Consensus practice guidelines on interventions for cervical spine (facet) joint pain from a multispecialty international working group

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ABSTRACT

Background The past two decades have witnessed a surge in the use of cervical spine joint procedures including joint injections, nerve blocks and radiofrequency ablation to treat chronic neck pain, yet many aspects of the procedures remain controversial.

Methods In August 2020, the American Society of Regional Anesthesia and Pain Medicine and the American Academy of Pain Medicine approved and charged the Cervical Joint Working Group to develop neck pain guidelines. Eighteen stakeholder societies were identified, and formal request-for-participation and member nomination letters were sent to those organizations. Participating entities selected panel members and an ad hoc steering committee selected preliminary questions, which were then revised by the full committee. Each question was assigned to a module composed of 4–5 members, who worked with the Subcommittee Lead and the Committee Chairs on preliminary versions, which were sent to the full committee after revisions. We used a modified Delphi method whereby the questions were sent to the committee en bloc and comments were returned in a non-blinded fashion to the Chairs, who incorporated the comments and sent out revised versions until consensus was reached. Before commencing, it was agreed that a recommendation would be noted with >50% agreement among committee members, but a consensus recommendation would require ≥75% agreement.

Results Twenty questions were selected, with 100% consensus achieved in committee on 17 topics. Among participating organizations, 14 of 15 that voted approved or supported the guidelines en bloc, with 14 questions being approved with no dissensions or abstentions. Specific questions addressed included the value of clinical presentation and imaging in selecting patients for procedures, whether conservative treatment should be used before injections, whether imaging is necessary for blocks, diagnostic and prognostic value of medial branch blocks and intra-articular joint injections, the effects of sedation and injectate volume on validity, whether facet blocks have therapeutic value, what the ideal cut-off value is for designating a block as positive, how many blocks should be performed before radiofrequency ablation, the orientation of electrodes, whether larger lesions translate into higher success rates, whether stimulation should be used before radiofrequency ablation, how best to mitigate complication risks, if different standards should be applied to clinical practice and trials, and the indications for repeating radiofrequency ablation.

Conclusions Cervical medial branch radiofrequency ablation may provide benefit to well-selected individuals, with medial branch blocks being more predictive than intra-articular injections. More stringent selection criteria are likely to improve denervation outcomes, but at the expense of false-negatives (ie, lower overall success rate). Clinical trials should be tailored based on objectives, and selection criteria for some may be more stringent than what is ideal in clinical practice.

INTRODUCTION

There are few subjects in interventional pain and spine medicine as controversial as the diagnosis, etiology, and treatment of neck pain. Neck and posterior head pain have a high prevalence rate in both developed and undeveloped regions, being particularly common in the USA, Western Europe, East Asia, Northern Africa, and the Middle East. A systematic review estimated the annual and lifetime prevalence rates to be 37.2% (range 16.7–75.1%), and 48.5% (range 14.2–71%), respectively. According to the Global Burden of Disease 2016 study, spine pain (including neck and low back) is the most common cause of disability in North America and globally for people 25–64 years of age. Age is positively related to the risk of neck pain, obesity is probably unrelated, and women are more likely to experience neck pain. When prevalence is broken down by spine joint or segment, the cited frequency of atlanto–axial (AA) joint pain ranges from as low as 16% to as high as 60% in patients with suspected cervicogenic headaches. Cervical facet (also known as zygapophysial or zygapophyseal) joints are considered to be the primary source of pain in 26–70% of patients with chronic neck pain and 54–60% of neck pain following whiplash injury. The C2–3 and C5–6 joints are the most common clinically implicated levels.
in neck pain, with C2–3, C3–4, and C4–5 being the most radiologically affected. The wide disparity in reported prevalence raises questions regarding the use and accuracy of historical and physical exam signs as non-interventional diagnostic reference standards. The poor correlation between facet joint pathology on imaging and neck pain provokes further debate and disagreements with insurance payers. For diagnostic and/or prognostic criteria, the literature on the ideal patient response for designating a block as ‘positive’ and the optimal number of blocks that should be performed before cervical medial branch radiofrequency ablation (RFA) treatment is contentious and inconsistent, with no consensus emerging.

