



Internal Neurolysis (Nerve Combing) for Trigeminal Neuralgia without Neurovascular Compression

Neurólise interna (nerve combing) para neuralgia do trigêmeo sem compressão neurovascular

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Abstract

Objective The aim of the present study was to describe and evaluate the initial and the long-term clinical outcome of internal neurolysis (IN) for trigeminal neuralgia (TN) without neurovascular compression (NVC).

Methods A total of 170 patients diagnosed with TN were treated by posterior fossa exploration, during the period between April 2012 and October 2019. The patients were divided into two groups: Group A (50 patients) was treated by IN and Group B (120 patients) received microvascular decompression (MVD). Surgical outcomes and post-operative complications were compared between the two groups. Pain intensity was assessed by the Barrow Neurological Institute (BNI) pain intensity score and BNI facial numbness score. Pain recurrence was statistically evaluated with Kaplan-Meier analysis.

Results Pain was completely relieved in 44 patients (88%) who underwent IN (group A); 3 (6%) experienced occasional pain but did not require medication (BNI 2). In group B, 113 (94%) experienced immediate pain relief after MVD. The median duration of follow-ups was 4 years (6 months to 7.5 years). In Group A, there was a meantime recurrence of 27 months in 3 patients (6%). The recurrence in Group B was of 5.8% during the follow-up period. There were no statistically significant differences in the surgical outcomes between the two groups. All patients with IN experienced some degree of numbness, 88% of the cases resolved in 6 months, on average.

Conclusion Internal neurolysis is an effective, safe and durable treatment option for trigeminal neuralgia when NVC is absent.

Keywords

- ▶ internal neurolysis
- ▶ nerve combing
- ▶ trigeminal neuralgia
- ▶ root entry zone
- ▶ neurovascular compression
- ▶ microvascular decompression

Resumo

Objetivo O propósito do presente estudo foi descrever e avaliar o resultado clínico inicial e a longo prazo da neurólise interna (IN, na sigla em inglês) para neuralgia do trigêmeo (TN, na sigla em inglês) sem compressão neurovascular (NVC, na sigla em inglês).

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Palavras-chave

- ▶ neurólise interna
- ▶ nerve combing
- ▶ neuralgia do trigêmeo
- ▶ zona de entrada da raiz
- ▶ compressão neurovascular
- ▶ descompressão microvascular

Métodos. Um total de 170 pacientes diagnosticados com TN foram tratados pela exploração da fossa posterior, durante o período entre abril de 2012 e outubro de 2019. Os pacientes foram divididos em dois grupos: o grupo A foi tratado com IN (50 pacientes) e o grupo B recebeu descompressão microvascular (DMV) (120 pacientes). Resultados cirúrgicos e complicações pós-operatórias foram comparados entre os dois grupos. A intensidade da dor foi avaliada pelo escore de intensidade de dor do Barrow Neurological Institute (BNI, na sigla em inglês) e pelo escore de dormência facial do BNI. A recorrência da dor foi avaliada estatisticamente com a análise de Kaplan-Meier.

Resultados A dor foi completamente aliviada em 44 pacientes (88%) submetidos a IN (grupo A); em 3 (6%) houve dor ocasional, mas não necessitaram de medicação (BNI 2). No grupo B, 113 (94%) experimentaram alívio imediato da dor após DMV. A duração média do acompanhamento foi de 4 anos (6 meses a 7,5 anos). No grupo A, houve recorrência em três pacientes (6%). O tempo médio de recorrência foi de 27 meses. A recorrência no grupo B foi de 5,8% nesse período de acompanhamento. Não houve diferenças estatisticamente significativas nos resultados cirúrgicos entre os dois grupos. Todos os pacientes com IN experimentaram algum grau de dormência, em 88% dos casos, resolvidos em 6 meses, em média.

Conclusões A IN é uma opção de tratamento atraente, eficaz, segura e durável para a neuralgia do trigêmeo quando a NVC está ausente.

Introduction

Vascular arterial contacts with the dorsal root of the trigeminal nerve were first described in 1929, by Dandy.¹ These findings were characterized in subsequent communications.^{2,3} In his 1934 publication, "Concerning the cause of trigeminal neuralgia," Dandy describes the total or subtotal division of the trigeminal root in 215 cases.³ Through a lateral suboccipital craniectomy without the aid of illumination and magnification afforded by an operating microscope, Dandy was able to identify compression on the trigeminal nerve root by a superior cerebellar artery (SCA) loop, in 66 cases (30.7%); compression by a branch of the superior petrosal vein in 30 (14%), and a dolichoectatic basilar artery in 6 (2.7%), for a total of 102 cases (47%).³⁻⁵

Dandy postulated that the vascular compression of the trigeminal roots was a major cause of tic douloureux.^{3,6} However, vascular transposition was not used by Dandy to attempt to relieve the pain; rather, selective section of the trigeminal nerves was performed.

In 1959, Gardner et al. reported a series of 100 patients in whom an extradural subtemporal approach was used to manipulate and free up the sensory root of the trigeminal nerve from its dural sleeve. Exploration of the trigeminal nerve through the posterior fossa, without magnification, was performed in two patients with recurrence. In one patient, an arterial loop was found lying against the nerve at the pons. Pain was completely relieved by separating the vessel from the nerve with a piece of Gelfoam. The second patient was relieved of pain following removal of a large meningioma compressing the trigeminal nerve. It was concluded that the critical part of the operation appeared to be a neurolysis or manipulation of the sensory root of the trigeminal nerve at the point where it crosses the apex of the petrous bone. Gentle manipulation of

the sensory root could be done through a middle or a posterior fossa approach. "The cause of trigeminal neuralgia, therefore, presumably lies in the sensory root."⁷

