# The Relationship Between Visual Pain Score and Neutrophil-to-Lymphocyte Ratio in Cases of Radiculopathy Treated with Transforaminal Epidural Steroid Injections

## 🕑 Eylem Burcu Kahraman Özlü, 🕲 Dilara Başgül Sagiri, 🕲 Arif Tarkan Çalışaneller

Department of Neurosurgery, Haydarpaşa Numune Training and Research Hospital, İstanbul, Türkiye

#### ABSTRACT

**Objective:** This study aimed to evaluate whether there was a significant relationship between visual pain score and neutrophil/lymphocyte ratio values over time in patients who were treated with transforaminal epidural steroids for radicular pain.

**Materials and Methods:** The study included 32 cases admitted to our hospital's neurosurgery clinic with the complaint of radicular pain. Patients' age, gender, visual pain scale (VPS) and NLR values on the operation day (Day 0), 15<sup>th</sup> and 30<sup>th</sup> days of the procedure were evaluated retrospectively.

**Results:** The mean age of 11 male (34.4%) and 21 female (65.6%) patients was found to be 54.8±11.7 (31–81). For all cases, the mean VPS score on the day 0 was 7.13±1.60 (4–10), 4.84±1.94 (1–8) on the day 15, and 4.38±2.28 (1–10) on the day 30. the NLR value measured on the 15<sup>th</sup> day did not show a significant (p>0.05) change compared to the measurement made on the day of the procedure (Day 0). However, the NLR value of the 30<sup>th</sup> day showed a statistically significant decrease (p<0.05) compared to the 0th and 15<sup>th</sup> day measurements.

**Conclusion:** It is thought that the NLR value, which is used as a marker in the literature to evaluate the prognosis of many inflammatory processes, may be a marker associated with inflammatory processes in the epidural space in cases with lumbar radicular pain and that TESI is effective in pain control by suppressing the inflammatory process in the epidural space.

Keywords: Inflamation, neutrophil-to-lymphocyte ratio, transforaminal epidural steroid injections

How to cite this article: Kahraman Özlü EB, Başgül Sagiri D, Çalışaneller AT. The Relationship Between Visual Pain Score and Neutrophil-to-Lymphocyte Ratio in Cases of Radiculopathy Treated with Transforaminal Epidural Steroid Injections. CM 2024;16(3):180-184

## **INTRODUCTION**

It is thought that 80–85% of the population has experienced low back pain at least once in their lives. While 80–90% of acute low back pain regresses within 6–8 weeks even if no treatment is given, 20–50% of it recurs in the following year, and 5% of it becomes chronic and lasts longer than 6 months.<sup>[1]</sup> Intervertebral disc disorders are considered to be the most common cause of low back pain. In addition, 10– 15% of these cases need surgical treatment.<sup>[2]</sup> In other cases, resting, exercise, lifestyle changes, medical treatments and physical therapy can be effective in pain control. Even if radicular pain is associated with root irritation in many cases, the inability to achieve pain control by removal of the pressure through surgery in some patients and the reduction of pain without operation suggested that there were also causes of pain other than mechanical pressure. In the literature, the existence of local inflammation is discussed at this point. The emergence of inflammatory agents in the degenerated disc, facet joint or epidural distance increases the sensitivity of the root towards irritation.<sup>[3,4]</sup> The presence of t-lymphocytes, macrophages, phospholipase A-2 and pain-related neuropeptides in the epidural space and degen



Address for Correspondence: Eylem Burcu Kahraman Özlü, Department of Neurosurgery, Haydarpaşa Numune Training and Research Hospital, İstanbul, Türkiye E-mail: burcu.kahraman@gmail.com ORCID ID: 0000-0003-1662-1103 Received date: 10.05.2024 Revised date: 23.07.2024 Accepted date: 29.07.2024 Online date: 31.07.2024

Comprehensive Medicine published by Kare Media. OPEN ACCESS This is an open access article under the CC BY-NC license (http://creativecommons.org/licenses/by-nc/4.0/). erated disc material in the studies also supports the opinions on the relationship between radiculopathy and the inflammatory process.<sup>[5]</sup> Leukocyte, neutrophil, lymphocyte, platelet and monocyte counts are frequently used as inflammation markers. In recent years, neutrophil-to-lymphocyte ratio (NLR) has been frequently mentioned as an inflammatory marker for many pathologies. There are numerous publications recommending the use of NLR as an important marker indicating the severity of endothelial dysfunction and inflammation in the acute period in different clinical situations.<sup>[6,7]</sup>