Cervical spine joint interventions are commonly performed in interventional pain practices, with hundreds of thousands per year being performed in the USA alone. For cervical medial branch RFA, a recent review of the Medicare claims and encounters databases from 2000 to 2018 demonstrated a 112% overall increase in utilization (8.7% annually) over the past 9 years. Along with increased utilization, there was also a reciprocal increase in expenditures on cervical facet interventions of 53% from 2009 to 2018; however, the cost per patient declined 7% over this same time interval (0.8% annual reduction). The utilization of facet interventions is considerably higher than the most commonly cited prevalence rates. Although overall utilization of facet interventions is increasing at a rapid pace, there is a discrepancy in the growth of medial branch blocks (MBB) and intra-articular (IA) joint injections (0.5% annual growth) and cervical medial branch RFA (8.7% annual growth). This disconnect may reflect practice changes such as decreased use of cervical facet IA joint injections, a reduction in the number of diagnostic blocks used before medial branch RFA, or a higher rate of prognostic blocks designated as positive. Increasing utilization alters the risk to benefit ratio of treatments; this, along with inconsistencies in practice and the lack of widely accepted consensus guidelines, has led to increased scrutiny on the part of government regulatory agencies and insurance payers. The Spine Intervention Society (SIS) and the American Society of Interventional Pain Physicians (ASIPP) have published guidelines on the performance of cervical facet blocks and RFA but these rigorous criteria have not been followed in recent randomized controlled and uncontrolled trials (RCTs).

Whereas stringent selection criteria have been associated with high medial branch RFA success rates, the increased false-negative rate that inevitably accompanies strict diagnostic criteria and a host of other factors have resulted in an urgent need for guidelines to inform cervical joint interventions in clinical practice and trials. These factors include the absence of safer and more effective alternatives for neck pain (ie, spinal fusion and chronic opioid therapy were less scrutinized when many of the previous cervical facet studies were published), the publication of few high-quality clinical trials, rising utilization which alters the risk to benefit ratio, and questions surrounding the cost-effectiveness of diagnostic paradigms, which vary from country to country. We aimed to develop pragmatic guidelines that can be used to inform clinical care, improve the quality of research, and assist payers with clinical practice pathways and authorization decisions.

METHODS
The decision to convene a multispecialty and multinational Cervical Joint Working Group to develop atlanto–occipital (AO), atlanto–axial (AA), and cervical facet joint intervention guidelines was approved by the American Academy of Pain Medicine (AAPM) and American Society of Regional Anesthesia and Pain Medicine (ASRA-PM) in August 2020. Fifteen stakeholder academies and societies as well as other organizations (eg, US Departments of Defense and Veteran Affairs) with a vested interest in cervical spine joint interventions were identified, and formal request-for-participation and member nomination letters were sent to those societies who approved involvement in September 2020. A single pain society (US Association for the Study of Pain, USASPS) declined to participate. Organizations were asked to consider a candidate’s expertise, clinical experience, academic interests, and diversity in their nomination process. Each sponsoring society (AAPM and ASRA-PM) nominated two members and participating organizations nominated one member (see online supplemental appendix A for a list of participating academies, societies, and respective representatives). The sole ad hoc member (MSW) had been preliminarily designated to represent USASPF before their Board of Directors declined to participate. For the Departments of Defense and Veterans Affairs representatives, the Chairperson of the Department of Anesthesia at the Uniformed Services University of the Health Sciences and Director of the VA National Pain Management Program nominated individuals.

The Cervical Joint Working Group was charged with preparing guidelines on the use of AO, AA, medial branch and facet joint blocks, and medial branch RFA that spanned the entire spectrum of care to include patient selection, optimizing accuracy, interpreting results, and risk mitigation. Questions and formats were developed by the committee co-chairs (RWH, SPC) based on input from the working group and refined during the initial video-conference call. Guidelines for individual study questions were developed by subcommittees composed of 4–5 members, with one or two persons designated as the ‘leads’ responsible for task delegation. Once a subcommittee came to a consensus on an answer, the working group chairs assisted with editing and formatting, and the section was sent to the entire committee for open-forum comments and revisions. A modified Delphi method was used to tabulate comments, incorporate changes, and converge the answers towards consensus over rounds of teleconference or electronic correspondence. At the initial conference call, the working group decided that >50% panel agreement was sufficient to report a recommendation, but ≥75% agreement was required for consensus, consistent with the Lumbar Facet Intervention Guidelines. After the working group completed the guidelines, the document was sent to participating organizations’ boards of directors for approval, with only minor changes permitted at this stage. For organizational agreement, we determined that consensus required at least ≥75% agreement. At both the committee and organizational levels, dissensions and absences were tabulated for each question.