In 1962, Gardner published "Concerning the mechanism of trigeminal neuralgia and hemifacial spasm." He reported a series of suboccipital approaches that followed a failed middle fossa procedure in 18 cases for trigeminal neuralgia (TN) from 1955 to 1961. Although in the era that predated neurosurgical use of the operative microscope, he found an artery loop that compressed and encircled the nerve in 6 cases.⁸ There was no mention of the treatment those 6 patients received. Rand reported in 1981 a personal communication with Gardner, where he was informed that the treatment of TN involved either removing the offending lesion such as a tumor, or performing a vascular decompression by dissecting the arterial compressing loop away from the trigeminal root, and placing Gelfoam in between at the level of the pons.⁹

Jannetta, in 1967, still a neurosurgical resident at the time, participated in a workshop titled "Structural Mechanisms of Trigeminal Neuralgia."¹⁰ Reporting five patients with TN, he operated using a binocular dissecting microscope following exposure of the nerve through the transtentorial subtemporal approach to the cerebellopontine angle. He found that the trigeminal nerve was mildly to severely distorted and compressed by one or more small tortuous arteries that appeared to be branches of the SCA. In four out of five patients, he was able to free the artery from the fine to dense arachnoidal membranes, allowing the vessel to assume a new position away from the nerve. He performed a partial to total section of the portio major in all cases. Jannetta hypothesized that it was possible that this minor arterial distortion of the trigeminal nerve at the pons may be a contributing factor in TN. "This possible definitive procedure, namely, release of the artery without nerve section, is planned in a future series of patients."¹⁰

Table 1 Incidence of neurovascular compression

Authors	Patients	With Vascular Conflict (%)	No vascular Conflict (%)
Van Loveren et al. ¹⁷ 1982	50	82	18
Piatt et al. ¹⁸ 1984	103	76.7	23.3
Zorman et al. ¹⁹ 1984	118	76.3	23.7
Benderson et al. ²⁰ 1989	243	87.7	12.3
Klun ²¹ 1992	215	80.5	19.5
Baechli et al. ²² 2007	40	87.5	12.5
Leal et al. ²³ 2010	100	91	9
Revuelta et al. ²⁴ 2013	271	83.8	16.2
Ko et al. ²⁵ 2015	156	82.7	17.3
Hitchon et al. ²⁶ 2019	69	87	13
Yang et al. ²⁷ 2019	298	88.6	11.4

In a 1974 letter published in JAMA, Jannetta described his 8-year experience with 150 cases of tic douloureux. He reported having found compression-distortion of the nerves with resultant pain control after vascular decompression.¹¹ The approach used was not described. This was followed by complete manuscripts published in 1976 and 1977.^{12,13}

Jannetta became the first neurosurgeon to explore the cranial nerves within the posterior fossa by using an operating microscope, modifying Dandý's original suboccipital approach and developing a set of microneurosurgical instruments. Therefore, the idea that TN was a neurovascular compression was revived, and the treatment of this problem by microvascular decompression (MVD) was developed.

In spite of some initial controversy, today MVD represents the best and the most widely used surgical treatment for TN, because it offers the best long-term cure rates and preserves facial sensation.¹⁴⁻¹⁶

Neurovascular compression (NVC) of the trigeminal nerve has been identified in most cases of TN; however, the absence of NVC in TN is also common and has been described in the literature in 9 to 23.7% of the cases (→ **Table 1**).¹⁷⁻²⁷ Many surgical modalities are appropriate for MVD candidates who exhibit no NVC on magnetic resonance imaging (MRI), such as percutaneous balloon compression, radiofrequency thermocoagulation, glycerol rhizolysis, and radiosurgical techniques.

In the case of negative microsurgical suboccipital retrosigmoid exploration of the trigeminal root, the intraoperative management options have not been established. Surgical strategies include internal neurolysis (IN), trigeminal root compression (TRC) and partial sensory rhizotomy (PSR).

Internal neurolysis, also referred to as "nerve combing" or "neurocombing", is a procedure in which all or portions of the trigeminal nerve are divided longitudinally along its fibers between the pons and the porus trigeminus.^{28,29} The objective of the present study is to evaluate the efficacy of IN in patients with TN without NVC and evaluate the long-term outcome.

Materials and Methods

This is an observational prospective cohort study, based on the prospective observation of 184 patients diagnosed with TN (TN Type 1). The patients were treated by posterior fossa exploration, during the period between April 2012 and October 2019.

Sixteen of the patients were excluded from the present study due to the diagnosis of a cerebellopontine angle tumor during the preoperative evaluation, using magnetic resonance imaging (MRI) of the brain. Data was obtained from hospital records and patient interviews.

Patients included in the cohort study were diagnosed with TN (TN type 1), characterized by paroxysmal and lancinating pain in the distribution area of the trigeminal nerve branches. These patients experienced unilateral, recurrent, stereotyped attacks, generally provoked by stimulated trigger points, including facial movements and changes in temperature.

Indications for surgery included insufficient pain relief with drug treatment and or unacceptable medication side effects. Surgical procedures involved MVD and percutaneous procedures (balloon compression and radiofrequency thermocoagulation).

General physical and neurological examinations were clinically performed to evaluate the patients. In addition, brain MRI, in combination with three high-resolution sequences (T2-weighted 3D, 3D TOF-MRA, and 3D T1-Gad) with fine cuts in the trigeminal nerves, were used for detection of possible neurovascular contact. Routine presurgical examinations were performed.

A surgical treatment algorithm was used for patients with TN (→ **Fig. 1**). The advantages and disadvantages of the different surgical treatment options were explained to the patients, and they decided according to their preference. Only those who chose MVD were included in the present study.