Transforaminal epidural steroid injection (TESI) is a non-surgical minimally invasive treatment method that has been often used in recent years for patients with lumbar discopathy without progressive motor defects presenting with radicular symptoms. Studies have also revealed that transforaminal epidural steroid and local anesthetic injection is an effective and safe treatment method in radicular pain control.<sup>[8]</sup> It is known that prostaglandins, tumor necrosis factor (TNF), Interleukin 1 (IL-1) and IL-6 inflammatory markers occur in the epidural space in intervertebral disc disorders. Epidural steroid is thought to be effective in the treatment of local inflammatory processes that cause radicular pain. Transforaminal epidural injection is often preferred to transfer high concentrations of steroids to the target tissue.

In our study, it was aimed to evaluate whether there was a significant relationship between visual pain score (VPS) and NLR values over time in patients receiving TESI for radicular pain.

## MATERIALS and METHODS

This study was discussed and approved at Training Planning Board (TPB) meeting of our hospital on 26.03.2024. Study consent for retrospective studies is evaluated by the Training Planning Board (TPB) in our hospital.

Thirty-two cases who received TESI in the Neurosurgery Clinic of our hospital between 2023 and 2024 were evaluated retrospectively based on the hospital information system data.

The study included patients, who were admitted to the neurosurgery outpatient clinic of our hospital with the complaint of radicular pain, whose examinations revealed degenerative findings such as disc herniation and spinal stenosis, who did not have motor defects and progressive neurological deficits, and who received transforaminal epidural steroid injection for pain treatment. Cases with extruded, sequestered disc herniation and motor deficits or progressive neurological deficits underwent surgical intervention and were excluded from the study. In addition, cases with spinal tumor, spinal fracture, uncontrolled diabetes, uncontrolled hypertension, heart diseases,

malignancy and bleeding diathesis, infection, clotting disorder, and the presence of a history of TESI in the last 3 months were accepted as the exclusion criteria of the study. Patients' other comorbidities were not used as exclusion or inclusion criteria.

Scopy-controlled TESI was applied to the distance determined by the radiological imaging and neurological examination findings of the patients evaluated during the brain and nerve surgery polyclinic admissions of our hospital. All cases were evaluated with a standard VPS scale on the day of the procedure and during follow-up examinations.

Patients' age, gender, visual pain scale (VPS) and NLR values on the operation day (Day 0), 15<sup>th</sup> and 30<sup>th</sup> days of the procedure were evaluated retrospectively. NLR values were obtained by our hospital laboratory complete blood count examination, and the normal NLR range in our hospital laboratory was accepted to be 0.00–3.13.

All patients were verbally informed in detail regarding the procedure during their admissions to the outpatient clinic, and any questions they had before the procedure were answered, and their written consent was obtained. The detailed surgical technique of the TESI procedure was described in another article.<sup>[9]</sup> Briefly, the patients were not sedated during the procedure, all procedures were performed under local anesthesia. The procedure was transforaminally performed in all patients. TESI procedure was performed in the operating room with C-arm fluoroscopy. After sterile draping, 5 mg of 0.5% bupivacaine (1 ml) and 40 mg methylprednisolone acetate (1 ml) (2 ml in total) were mixed for each level and injected into the target distance stained with contrast material with fluoroscopy control. Following the procedure, the patients were monitored in the clinic for 1 hour. Their medications, except for anticoagulant-antiaggregant treatments, were continued including the day of the procedure. Anticoagulant and antiplatelet treatments were recommended to be initiated the next day. Patients were called for control on the 15<sup>th</sup> and 30<sup>th</sup> days, which we also use in routine clinical follow-up, and their VAS and NLR values were checked.

## **Statistical Analysis**

In the descriptive statistics of the data, mean, standard deviation, median, lowest, highest, frequency and ratio values were used. The distribution of the variables was measured with Kolmogorov Smirnov and Shapiro-Wilk tests. Wilcoxon test was used to analyze the dependent quantitative data of the study. Spearman correlation analysis was used for the correlation analysis. In the analyses, IBM SPSS 27.0 software based in USA was used.