Search engines used during composition of the various sections included PubMed, Embase, Google Scholar, SCOPUS, and Cochrane Database of Systematic Reviews, in addition to examination of the reference sections of all manuscripts. Additional articles were identified by searching in topically related new journals that are not yet indexed by Science Citation Index or found within PubMed. There were no limitations on language or types of articles used to develop the guidelines, such that experimental studies were considered for the sections on physical examination, anatomy and technical parameters, and case reports were considered for sections pertaining to risk mitigation and complications. Keywords used to address guideline topics were tailored to individual questions and included ‘atlanto–occipital’, ‘atlantooccipital’, ‘atlanto–axial’, ‘atlantaoxial’, ‘cervicogenic’, ‘headache’, ‘facet’, ‘neck pain’, ‘zygapophysial’, ‘zygapophysyal’, ‘radiofrequency’, ‘denervation’, ‘ablation’,
and ‘arthritis’, among others. In accordance with the Lumbar Facet Intervention Guidelines, conclusions for each topic were graded on a scale from A to D, or as insufficient, according to the US Preventative Services Task Force grading of evidence guidelines, with the level of certainty rated as high, moderate, or low (tables 1–3).30

This system, which has been modified for use in interventional pain management guidelines drafted by the ASRA-PM, AAPM, American Society of Anesthesiologists (ASA), ASIPP, and the International Neuromodulation Society (INS), was chosen over others because of its flexibility which permits high-grade recommendations in the absence of high-quality level I studies, which are challenging to conduct for invasive procedures.37

**QUESTION 1: CAN HISTORY AND PHYSICAL EXAMINATION BE USED TO IDENTIFY PAINFUL AO OR AA JOINTS OR TO SELECT PEOPLE FOR PROGNOSTIC BLOCKS?**

AO and AA joint disease may be a source of both neck pain and headache. Accurate diagnosis and management of neck and head pain can be challenging. Pain may be referred from other sources including cervical intervertebral discs, cervical facet joints, ligaments, fascia, and muscles. Detailed history and physical examination can be valuable to help distinguish the etiology of the pain and to target diagnostic and therapeutic injection targets.38 39

**Relevant anatomy of the AO and AA joints**

The AO and AA joints are unique in the cervical spine. The AO and AA joint complexes allow for a significant range of motion (ROM) between the head and mid-cervical spine. The AO and AA joints are innervated by the ventral rami of C1 and C2, respectively.40–42 The AO joint is a synovial articulation between the occipital bone and the first cervical vertebra (the atlas). The joint is formed superiorly by the convex occipital condyle and the occipital bone and the first cervical vertebra (the atlas). The AA joint complex consists of three joints, lateral mass. The AA joint complex is formed superiorly by the convex occipital condyle and the occipital bone and the first cervical vertebra (the atlas). The AA joint complex consists of three joints, lateral mass.

**Table 1** Levels of evidence for guidelines and recommendations

<table>
<thead>
<tr>
<th>Certainty of net benefit</th>
<th>Substantial</th>
<th>Moderate</th>
<th>Small</th>
<th>Zero/negative</th>
</tr>
</thead>
<tbody>
<tr>
<td>High</td>
<td>A</td>
<td>B</td>
<td>C</td>
<td>D</td>
</tr>
<tr>
<td>Moderate</td>
<td>B</td>
<td>B</td>
<td>C</td>
<td>D</td>
</tr>
<tr>
<td>Low</td>
<td>Insufficient</td>
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</tbody>
</table>

One of the main ligamentous structures of the spinal column (anterior longitudinal ligament, posterior longitudinal ligament, ligamentum flavum, interspinous ligament, and supraspinous ligament), the AO joint complex has additional overlying ligaments including the AO ligament, tectorial membrane, apical ligament, and the cruciate ligaments (comprised of the transverse ligament and a superior and inferior band) which provide stability and flexibility, but can also be a pain generator.43 The AA joint complex has additional ligaments as well, namely the anterior and posterior AA ligaments, transverse ligament of the atlas, apical ligament, alar ligaments, and tectorial membrane. These ligaments can become calcified in elderly people, leading to decreased ROM and increased neck pain.44

The AO and AA joints, as described above, provide mechanical strength to stabilize the head, while allowing for complex movements of the cervical spine. Approximately 50% of total cervical spine flexion and extension occurs at the AO articulation.45 46 Over 50% of all cervical spine rotation is provided by the dens of C2 which articulates with C1 and transverse ligaments.45 46 The synovial joints at C1 and C2 rely more on ligamentous stabilizers because they do not have intervertebral discs to provide stabilization.47 48 The weight from the occipital condyles transfers the load from the occiput to the C1 lateral masses and then onto the C2 lateral masses.46

**Referral patterns for pain arising from AO and AA joints**

The diagnosis of pain arising from the AO and AA joints has been less well studied than C2–3 through C7–T1 cervical facet joint pain. As seen in figure 1A,B, pain arising from C1–C2 most often occurs in the suboccipital region, commonly extending cephalad into the head or caudad into the upper neck. Referred pain patterns have been studied in healthy volunteers without neck pain as well as in those with proven cervical joint pain.49–51

Dreyfuss et al.50 studied pain referral patterns in asymptomatic patients from provocative testing of the AO and lateral AA joints via fluoroscopically-guided IA injections. The authors confirmed the nociceptive ability of the AO and AA synovial joints and found that AA injections resulted in consistent referral patterns whereas the AO referral patterns varied significantly.