The 170 patients with trigeminal neuralgia were divided into two groups. The first group, Group A, consisted of 50 patients who were treated with internal neurolysis (IN). Group B was made up of 120 patients treated with MVD (4 patients treated with MVD and IN). Each group had one patient who underwent two procedures at different times for bilateral trigeminal neuralgia, which was considered to be a separate case for the purposes of analysis. Only one patient (2%) in Group A had a previous surgery. This patient had undergone two percutaneous balloon surgeries without adequate response in pain management. The follow-up period was from 6 months to 7.5 years (average 4 years). In patients with recurrence, the time at which the recurrence occurred was considered a follow-up endpoint.

The Barrow Neurological Institute (BNI) pain intensity score and facial numbness score table was used in the postoperative evaluation (→ **Table 2**). The grade of NVC was established on a scale from I to III. In grade I, the vessel was in contact with the root without any visible indentation at the root. In grade II, there was root displacement and/or distortion, and in grade III there was marked indentation in the root. The absence of contact or compression in the root was recorded as grade 0.

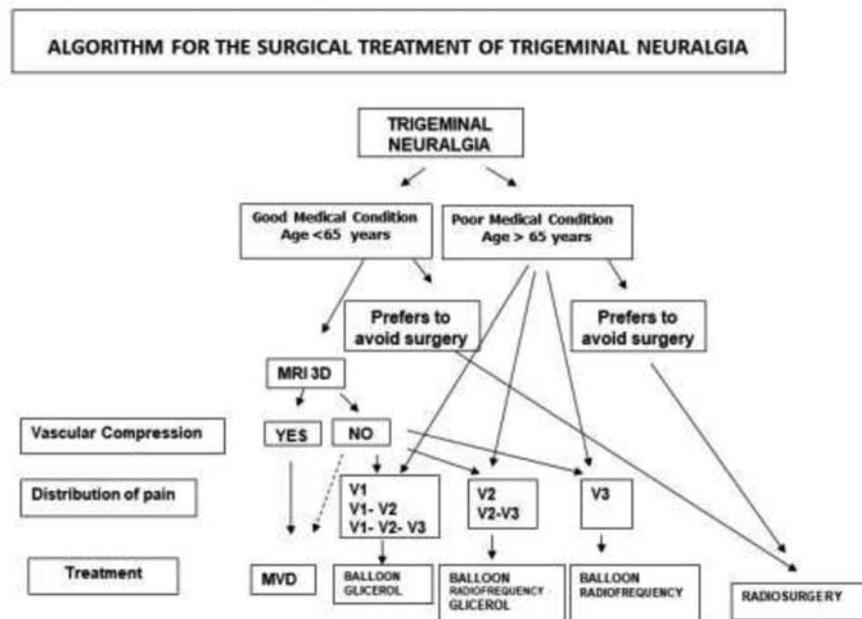


Fig. 1 Algorithm of surgical treatment for trigeminal neuralgia.

Table 2 Barrow Neurological Institute (BNI) pain intensity score and facial numbness score

(P) Evaluation of pain relief by the BNI pain intensity score	
1	No pain, no medication
2	Occasional pain, not requiring medication
3	Some pain, adequately controlled with medication
4	Some pain, not adequately controlled with medication
5	Severe pain / no pain relief
(N) Evaluation of numbness by the BNI facial numbness score	
1	No facial numbness
2	Mild facial numbness, not bothersome
3	Facial numbness, somewhat bothersome
4	Facial numbness, very bothersome

Statistical Analysis

Epidemiological data, clinical history, operative findings, and clinical results were analyzed.

The data was stored on a Microsoft Excel (Microsoft Corporation, Redmond, WA, USA) spreadsheet. The mean, standard deviation (SD), minimum and maximum values were used to summarize the quantitative variables. To summarize the qualitative variables, frequency tables (simple and crossed) were used, expressing the final results in absolute and relative frequency. Ratio comparison test and Kaplan-Meier curves were calculated to determine the ratio of patients who responded to the operation. All hypothesis tests were performed considering a type I error (α) equal to 0.05. Statistical analysis was performed by the R version 4.0.2 statistical software (R Foundation, Vienna, Austria).

Surgical Procedure

The patients were placed in supine or in lateral decubitus position with the affected side upward, the ear positioned parallel to the floor, and the chin flexed. A linear 3.5 cm incision was made behind the ear, starting at the top of the pinna and extending to the mastoid tip within the hairline, between ~ 4 and 5 cm.

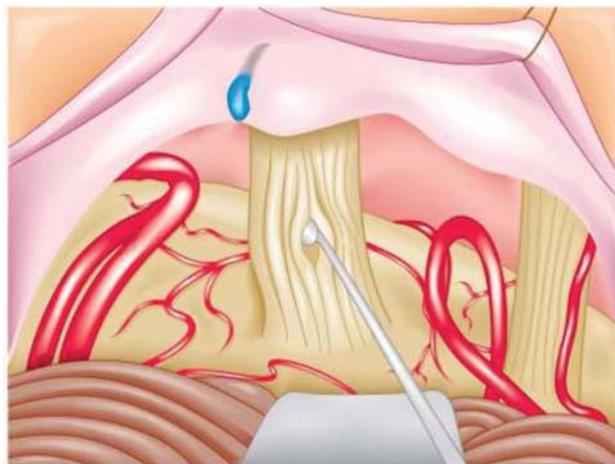
The burr hole was made in the asterion, and a 3×3 cm retrosigmoid craniotomy is performed using a high-speed drill and Kerrison Rongeurs to expose the borders of the transverse sinus, the sigmoid sinus, and their intersection. Any exposed mastoid air cells were carefully waxed. The dura was opened in a rectangular fashion along the inferior border of the transverse sinus with the base toward the sigmoid sinus. The dural edges were elevated with sutures extended over the craniotomy dressings. Under direct microscopic visualization, gentle traction of the cerebellum took place with an aspirator and bipolar, protected by cottonoids. Meticulous arachnoid dissection was made and cerebral spinal fluid (CSF) was aspirated.