#### Table 1. Demographics of cases

	Min-max	Median	Mean±SD	n	%
Age Gender	31.0-81.0	52.5	54.8±11.7		
Male				11	34.4
Female				21	65.6

SD: Standard deviation

## RESULTS

The study included 32 patients (11 males, 21 females) who received TESI for the treatment of radicular pain in our clinic between 2023-2024 and whose VPS and NLR values could be reached on days 0, 15 and 30. The mean age of the patients was found to be  $54.8\pm11.7$  (Table 1).

For all cases, the mean VPS score on the day 0 was  $7.13\pm1.60$  (4–10),  $4.84\pm1.94$  (1–8) on the day 15, and  $4.38\pm2.28$  (1–10) on the day 30. The reference NLR range used in the laboratory tests of our hospital was determined as 0.00-3.13. The mean NLR values for all cases were calculated to be  $2.70\pm1.83$  on the day 0,  $2.46\pm1.09$  on day the 15, and  $1.99\pm0.53$  on the day 30.

The VPS score values of the patients on the  $15^{th}$  and  $30^{th}$  days decreased statistically and significantly (p<0.05) compared to the values on the day of the procedure (Day 0). The VPS score of the  $30^{th}$  day did not show a significant (p>0.05) change compared to the  $15^{th}$  day evaluation.

In addition, the NLR value measured on the  $15^{th}$  day did not show a significant (p>0.05) change compared to the measurement made on the day of the procedure (Day 0). However, the NLR value of the  $30^{th}$  day showed a statistically sig-

Table 2. VPS and NLR values by days D\*\* Min-max Median Mean±SD p\* VPS 0<sup>th</sup> day 4.00-10.00 7.00 7.13±1.60 15<sup>th</sup> day 1.00-8.00 5.00 4.84±1.94 0.000 W 4.00 30<sup>th</sup> day 1.00-10.00  $4.38 \pm 2.28$ 0.000 0.115 W NLR 0<sup>th</sup> day 0.60-9.40 2.21 2.70±1.83 15<sup>th</sup> day 1.25-6.86 2.14 2.46±1.09 0.568 W 30<sup>th</sup> day 0.91-3.45 2.02  $1.99 \pm 0.53$ 0.044 0.001 w

w: Wilcoxon test. \*: Change according to day 0; \*\*: Change according to day 15. VPS: Visual pain score; NLR: Neutrophil-to-lymphocyte ratio; SD: Standard deviation

nificant decrease (p<0.05) compared to the  $0^{th}$  and  $15^{th}$  day measurements (Table 2) (Fig. 1).

No significant (p>0.05) correlation was observed between the VPS score and NLR value evaluated on days 0, 15 and 30. No significant (p>0.05) correlation was observed between the 1<sup>st</sup> and 2<sup>nd</sup> measurements, 1<sup>st</sup> and 3<sup>rd</sup> measurements, 2<sup>nd</sup> and 3<sup>rd</sup> measurements regarding VPS scores and NLR changes (Table 3).

## DISCUSSION

Radicular pain is a commonly observed complaint in the daily life of individuals, disrupting the quality of life and causing serious labor loss. The estimated one-year prevalence of lumbosacral radicular pain varies from 3% to 14%. <sup>[10]</sup> Causes such as mechanical root compression, infection and instability due to intervertebral disc protrusion are the main pathologies that are associated with radicular pain.



Table 3. VPS scores and NLR changes										
	NLR									
	Day 0	Day 15	Day 30	0/15 day change	0/30 day change	15/30 day change				
VPS										
r	-0.310	-0.028	-0.095	-0.166	-0.343	-0.293				
р	0.085	0.880	0.606	0.363	0.055	0.103				

Spearman corelation. VPS: Visual pain score; NLR: Neutrophil-to-lymphocyte ratio

However, it may not always be accurate to associate radicular pain with root compression. In McRae's study,<sup>[11]</sup> a postmortem case with roughly 40% disc protrusion but no history of sciatalgia is mentioned. In MRI imaging, up to 60% of radiological root compression findings that do not reveal clinical findings are mentioned.<sup>[12]</sup> Even if mechanical compression is not surgically decompressed in the majority of cases without neurological deficits with complaints of sciatalgia, the findings may regress within 4-6 weeks through conservative methods.[3] Apart from medical treatments, physical therapy applications, and conservative methods such as rest, TESI is a common non-surgical intervention in cases with radicular pain. In our clinic, we frequently perform the TESI procedure for radicular pain control in patients with radicular pain for various reasons and without accompanying progressive motor deficit.