Referral patterns from asymptomatic patients based on pain provocation are consistent with those from symptomatic patients, based on pain relief after injection. AO-mediated pain has consistently been reported as suboccipital, but may extend process and the posterior surface of the anterior arch of the atlas anteriorly and the transverse ligament posteriorly. In this document, the AA joint refers to the lateral AA joint, unless otherwise specified.

C1 does not have a vertebral body and is not separated from adjacent levels by an intervertebral disc.43 In addition to the five main ligamentous structures of the spinal column (anterior longitudinal ligament, posterior longitudinal ligament, ligamentum flavum, interspinous ligament, and supraspinous ligament), the AO joint complex has additional overlying ligaments including the AO ligament, tectorial membrane, apical ligament, and the cruciate ligaments (comprised of the transverse ligament and a superior and inferior band) which provide stability and flexibility, but can also be a pain generator.43 The AA joint complex has additional ligaments as well, namely the anterior and posterior AA ligaments, transverse ligament of the atlas, apical ligament, alar ligaments, and tectorial membrane. These ligaments can become calcified in elderly people, leading to decreased ROM and increased neck pain.44

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**Table 2** What the grades of evidence mean and suggestions for practice

<table>
<thead>
<tr>
<th>Grade</th>
<th>Definition</th>
<th>Suggestions for practice</th>
</tr>
</thead>
<tbody>
<tr>
<td>A</td>
<td>Our committee recommends this treatment, test or strategy to improve outcomes. There is high certainty that the net benefit is substantial</td>
<td>Offer or provide this service</td>
</tr>
<tr>
<td>B</td>
<td>Our committee recommends this treatment, test or strategy to improve outcomes. There is high certainty that the net benefit is moderate or there is moderate certainty that the net benefit is moderate to substantial</td>
<td>Offer or provide this service</td>
</tr>
<tr>
<td>C</td>
<td>Our committee recommends selectively offering or providing this treatment, test or strategy to improve outcomes to individual patients based on professional judgment and patient preferences. There is at least moderate certainty that the net benefit is small</td>
<td>Offer or provide this service for selected patients depending on individual circumstances</td>
</tr>
<tr>
<td>D</td>
<td>Our committee recommends against the treatment, test, strategy or intervention. There is moderate or high certainty that the service has no net benefit or that the harms outweigh the benefits</td>
<td>Discourage the use of this service</td>
</tr>
<tr>
<td>I</td>
<td>Our committee concludes that the current evidence is insufficient to assess the balance of benefits and harms of the intervention. Evidence is lacking, of poor quality, or conflicting, and the balance of benefits and harms cannot be determined</td>
<td>Read the clinical considerations section of the recommendation. If the treatment or service is offered, patients should understand the uncertainty about the balance of benefits and harms</td>
</tr>
</tbody>
</table>

Special article

Table 3  Levels of certainty regarding net benefit

<table>
<thead>
<tr>
<th>Level of certainty</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>High</td>
<td>The available evidence usually includes consistent results from well-designed, well-conducted studies in representative populations with suspected cervicogenic headache and/or facetogenic pain and/or compelling evidence from non-randomized studies. The studies assess the effects of the treatment, test, or other intervention on treatment or other relevant outcomes. The conclusion is therefore unlikely to be strongly affected by the results of future studies.</td>
</tr>
</tbody>
</table>
| Moderate           | The available evidence is sufficient to determine the effects of the intervention on outcomes, but confidence in the estimate is constrained by such factors as:  
  - The number, size, or quality of individual studies  
  - Inconsistency of findings across individual studies  
  - Limited generalizability of findings to individuals with suspected cervicogenic headache and/or cervical facetogenic pain  
  - High likelihood of bias  
  - Lack of coherence in the chain of evidence  
  - As more information becomes available, the magnitude or direction of the observed effect could change, and that change may be large enough to alter the conclusion. |
| Low                | The available evidence is insufficient to assess effects on treatment and other outcomes of interest. Evidence is insufficient because of:  
  - The limited number or size of studies  
  - Important flaws in study design or methods  
  - Inconsistency of findings across individual studies  
  - Gaps in the chain of evidence  
  - High likelihood of bias  
  - Findings not generalizable to individuals with suspected cervicogenic headache and/or cervical facetogenic pain  
  - Lack of information on important outcome measures  
  - More information may allow estimation of effects on treatment outcomes. |