The superior petrosal vein and its main branches were dissected free from the surrounding arachnoid membranes and preserved as much as possible. The entire length of the trigeminal nerve from the pons to the Meckel cave was carefully explored even if a definite vessel was located. Conflicting vessels were dissected and separated from the nerve, a teflon felt was implanted between them. Particular attention was made to prevent a "neo-compression" between the prosthetic material and the nerve root.

When a vascular compression was not found; veins with simple contact or thickness arachnoid adhesions affected branches of the trigeminal nerve, according to preoperative pain location and intraoperative findings, were longitudinally divided along its fibers, using a Rothon microdissector, into 3 to 4 bundles from the root entry point (REP) to the porus



A



B

Fig. 2 Internal neurolysis (IN) for trigeminal neuralgia without neurovascular compression. (A) IN procedure with longitudinal division of the trigeminal nerve using a microdissector. (B) An anatomical drawing of the IN procedure.

trigeminus (→ **Fig. 2**). Finally, compression of the root at the root entry zone (REZ) with bipolar forceps for 10 seconds took place. The dura was closed, filling the craniotomy defect with autogenous bone. Closure of muscle, soft tissue, and skin was performed in layers in a standard fashion.

Results

In the present study, 50 internal neurolysis procedures were performed in patients with TN during the period between April 2012 and October 2019. Of these patients, 40 were female (80%), the age ranged from 25 to 75 years old (mean of 49 years old).

The age distribution according to gender was unimodal (→ **Fig. 3**). The most affected side of the face was the right (68%), the most affected branches being the combination V2, V3 (50%). These clinical and demographic characteristics are shown in → **Table 3** and → **Table 4**. The degree of NVC severity is shown in → **Table 5**. In this series, we found 42 cases (24.7%) with absence of neurovascular conflict.

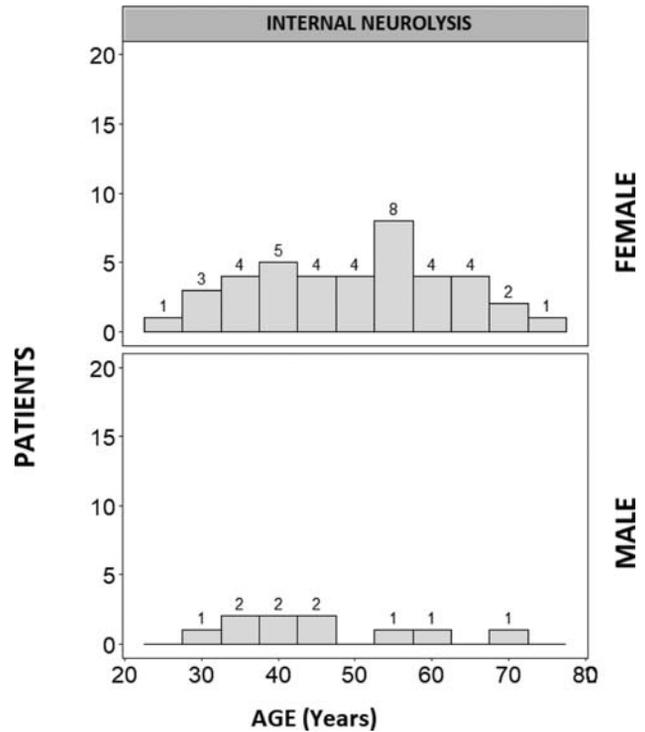


Fig. 3 The Hartigan's Dip Test for Unimodality indicates that the age distribution according to gender was unimodal, female ($p = 0.227$) and male ($p = 0.220$).

Table 3 Demographic and clinical information of the two groups

FEATURE	GROUP A (n = 50)	GROUP B (n = 120)
AGE (years old)	49 (25–75)	50.8 (23–76)
SIDE (Right/Left)	34/16	87/33
GENDER (Female /Male)	40/10	79/41
DISTRIBUTION OF PAIN		
V1	0	0
V2	6 (12%)	13 (10.8%)
V3	9 (18%)	27 (22.5%)
V1,V2	2 (4%)	10 (8.3%)
V2,V3	25 (50%)	51 (42.5%)
V1,V2,V3	8 (16%)	19 (15.8%)

Group A: Patients without vascular compression, Group B: Patients with vascular compression.

Surgical findings in Group A included 42 patients (84%) with no neurovascular conflicts found. Although no neurovascular conflicts were detected, there was presence of arachnoid thickening in 11 patients. The remaining 8 patients (16%) presented with a vein with simple contact (grade 1) to the trigeminal root; 2 of those patients also presented thickened arachnoid adhesions.

In Group B, 116 patients (96.7%) with distortion/displacement (grade II) or indentation (grade III) were surgically treated with MVD; while 4 patients (3.3%) with touching or less severe compression (grade I) underwent an MVD in combination with an IN (→ **Table 5**).

Table 4 Distribution of patients with internal neurolysis according to age

AGE (years old)	Patients	%
≤ 30	4	8
31–40	9	18
41–50	12	24
51–60	15	30
61–70	8	16
71–80	2	4
TOTAL	50	100

Table 5 Distribution of patients according to the grade of severity of neurovascular contact

GRADE	Group A (n = 50)	Group B (n = 120)	TOTAL (n = 170)
0	42 (84%)	0	42 (24.7%)
I	8 (16%)	4 (3.3%)*	12 (7%)
II	0	99 (82.5%)	99 (58.2%)
III	0	17 (14.2%)	17 (10%)

Group A: Internal neurolysis, Group B: Microvascular decompression. *MVD and internal neurolysis were performed.

During the manipulation of the trigeminal nerve, and while IN was performed; some patients experienced bradycardia and arrhythmias that reversed spontaneously. None of these cases reported recurrent bradycardia or arrhythmias requiring any additional treatment.

