisation of the lumbar spine with an intensive rehabilitation programme for patients with chronic low back pain: the MRC spine stabilisation trial," *BMJ*, vol. 330, no. 7502, p. 1233, 2005.

[21] G. M. Manzano, L. M. P. Giuliano, and J. A. M. Nóbrega, "A brief historical note on the classification of nerve fibers," *Arquivos de Neuro-Psiquiatria*, vol. 66, no. 1, pp. 117–119, 2008.

[22] Y. Fukuda, Y. Li, and R. A. Segal, "A mechanistic understanding of axon degeneration in chemotherapy-induced peripheral neuropathy," *Frontiers in Neuroscience*, vol. 11, 2017.

[23] S. L. Wolf, D. L. Barton, R. Qin et al., "The relationship between numbness, tingling, and shooting/burning pain in patients with chemotherapy-induced peripheral neuropathy (CIPN) as measured by the EORTC QLQ-CIPN20 instrument, N06CA," *Supportive Care in Cancer*, vol. 20, no. 3, pp. 625–632, 2012.

[24] Y. Iizuka, H. Iizuka, S. Tsutsumi et al., "Foot drop due to lumbar degenerative conditions: mechanism and prognostic factors in herniated nucleus pulposus and lumbar spinal stenosis," *Journal of Neurosurgery: Spine*, vol. 10, no. 3, pp. 260–264, 2009.

[25] J. D. Stewart, "Foot drop: where, why and what to do?" *Practical Neurology*, vol. 8, no. 3, pp. 158–169, 2008.

[26] M. Buonocore and C. Bonezzi, "Are hot-burning sensations produced by the axonal damage of afferent unmyelinated fibres?" *Minerva Anestesiologica*, vol. 72, no. 5, pp. 321–327, 2006.