Note, the levels of certainty described in the definitions for the grading of evidence in table 2 refer solely to the magnitude of benefit attributed to the intervention, while the levels of certainty in table 3 consider the basis of evidence for the recommendation and the likelihood the recommendation will be affected by future studies. Whereas the two are related, the grading of evidence and rating of certainty were considered separately.

to the frontal area, slightly anterior to the vertex. The referral zone approaches—but does not include—the ear in most cases. Other patterns that have been described are isolated suboccipital pain, suboccipital and supraorbital pain, and rarely the entire hemicranium.

The spinal nucleus of the trigeminal nerve extends caudally to the dorsal horn of the first 3–4 cervical spinal nerves. The trigeminal nerve and the upper three cervical nerves provide afferent fibers to the trigeminocervical nucleus, which may account for the overlapping pain patterns described in AO and AA joint pain which include upper neck pain that spreads to the oculofrontotemporal area.

The pain referral patterns of the AA joint reported by Dreyfuss et al. are consistent with prior studies. Pain emanating from the AA joint was described as discrete unilateral pain at the occipito–cervical junction, retro–mastoid area, and in the upper cervical region. This is in contrast to pain from the AO joint reported in the same study which tended to radiate more cephalad towards the vertex of the head, and occasionally into the temporal and posterior auricular areas. Cooper et al. reported that AA pain often encompassed the region of the posterior ear and orbit. It sometimes encompassed the ear itself and was rarely experienced in the temporoparietal area. The pain quality has been described as ‘deep’, ‘boring’, and ‘aching’. Patients with AA joint pain often report occipital headaches, suboccipital neuralgia, and sometimes pain radiating to the shoulder. Radicular pain or a history suggestive of myelopathy is an uncommon finding; however, these have been reported in rare cases of C1–C2 pseudoarticulation.

Figure 1  Posterior (A) and lateral (B) segmental maps showing the typical pain referral patterns of the atlanto–occipital (C0–C1, blue) and atlanto–axial (C1–2, red) joints. Striped areas (blue/red hash marks) represent overlapping atlanto–occipital and atlanto–axial pain maps.
nerves including the AO joint, median AA joint, C2–3 disc, and from any structure innervated by the upper three cervical spinal regions. This low PPV may be explained by the fact that cervicogenic headache can be referred pain.67 Paradoxically, greater AA joint ventral subluxation was associated with less pain. Alternative imaging including bone window CT views of the AA joint, MRI, or cervical myelography may be needed to rule out concomitant alternative cervical spine pathology.68

**Recommendations**

In summary, there are no pathognomonic historical signs or physical examinations that can reliably predict response to AA or AO joint blocks in individuals with chronic neck pain. AA and AO joint pain typically manifest in the C1, C2, or trigeminal nerve process of the axis and the ring formed by the transverse ligament of the atlas and the anterior arch. Common historical and physical examination findings of AA dysfunction include limited ROM during rotation as well as flexion and extension depending on the extent of tectorial membrane impairment. In more advanced cases, examination signs can include severely restricted rotation and lateral flexion of the cervical spine to the affected side.63 crepitus, prominent tenderness at the occipito–cervical junction, cranio cervical kyphosis, and torticollis.64 The presence of gait abnormalities, radicular symptoms, and audio-visual symptoms are unlikely to be related to isolated AO or AA dysfunction.

Although the AO and AA joints can be visualized on imaging, including plain radiographs, imaging cannot confirm the origin of pain. Normal imaging does not rule out arthropathy and radiographic joint abnormalities are incapable of identifying a painful joint. For example, in a study involving 400 patients with rheumatoid arthritis, 45.8% had radiographic evidence of AA involvement but only 45.4% of those individuals had neck pain.65 Paradoxically, greater AA joint ventral subluxation was associated with less pain. Alternative imaging including bone window CT views of the AA joint, MRI, or cervical myelography may be needed to rule out concomitant alternative cervical spine pathology.68