In the postoperative period, 44 patients (88%) who underwent IN surgical treatment were pain free without any medication, 3 (6%) experienced occasional pain, but did not require medication. Mild facial numbness not bothersome to the patient was presented in 88% of the cases, which resolved in 6 months, on average. (► **Table 6**). In Group B, 94.2% of the

Table 6 Surgical results after surgery using BNI pain intensity score and facial numbness score

SURGICAL RESULTS	GROUP A (n = 50)	GROUP B (n = 120)
(P) Pain intensity score		
1	44 (88%)	113(94.2%)
2	3 (6%)	2 (1.7%)
3	0	1 (0.8%)
4	0	1 (0.8%)
5	3 (6%)	3 (2.5%)
(N) Facial numbness score		
1	0	104 (86.7%)
2	44 (88%)	11 (9.2%)
3	4 (8%)	5 (4.1%)
4	2 (4%)	0

Group A: Internal neurolysis, Group B: Microvascular decompression.

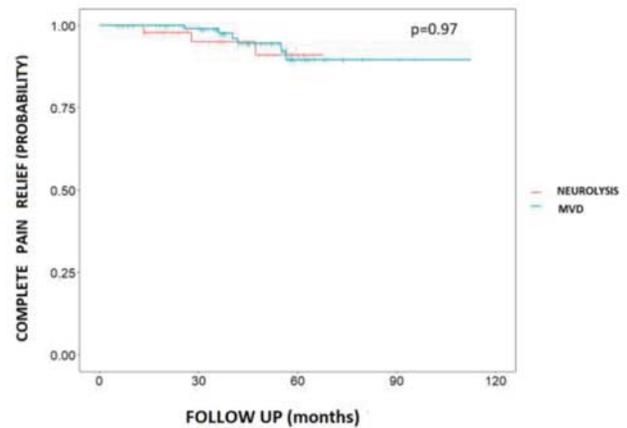


Fig. 4 Kaplan-Meier curves showed that the mean pain-free time was 84.9 (standard error: 2.82) months for neurolysis. For microvascular decompression, the mean was 85.4 (standard error: 1.86) months. The logarithm of range test (“logrank”) shows that the treatments did not show differences throughout the study time with respect to complete pain relief (p = 0.97).

patients who underwent MVD were pain-free, and 16 patients (13.3%) experienced some grade of facial numbness, 4 of them having been treated with DMV and IN surgeries.

Recurrence was defined as transition from pain-free or occasional pain without medication (BNI I and II) to severe pain or requirement of medication (BNI III–V).

The follow-up period was from 6 months to 7.5 years (average 4 years). In group A there was recurrence in 3 patients (6%), one of them with intraoperative finding of a small vein that contacted the trigeminal root, occurred at 12 months; a second patient in whom only adhesions were found recurred at 25 months, and a third patient in whom no neurovascular compression was identified, it recurred at 44 months. Two out of three patients underwent successful balloon compression, and the third continued with medication.

The mean time of recurrence was 27 months. Kaplan-Meier analysis showed that the mean pain-free time was 84.9 (standard error: 2.82) months (► **Fig. 4**).

The recurrence in Group B was of 5.8% in this follow-up period.

The logarithm of range test (“Logrank”) shows that the treatments in the 2 groups did not show differences over time in the study with respect to complete pain relief (p = 0.97).

Complications are summarized in ► **Table 7**. Cerebellar infarction occurred in one patient who also had hemifacial

Table 7 Postoperative complications

COMPLICATION	Group A	Group B
Cerebellar infarction	1 (2%)	
Transient hearing loss	1 (2%)	2 (1.7%)
Meningitis		1 (0.8%)
Transient diplopia		1 (0.8%)
Wound cerebrospinal fluid leak		2 (1.7%)
Nasal cerebrospinal fluid leak		2 (1.7%)

Group A: Internal Neurolysis, Group B: Microvascular decompression.

spasm and had to undergo MVD of the facial nerve during the same procedure. Most of the patients experienced mild transient hearing loss, with resolution in 2 months; the diplopia was caused by paresis of the VI cranial nerve, with complete resolution after 3 months. In cases of nasal CSF leaks, external lumbar drainage was placed for between 5 and 7 days. In 2 cases of wound CSF leaks, placement of new sutures were required, as well as compression and use of acetazolamide. Placement of external lumbar drain was not necessary.

Discussion

Pathophysiology

The leading theory for the pathophysiology of TN is the “ignition” hypothesis proposed by Devor et al in 2002, which proposes three key elements in the genesis of TN: a trigger, an amplifier, and a stop mechanism.³⁰

According to the ignition hypothesis, a neurovascular contact is thought to damage the myelin sheath in the trigeminal nerve, which renders the sensory afferents hyperexcitable by means of ectopic pacemaker sites and ephaptic cross-talk and crossed after discharge between axons.^{30,31} Anatomical studies confirmed that the root entry zone or transition zone of the trigeminal nerve, where the myelination changes from peripheral Schwann cell myelination to central oligodendrocyte myelination, is a site of demyelination in patients with classical TN with a neurovascular contact.^{32,33}

Currently accepted, the decompression of the central myelin alone or decompression of the central-peripheral myelin transitional zone was the mechanism that provided pain relief.