The mechanism of radicular pain is thought to be increased vascular permeability due to inflammation in the facet joint, intervertebral disc and epidural space, and secondary root edema emergence. The increased amount of neutrophils in this inflammatory environment causes the production of large amounts of nitric oxide, which has been revealed to be associated with pain and worsens the clinical situation.<sup>[13]</sup> The steroid applied in TESI directly and/or indirectly reduces the synthesis of inflammatory agents such as arachidonic acid, phospholipases, prostaglandins, leukotrienes and prevents their accumulation in the epidural space. The steroids used improve both early inflammation (edema, fibrin accumulation, capillary dilatation, leukocyte aggregation and phagocytosis) and late effects (capillary and fibroblast proliferation, collagen accumulation and cicatrization).<sup>[3]</sup> In this way, acute inflammation is limited, and pain control is enabled.<sup>[14]</sup> Steroids also modulate inflammatory responses by suppressing pro-inflammatory genes, reducing leukocyte migration into the environment, and preventing the accumulation

of macrophages in the environment. These effects can prevent local inflammation by using TESE.

Neutrophil-to-lymphocyte ratio (NLR) is a practical and inexpensive inflammatory marker that can be reached by a simple laboratory examination such as a complete blood count. NLR value is also used as a prognostic marker for cardiac events, ischemic stroke, chronic diseases, cancers and infectious diseases in many studies.<sup>[15–17]</sup> There are also studies focusing on the presence of inflammation in intervertebral disc pathologies and changes reflected in inflammatory markers. Bozkurt et al.<sup>[18]</sup> also reported a significant positive correlation between VPS score, NLR value and pain severity evaluated in the preoperative and postoperative period in patients with lumbar disc hernia.<sup>[13]</sup>

In our study, after TESI procedure, a significant regression and correlation was found in NLR and VPS values in the 1<sup>st</sup> month compared to the evaluation made on the day of the procedure. In the evaluation made on the 15<sup>th</sup> day, although the VPS scores decreased arithmetically compared to the evaluation made on the 30<sup>th</sup> day, it did not show a statistically significant decrease. However, NLR values decreased statistically significantly on the 30<sup>th</sup> day compared to the values measured on the operation day and 15<sup>th</sup> day. It was thought that the VPS values that regressed in the early period after the procedure may be related to the early effect of lidocaine, which is a short-acting local anesthetic used in TESI led to a decrease in NLR and VPS values on the 30<sup>th</sup> day.

Not excluding the other pathologies that may affect NLR, lack of a control group and limited number of cases stand out as the limitations of our study. It is thought that more significant results can be obtained in a larger patient series in which all pathologies that will affect the NLR value are excluded.

# CONCLUSION

As a result, it can be put forward that the NLR value, which is used as a marker in evaluating the prognosis of many inflammatory processes in the literature, can also be considered as a marker in lumbar radicular pain cases associated with inflammatory processes in the epidural distance, and that TESI is an effective application in pain control by suppressing this inflammatory process. In cases where high NLR values were reported before the procedure, inflammation suppressed by TESE may provide better pain control, which can be evaluated through larger series of studies.

#### Disclosures

**Ethics Committee Approval:** The study was approved by the Haydarpaşa Numune Training and Research Hospital Ethics Committee (No: E-62977267-771-240825582, Date: 26/03/2024).

**Authorship Contributions:** Concept: A.T.Ç., E.B.K.Ö.; Design: E.B.K.Ö.; Supervision: A.T.Ç.; Materials: D.B.S., E.B.K.Ö.; Data Collection or Processing: D.B.S., E.B.K.Ö.; Analysis or Interpretation: E.B.K.Ö., A.T.Ç.; Literature Search: D.B.S., E.B.K.Ö.; Writing: E.B.K.Ö.; Critical review: A.T.Ç., E.B.K.Ö.

**Conflict of Interest:** No conflict of interest was declared by the authors.

**Informed Consent:** Written informed consent was obtained from all patients.

**Use of AI for Writing Assistance:** No support was received from any artificial intelligence-supported source for the study.