**Table 4**  
Studies evaluating pain referral patterns for atlanto–occipital and atlanto–axial injections

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patients</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Busch and Wilson, 1989</td>
<td>25 pts with head and neck pain of which 2 pts received LA and steroid AO and AA joints injections</td>
<td>Case series/retrospective</td>
<td>Pain relief in both cases ranged from 3 weeks to 4 months (one patient received serial injections). AA and AO joints may be a source of occipital headache refractory to conservative therapy</td>
<td>25 pts treated but only 2 cases described</td>
</tr>
<tr>
<td>Aprill et al 2002</td>
<td>34 pts with history of occipital or SO pain, tenderness over C1 and decreased ROM of AA joint treated with IA injection of LA and steroid</td>
<td>Prospective observational</td>
<td>21 of 34 (62%) pts experienced complete relief (≤1/10 on VAS) for 2 hours following the injection</td>
<td>No control group, no sham injection or other joints examined</td>
</tr>
<tr>
<td>Narouze et al 2007</td>
<td>32 pts with clinical exam consistent with AA-mediated pain treated with AA IA injection of LA and steroid</td>
<td>Retrospective study</td>
<td>15 of 32 (46.8%) pts obtained complete pain relief (NRS pain score=0), 26/32 (81.2%) pts with ≥50% pain relief</td>
<td>No control or comparison groups or other joints examined</td>
</tr>
<tr>
<td>Lee et al 2015</td>
<td>24 pts with headache and/or SO pain, SO tenderness, and limited range of lateral bending with rotation at the AO joint, treated with IA injection of LA. Responders received IA LA and steroid injection</td>
<td>Prospective observational</td>
<td>20 of 24 (83%) pts had ≥50% pain relief for 30 min following the injection. 18 of 20 (90%) pts had a 2-point or greater reduction in pain score</td>
<td>No control or sham groups, 2 month follow-up. 14 of 14 pts with headache had ≥50% relief. 15 of 20 pts with posterior neck pain had ≥50% relief.13 of 17 pts with shoulder/arm pain had ≥50% relief</td>
</tr>
<tr>
<td>Dreyfuss et al 1994</td>
<td>5 asymptomatic volunteers (no history of headache or neck pain) received one AO and and AA provocative injection with contrast</td>
<td>Prospective observational</td>
<td>AO injections provoked variable referred pain from the level of the C5 spinal segment through the vertex of the head. AA injections were more consistent, producing pain only in the SO region</td>
<td>Provoked pain described as dull, deep ache, or heavy pressure</td>
</tr>
<tr>
<td>Cooper et al 2007</td>
<td>5 pts with neck or SO headache treated with AA injection of LA (one patient had bilateral injections)</td>
<td>Prospective observational</td>
<td>Responders had 100% pain relief or relief in a definable portion of the patient’s area of pain. 5 out of 134 (3.7%) total pts with positive cervical diagnostic blocks had C1–2 as a source</td>
<td>AA injections relieved pain in the neck up to the vertex of the head, occasionally in the region of the ear and orbit</td>
</tr>
<tr>
<td>Fukui et al 1996</td>
<td>10 pts with neck pain were treated with AO and 10 pts with neck pain were treated with AA IA injections with LA and steroid</td>
<td>Prospective observational</td>
<td>AO and AA injections produced pain in the posterior occiput and posterolateral cervical spine (areas over the mastoid process)</td>
<td>Pain relief was not reported, although LA and steroid was used in the provocative injection</td>
</tr>
<tr>
<td>Ehni and Benner, 1984</td>
<td>7 pts with SO pain, tenderness and pain during rotation were treated with AA IA injection of LA and steroid</td>
<td>Case series</td>
<td>AA injections produced immediate relief</td>
<td>No details provided regarding injection technique. No quantification of pain relief was provided</td>
</tr>
</tbody>
</table>

AA, atlanto-axial (C1–C2) joint; AO, atlanto-occipital (C0–C1) joint; IA, intra-articular; LA, local anesthetic; MBB, medial branch blocks; NRS, numerical rating scale; pts, patients; ROM, range of motion; SO, suboccipital.

Historical features suggestive of lateral AA joint pain include occipital or suboccipital pain, focal tenderness over the suboccipital area, focal tenderness over the transverse process of C1, and pain provoked by active or passive rotation of C1 on C2.62 Using these features, Narouze et al62 treated 32 patients who were screened from a total of 115 patients referred with cervicogenic headache. Only 15 of those 32 patients experienced complete pain relief following an IA block, thereby confirming the presumptive pain generator with a diagnostic IA block, especially before considering surgical options. Based on a cohort study involving 34 patients in which 21 responded to lateral AA joint injections, Aprill et al63 concluded that the only way to confirm whether a joint is painful is by anesthetizing the joint. They found that history in conjunction with physical examination has a PPV of only 60% for pain stemming from the AA joint, meaning that without diagnostic blocks a substantial proportion of patients will be misdiagnosed (table 4).