McLaughlin et al., in 1999, postulated that the central myelin of the sensory root of the trigeminal nerve could extend along the entire cisternal segment of this branch. He stated that not only would vascular decompression of the REZ relieve the pain of patients with TN, but also that decompression performed anywhere along the entire length of the nerve (from the pons to entry into the Meckel cave) would be effective.³⁴

Sindou et al., in 2002, suggested that NVC can be located not only at the REZ but also all along the root, and because offending vessels can be multiple in the same patient, he strongly advised that the entire trigeminal nerve from the Meckel cave to the pons should be exposed. He proposed that NVC probably does not entirely explain all the pathogenesis of TN. He thought that there might be several other important factors. Sindou concluded, “It is wise to admit that besides peripheral mechanisms, central phenomena probably play an important additional role.”¹⁶

Peker et al., in 2006, reported his study of one hundred trigeminal nerves from 50 cadaver heads that were examined. “The measurements showed that the central myelin occupies only the initial one-fourth of the trigeminal nerve length. If trigeminal neuralgia is caused exclusively by vascular compression of the central myelin, the problem vessel would always have to be located in this region. However, it is well known that pain from trigeminal neuralgia can resolve after vascular decompression at more distal sites. This sug-

gests that the effects of surgical decompression are caused by another mechanism.”³²

Ko, Lee et al. observed that it is increasingly clear that TN occurs and recurs in the absence of NVC,³⁵ and that NVC of the trigeminal nerve exists in a sizeable population without TN.³⁶ He suggested that NVC is neither a necessary nor sufficient condition for the development of TN. Nevertheless, NVC does play a role in disease pathogenesis.

It is becoming increasingly understood that the pathophysiology of TN is associated with both trigeminal nerve and brain abnormalities.

Importance of Degree of Neurovascular Compression

Increasingly, the contributions of NVC severity is beginning to be better understood.

Sindou reported a series of 362 patients who underwent MVD. Concerning anatomical factors, neither the type of the compressive vessel nor its location along or around the root was found to be significant. However, the severity of compression was important – the more severe the degree of compression, the better the outcome.³⁷

Maarbjerg et al., in 2015, with 3T MRI, evaluated 135 patients with classical trigeminal neuralgia. Neurovascular contact was prevalent both on the symptomatic and asymptomatic side, 89 versus 78%, while severe neurovascular contact (displacement or atrophy of the trigeminal nerve) was highly prevalent on the symptomatic compared with the asymptomatic side (53 versus 13%). Severe neurovascular contact was caused by arteries in 98%.

This concluded that neurovascular contact causing displacement or atrophy of the trigeminal nerve is highly associated with the symptomatic side in classical TN, as opposed to neurovascular contact in general, suggesting that the degree of NVC could thus be important when selecting patients for surgery.³⁸

Hughes et al. published in 2019 the significance of degree of NVC in surgery for TN. Seventy-nine patients were studied on a 3-T MRI. Vascular contact was more common on the asymptomatic side (43/79, 54.4%) compared with the symptomatic side (30/79, 38%). Severe NVC (indentation/deformity of the of the trigeminal nerve) was much more commonly identified on the symptomatic side (47/79, 59.5%) compared with the asymptomatic side (7/79, 8.9%). In his study, the only imaging variable that was a statistically significant predictor of being pain-free without medication following MVD was severe NVC. Patients with severe NVC were 6.36 times more likely to be pain-free following MVD when compared with those without severe NVC.³⁹

According to various studies published with high resolution MRI, the presence of NVC has a sensitivity of 87% to 97% and a specificity between 50% to 100% for TN.^{23,35,39,40}

Recently, it was reported that ultra-high field MRI enabled advanced Diffusion tensor imaging (DTI)-based tractographic reconstruction of the white matter tracts connecting the thalamus and primary somatosensory cortex. Reduced microstructural integrity of thalamic-somatosensory tracts was found ipsilateral to the site of NVC in patients with TN, whereas no differences were observed contralateral to the

site of NVC. These changes implicate a compensatory mechanism for reducing pain sensation, and may provide substantial evidence of secondary white matter disruption at the thalamic-somatosensory level in patients with TN.⁴¹

Trigeminal Neuralgia without Vascular Compression

The absence of vascular compression over the trigeminal nerve has been described in the literature in 9 to 23.7% of cases (– **Table 1**). In our series of 170 patients, 42 (24.7%) presented no vascular conflict, and in 12 cases (7%), a mild contact (grade I) was found; 8 with venous compression were coagulated and cut and IN was performed. In 4 cases with arterial compression, MVD in combination with IN was performed.

It is likely that the number of TN without neurovascular compression (WONVC) will increase more since it is increasingly recognized that vascular contact (grade I) should not be accepted as vascular compression and additional treatment must be done.

In the absence of NVC on imaging, the question arises as to whether recommend a posterior fossa exploration versus percutaneous procedures or radiosurgery.

However, it is challenging for neurosurgeons to deal with cases without visible vascular compression intraoperatively, and there are no well-designed surgical strategies for these patients, which ranged from no manipulation to complete section of the nerve.

In these cases, most neurosurgeons do a meticulous dissection of the arachnoid to expose the entire trigeminal root, followed by a massage and/or a compression with bipolar to the trigeminal nerve or a partial nerve section.

Based on the proposed mechanisms of production of TN related to the compression of the sensitive trigeminal root of the trigeminal nerve in its intracranial route, nondestructive open surgical techniques have historically been directed toward a decompression of the trigeminal nerve at some point in its intracranial trajectory.^{42,43}

In 1955, Shelden et al. suggested that the common denominator to all procedures is operative trauma, introducing a new technique such as pressing with a blunt dissector the portio major nerve fibers of the trigeminal nerve surgically exposed through a middle cranial fossa approach, suggesting that the compression of the trigeminal fibers during the operation is the factor that relieves the pain.⁴⁴

Partial sensory rhizotomy (PSR) is a classical alternative technique used in the absence of NVC.