**Financial Disclosure:** The authors declared that this study received no financial support.

Peer-review: Externally peer reviewed.

## REFERENCES

- 1. Reilly PA. Occupational low back pain. J Rheumatol 2001;28:225-6.
- Deyo RA, Mirza SK, Martin BI. Back pain prevalence and visit rates: estimates from U.S. national surveys, 2002. Spine (Phila Pa 1976) 2006;31:2724–7. [CrossRef]
- McLain RF, Kapural L, Mekhail NA. Epidural steroid therapy for back and leg pain: mechanisms of action and efficacy. Spine J 2005;5:191–201. [CrossRef]
- Manchikanti L, Cash KA, Pampati V, Falco FJ. Transforaminal epidural injections in chronic lumbar disc herniation: a randomized, double-blind, active-control trial. Pain Physician 2014;17:E489–501. [CrossRef]
- Saal JS, Franson RC, Dobrow R, Saal JA, White AH, Goldthwaite N. High levels of inflammatory phospholipase A2 activity in lumbar disc herniations. Spine (Phila Pa 1976) 1990;15:674–8. [CrossRef]

- Balta S, Cakar M, Demirkol S, Arslan Z, Akhan M. Higher neutrophil to lymhocyte ratio in patients with metabolic syndrome. Clin Appl Thromb Hemost 2013;19:579. [CrossRef]
- Manchikanti L, Datta S, Gupta S, Munglani R, Bryce DA, Ward SP, et al. A critical review of the American Pain Society clinical practice guidelines for interventional techniques: part 2. Therapeutic interventions. Pain Physician 2010;13:E215–64. [CrossRef]
- Cohen SP, Bicket MC, Jamison D, Wilkinson I, Rathmell JP. Epidural steroids: a comprehensive, evidence-based review. Reg Anesth Pain Med 2013;38:175–200. [CrossRef]
- Kahraman Özlü EB. Transforaminal epidural injections in neurosurgical clinical practice: a single surgeon's experience. Med J Bakirkoy 2023;19:319–23. [CrossRef]
- Younes M, Béjia I, Aguir Z, Letaief M, Hassen-Zrour S, Touzi M, et al. Prevalence and risk factors of disk-related sciatica in an urban population in Tunisia. Joint Bone Spine 2006;73:538–42. [CrossRef]
- 11. McRae DL. Asymptomatic intervertebral disc protrusions. Acta Radiol 1956;46:9-27. [CrossRef]
- Boden SD, Davis DO, Dina TS, Patronas NJ, Wiesel SW. Abnormal magnetic-resonance scans of the lumbar spine in asymptomatic subjects. A prospective investigation. J Bone Joint Surg Am 1990;72:403–8. [CrossRef]
- Firidin MN, Akyüz ME. Preoperative and postoperative diagnostic efficiency of multi-inflammatory index on pain scoring of degenerated intervertebral disc. Adv Clin Exp Med 2022;31:947–52. [CrossRef]
- 14. Viner N. Intractable sciatica. The sacral epidural injection; an effective method of giving relief. Can Med Assoc J 1925;15:630–4.
- Balta S, Cakar M, Demirkol S, Arslan Z, Akhan M. Higher neutrophil to lymhocyte ratio in patients with metabolic syndrome. Clin Appl Thromb Hemost 2013;19:579. [CrossRef]
- Lee DH, Lee BK, Lee SM, Cho YS, Yun SW. Association of neutrophil-to-lymphocyte and platelet-to-lymphocyte ratios with in-hospital mortality in the early phase of severe trauma. Ulus Travma Acil Cerrahi Derg 2021;27:290–5. [CrossRef]
- Lo YT, Lim VY, Ng M, Tan YH, Chiang J, Chang EWY, et al. A prognostic model using post-steroid neutrophil-lymphocyte ratio predicts overall survival in primary central nervous system lymphoma. Cancers (Basel) 2022;14:1818. [CrossRef]
- Bozkurt H, Arac D, Cigdem B. The effect of the preoperative uric acid level and neutrophil lymphocyte ratio on preoperative and postop- erative visual pain scores in patients with lumbar disc hernia: a cross- sectional study. Turk Neurosurg 2019;29:705–9. [CrossRef]