**Physical examination of the neck to diagnose AO and AA pain**

The AA joint complex accounts for 60% of cervical rotational movement.64 The pivot articulation occurs between the odontoid process of the axis and the ring formed by the transverse ligament of the atlas and the anterior arch. Common historical and physical examination findings of AA dysfunction include limited ROM during rotation as well as flexion and extension depending on the extent of tectorial membrane impairment. In more advanced cases, examination signs can include severely restricted rotation and lateral flexion of the cervical spine to the affected side.63 crepitus, prominent tenderness at the occipito-cervical junction, cranio cervical kyphosis, and torticollis.64 The presence of gait abnormalities, radicular symptoms, and audio-visual symptoms are unlikely to be related to isolated AO or AA dysfunction.
distribution, with AA pain having more reproducible and consistent symptoms than AO joint pain. We conclude that history and physical examination cannot reliably identify painful AO or AA joints, but can guide injection decisions which could confirm the AO and AA joints as pain generators: grade C recommendation, low level of certainty.

**QUESTION 2: CAN HISTORY AND PHYSICAL EXAMINATION BE USED TO IDENTIFY A PAINFUL FACET JOINT, OR TO SELECT PEOPLE FOR PROGNOSTIC BLOCKS?**

Cervical facet joints are proposed as the primary source of pain in 25–67% of patients with chronic neck pain. The C2–3 and C5–6 joints are the most common clinically implicated in neck pain, while C2–3, C3–4, and C4–5 joints are the most likely to display radiological features of degeneration; injury to the neck increases the probability of the facet joints being the source of chronic neck pain (see below for whiplash). Frequently used criteria for considering patients for prognostic blocks include neck pain of moderate-severe intensity (score ≥4 out of 10 on a pain intensity scale) radiating to the head, shoulder, or upper arm for at least 6 weeks in the absence of focal neurological findings. The targeted facet joints are usually decided on based on patient report and tenderness on examination, sometimes performed under fluoroscopy. In patients who are postsurgical, adjacent segments are often affected after arthrodesis, and treated levels bear more force during extension following disc arthroplasty. Factors that decrease the probability of the cervical facet joints being pain generators include prominent pathology of other likely sources of neck pain such as markedly degenerated or herniated intervertebral discs and symptomatic spinal stenosis. In a study by Cohen et al that evaluated historical and physical examination features associated with successful cervical medial branch RFA outcomes, pain radiating to the occiput and a history of headaches increased the probability of treatment failure. In this study, paraspinal tenderness, but not neck pain worsened by facet loading (extension and rotation of the cervical spine), was highly predictive of a successful outcome, with two-thirds of patients with paraspinal tenderness reporting a successful outcome versus the same proportion of those without tenderness failing RFA.

Importance of whiplash as a cause of cervical facet joint pain

Mechanical injury is often an initiating factor for cervical facet joint pain. The cervical facet joints were identified as a source of neck pain in 54% (95% CI 40% to 68%) of individuals (as confirmed by dual diagnostic MBB) in a study performed in patients with chronic whiplash-associated disorders (WAD). However, the incidence of cervical facet joint pathology contributing to neck pain as calculated based on patients who completed this study was 71%. WAD represent a spectrum of symptoms arising from an initial whiplash injury that usually occurs as a result of a motor vehicle collision (MVC). Smith and colleagues performed dual diagnostic MBB. They reported the most common cervical facet joints associated with neck pain were C2–3 (36%), followed by C5–6 (35%), and C6–7 (17%). Joints at C1–2, C3–4, C4–5, and C5–6, and were each symptomatic in less than 5% of cases. Among patients with cervical facet joint pain, 52% had only 1 symptomatic joint. In the remainder, multiple symptomatic joints occurred in various combinations. These included bilateral joints at the same segment (eg, C2–3 or C5–6), adjacent joints on the same side (eg, C5–6, C6–7), and non-adjacent joints on the same side (C2–3 and C5–6). When C3–4 and C4–5 facet joints were symptomatic, it was usually in combination with an adjacent joint (table 5).