The response rate is reported in 70 to 87% of patients (– **Table 8**), with a recurrence rate of 29% at 5 years of follow-up.^{21,45–48} The main disadvantages of this ablative treatment are sensory deficits, which occur in 22 to 100% of patients and can lead to additional complications as dysesthesias, corneal reflex impairment and anesthesia dolorosa. Furthermore, it has been reported that a lesser degree of patient satisfaction is associated with side effects after PSR.⁴⁷

Ishikawa et al., in 2002, speculated a different mechanism other than vascular compression.⁴⁹

They suggested that arachnoid thickening or granulomatous adhesion between the root and surrounding structures can cause a tethering effect to the trigeminal root, which

Table 8 Partial sensory rhizotomy. Results related to extent of the rhizotomy

Reference	Number of Cases	Pain Relief After Surgery (%)	Median Follow-Up (months)	Extent of Rhizotomy	Corneal Reflex Impairment/Hyesthesia (%)	Sensory Deficit (%)	Anesthesia dolorosa	Recurrence Rate
Adams et al. ⁴⁵ J Neurol Neurosurg Psychiatry 1982	54	87.3	54	30–100% of trigeminal Nerve	0	100 Painful dysesthesias 7.4%	1 case (1.8%)	11.8% at 5 years
Bederson et al. J Neurosurg 1989	30 (PSR) 56 (PSR + MVD)	70 (PSR) 80 (PSR + MVD)	61.2	Inferior 1/2 to 2/3 of pars Major	4.6	dysesthesias 8.1%	2.3%	NA
Klun et al. ²¹ Neurosurgery 1992	42	86	62.4	1/3 or less of the pars major	0	100	0	Total recurrence rate 49%
Young et al. ⁴⁶ J Neurosurg 1993	83	48 Excellent 22 Good	72	1/3 to 1/2 ventrolateral of nerve root (89%) 2/3 of nerve root (11%)	NA	67	NA	17% at 1 year and 2.6% each year thereafter
Zakrzewska et al. ⁴⁷ Neurosurgery 2005	60	88	68.4	NA	22 "eye problems"	48	NA	28% at 5 years
Terrier et al. ⁴⁸ 2017	22	86.4	67.3	2/3 ventrolateral of pars Major	0	22.7	9% in hemitongue	14.6% at 1 year 31.5% at 5 years

Abbreviations: MVD, microvascular decompression; NA, not available; PSR, partial sensory rhizotomy.

Table 9 Series of intraoperative trigeminal root manipulations

Reference	N° of cases	Treatment	Pain-free immediatly	Recurrence	Facial numbness
Ishikawa et al. ⁴⁹ Journal of Clinical Neuroscience 2002	5	Arachnoidal dissection	80%	20% in 15 months	NR
Revuelta et al. ²⁴ World Neurosurgery 2013	44	Intermittent compression on cisternal segment	100%	27.2% in 10 months	61.4% transient
Cheng et al. ⁵¹ Acta Neurochir 2015	28	Compression with bipolar forceps for 10 seconds	71.4%	38.4% in 46 months	21.4% transient 14.3% permanent
Guo et al. ⁶⁸ Journal of Clinical Neuroscience 2018	19	360-degree circumferential arachnoid dissection	100%	10.5% in 2 years	15.8% transient
Urculo et al. ⁵² Neurocirugia 2020	10	Trigeminal root massage/ intermittent compression	100%	40% in 5 years	NR

Abbreviation: NR, not reported.

would make pulsatile movement of the root more restricted or desynchronized. This could give the root an abnormally high stretching force, which might promote hyperexcitability of the nerve.

Dissection of thick arachnoid around the root along the whole length reversed the root to be straight and flaccid. This speculative mechanism suggests that it is important to make the root free along the entire length, especially at its distal portion in cases with no offending vessels.

Revuelta-Gutierrez et al., in 2006 and 2013, proposed their surgical strategy for TN without vascular compression. The surgery consists of meticulous dissection of the arachnoid, exposing the entire trigeminal root, followed by gentle compression with bipolar tips on its cisternal segment.^{24,50}

This nerve compression technique has been subsequently followed by other authors.^{51,52}

► **Table 9** shows the outcome of a series of intraoperative trigeminal root manipulations.

Internal Neurolysis

Nerve combing is a surgical strategy for trigeminal neuralgia that longitudinally splits branches of the trigeminal nerve using a special fiber knife in accordance with preoperative pain locations and intraoperative findings. This surgical strategy for TN was reported by Li et al. in 1995.⁵³

Ashkan et al., in 2004, reported their series of MVD for TN in 80 patients, "When no compressing vessel was identified, the trigeminal nerve was probed longitudinally by use of a microdissector and split into several fascicles. To the best of our knowledge, this is a novel technique and is preferred by the authors to the partial transverse sectioning of the nerve, which is usually associated with facial numbness."⁵⁴ About 13 patients were subjected to these techniques, without specific mention of the surgical outcome of these patients

Ma et al., in 2009, described 10 patients without vascular compression who underwent nerve combing and achieved satisfactory long-term relief by 70%. They postulated that traumatizing the nerve and dampening its abnormal activity in the brainstem could perhaps explain nerve combing to relieve pain.²⁸

The mechanism by which IN results in pain relief remains unclear. Studies suggest that IN destroys the continuity of axons and decreases the electrical excitability of abnormal afferent trigeminal fibers, permanently degenerating the partial axon tissue and myelin sheath after a period of time.^{29,55}

Internal neurolysis may disrupt interconnections between nerve fibers and communication between distributions of the trigeminal nerve prior to entrance into the brainstem.²⁵ Kline et al. suggested that partial nerve injury and damage to the root of the trigeminal nerve mediated pain relief.⁵⁶

Internal neurolysis resulted in immediate relief of all pain in 80 to 98% of patients (► **Table 10**).

The success rate is durable, with an estimated 72 to 82% pain-free in 5 years.^{25,27,29,55,57}

Microvascular decompression has an initial pain relief rate of 79 to 94%,^{14–16,39} with a pain-free rate of 72 to 84% at a 5-year follow-up.^{47,58–60}

In our series, the two proportion Z-Test showed no statistically significant difference in immediate ($p = 0.289$) and long-term outcomes ($p = 1.000$) between IN and MVD.

All of our patients with IN had some degree of hypoesthesia; in two, the facial numbness was very bothersome in the immediate postoperative period, but they improved afterwards.

In 88% of the cases, sensitivity was fully recovered within 6 months after the operation. Compared with our patients with balloon compression, the degree of numbness is generally much lower in patients with IN at the immediate postoperative. Reported rates of facial numbness after IN range between 60 and 96%.^{25,27,28,61–63} Most cases are transient and resolve over weeks to months.

A previous study reported that facial numbness after IN did not negatively impact the quality of life in the majority of cases when the pain was controlled.⁶²

One case of anesthesia dolorosa reported in the literature includes a patient with previous MVD for the treatment of TN, who presented with a sensory deficit.²⁵

Some studies found that patients with TN without NVC may represent a distinct population of younger, predominantly female patients in their mid-30s with a short symptom duration.^{64,65}

Table 10 Literature series of Internal Neurolysis (nerve combing) for trigeminal neuralgia

Reference	Number of cases	Outcome	Sensory deficit
Ma et al. ²⁸ Clin J Pain 2009	10	Initial pain-free: 80% 70% pain free, recurrence 10% in 3 years	90% transitory numbness 10% permanent numbness
Jie et al. ²⁹ Acta Neurochir 2013	28	82% pain-free, 4% recurrence in 4.3 years	10.5% numbness
Ko et al. ²⁵ JNS 2015	27	Initial pain-free 85%. 72% in 5 years	96% numbness 1 patient (4%) anesthesia dolorosa *
Zhou et al. ⁵⁷ Braz J Otorhinolaryngol 2016	50	Initial satisfactory relief 92% 82% in 7.5 years Recurrence 10%	16% dysesthesias
Zhao et al. ⁵⁵ J Craniofacial 2017	15	Initial pain-free 80% 73.3% pain-free in 4 years	13% transitory numbness 7% permanent numbness
Zhang et al. ⁶¹ World Neurosurgery 2017	86	Initial pain-free 98% 94% in 1 year	60% transitory numbness at 1 day 4% in 1 year
Liang et al. ⁶² Ir J Med Sci 2017	37	Initial pain free 94.6% In 3 years 78% pain free Recurrence 21.6%	91,9% transitory facial numbness In 6 months 67.6% facial numbness
Wu et al. ⁶³ Stereotact Funct Neurosurg 2018	27	Initial pain free 92.6%	88.9% facial numbness
Yang et al. ²⁷ Acta Neurol Belg 2019	34	Initial pain-free 88% In 5 years 73% pain-free Recurrence 6%	76% transitory facial numbness
González-Portillo et al. 2020	50	Initial pain-free 88% Recurrence 6%	100% facial numbness 88% transitory

*Previous MVD, with sensory deficit.

In our patients undergoing IN, the difference in the mean age according to gender was not statistically significant ($p = 0.335$). The age distribution of our female patients shows for the IN procedure an almost uniform dispersion throughout the range of 25 to 75 years old; no outstanding peak was observed. The distribution of the patients in the study of Liang et al. was normal and unimodal. The mean age of the patients was 50.19 years old.⁶²

The relapse of TN after MVD has been shown without recurrence of compression.^{35,66,67}

Internal neurolysis can be an important option in posterior fossa exploration for recurrent or persistent TN symptoms. Zhang et al. reported that, in patients with recurrent or persistent TN symptoms, IN plus MVD significantly improved the success rate of the operation compared with simple redo MVD.⁶¹

Surgical Description of the Grades of Compression and Location in Microvascular Decompression

We suggest a surgical description for posterior fossa exploration for TN. ▶ **Table 11**. It has two components: first, the degree of NVC, based in the studies of Sindou³⁷ and Maarbjerg,³⁸ and the report of Hughes, who concluded that “The only imaging variable that was a statistically significant predictor of being pain-free without medication fol-

Table 11 Surgical description of microvascular decompression for trigeminal neuralgia

MICROVASCULAR DECOMPRESSION GRADING SYSTEM	
GRADES OF COMPRESSION	
<input type="checkbox"/> Grade 0	No contact
<input type="checkbox"/> Grade I	Simple neurovascular contact: without visible alteration of the root
<input type="checkbox"/> Grade II	Severe neurovascular contact: displacement, distortion, indentation, engrooving or atrophy of the trigeminal nerve.
LOCATION OF NEUROVASCULAR COMPRESSION	
<input type="checkbox"/> Anterior Third	
<input type="checkbox"/> Middle Third	
<input type="checkbox"/> Posterior Third	

In this grading system, veins are graded according to the same scale as arteries.

The cisternal segment extends from the root entry point to the porus trigeminus.

Posterior Third: Defined as the proximal third of the cisternal segment of the nerve closest to the root entry point.

lowing MVD was severe NVC.”³⁹ The second component is the location of the MVD. The locations were divided in thirds based on the study of Peker et al.,³² who stated that the cisternal segment of the trigeminal nerve ranged from

8 mm to 15 mm long (mean, 12.3 mm; median, 11.9 mm). The proportion of the central myelin relative to the length of the cisternal portion of the trigeminal nerve was 25% in 74 specimens and between 26 and 33% in 20 specimens out of 100.

Conclusions

Internal neurolysis has become increasingly used as an adjunct or standalone therapy when a low grade or no NVC is identified intraoperatively, or as a salvage procedure in TN patients who are unresponsive to MVD or have a recurrence of symptoms.

Internal neurolysis is a viable treatment option, providing adequate initial pain relief and long-term efficacy.

Patients treated with IN may experience some degree of facial numbness, which was well tolerated by the patients in the present study. However, further studies are required to explore the exact mechanisms of pain relief.

In the present study, there were no statistically significant differences in the immediate and long-term surgical outcomes between IN and MVD.

Conflict of Interests

The authors have no conflict of interests to declare.

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