**C2–3 facet joint**

The C2–3 joint pathology is considered a valid etiology of cervicogenic headaches. The C2–3 facet joint is innervated by the third occipital nerve (TON); hence, headaches arising from C2–3 pathology were called third occipital headache in the past. Lord and colleagues evaluated the prevalence of TON headache among 100 patients with neck pain for at least 3 months following whiplash using history, physical examination, and diagnostic cervical MBB with local anesthetics (LA). The prevalence of TON headache in this cohort was 27% (95% CI 18% to 36%) and as high as 53% (95% CI 37% to 68%) among those with headache as the dominant symptom. No unique features on history or examination correlated with positive MBB results. Patients with a positive block were significantly more likely to be tender over the C2–3 facet joint, with a sensitivity of 85%.

Referral patterns of pain for pain arising from cervical facet joints

The referral patterns of pain arising from the cervical facet joints have been evaluated in volunteers and in patients with pain proven to arise from the cervical facet joints. Dwyer et al performed IA facet joint injections in four volunteers and one patient with neck pain to map the area of pain produced by injection into each joint (figure 2A,B). Stimulation of the C2–3 joint by capsular distension was associated with upper neck pain that extended into the head (often towards the ear, vertex, forehead, or eye). Stimulation of the C5–6 joint resulted in pain radiating into the lower neck, top of the scapula, and shoulder above the level of the scapular spine that was distinguishable from pain extending caudally to the scapular spine from irritation of the C6–7 joint. Injections into the C3–4 joint resulted in pain in the neck extending from the suboccipital region to the lower neck without involving the shoulder, whereas injection into the C4–5 joint caused pain that was more caudal, in the top of the shoulder and lower part of the neck.

Cooper et al conducted a study in 194 patients with neck pain who received dual LA diagnostic MBB. They reported the most common cervical facet joints associated with neck pain were C2–3 (36%), followed by C5–6 (35%), and C6–7 (17%). Joints at C1–2, C3–4, C4–5, and C5–6, and were each symptomatic in less than 5% of cases. Among patients with cervical facet joint pain, 52% had only 1 symptomatic joint. In the remainder, multiple symptomatic joints occurred in various combinations. These included bilateral joints at the same segment (eg, C2–3 or C5–6), adjacent joints on the same side (eg, C5–6, C6–7), and non-adjacent joints on the same side (C2–3 and C5–6). When C3–4 and C4–5 facet joints were symptomatic, it was usually in combination with an adjacent joint (table 5).
Physical examination of the neck to diagnose facetogenic pain

Physical examination of the neck was found to have a high sensitivity but low specificity in a study in which 77% of subjects were identified as having primarily facet joint pain. However, other studies have suggested that specific physical examination maneuvers can identify cervical facet joints as the primary cause of neck pain (Box 1).

In a study involving 125 patients who received dual LA diagnostic cervical MBB, a protocol consisting of manual spinal examination, palpation for segmental tenderness, and extension-rotation testing was found to have a specificity of 84% (95% CI 77% to 90%) and a positive likelihood ratio of 4.94 (95% CI 2.8 to 8.2) for identifying cervical facet joints as the principal source of neck pain. Table 5 summarizes the evidence for features on history and physical examination suggestive of cervical facetogenic pain.

Recommendations

In summary, there are no single pathognomonic historical symptoms or physical examination signs that can reliably predict the response to facet joint blocks in individuals with chronic neck pain, although a history of whiplash and the presence of paraspinal tenderness in the muscles overlying the facet joints appear to be associated with a positive response to facet joint interventions. Maneuvers associated with radicular signs may be predictive of negative diagnostic cervical MBB. There does not appear to be a difference between the psychological profiles of patients who respond and those who do not respond to interventions targeting the innervation to the cervical facet joints. When selecting targets for blocks, levels should be determined based on clinical presentation (tenderness on palpation [preferably performed under fluoroscopy], pain referral patterns); grade C recommendation, low level of certainty.

QUESTION 3: IS THERE ANY CORRELATION BETWEEN RADIOLOGICAL FINDINGS AND PROGNOSTIC BLOCK OR RFA OUTCOMES?

Radiological findings and painful facet joints

In order to correlate radiological findings with a painful facet joint, or outcomes of prognostic IA blocks, MBB or RFA, radiological findings must be compared with patient-reported pain outcomes. Degenerative changes noted in radiological studies may not always be symptomatic, and the presence of findings does not always correlate with clinical symptoms.

Plain film radiographic examinations of the cervical spine represent a simple imaging modality for the evaluation of spine pathology. However, research to date has not found a strong association between the presence of cervical spondylosis on x-rays and clinical pain symptoms. Heller et al described a retrospective case–control study in 653 patients referred for x-ray examination of the cervical spine for neck pain compared with 365 asymptomatic patients referred for barium studies. No significant differences were noted in the presence of cervical spondylosis between groups, and there were also no significant associations between pain in the arm, shoulder, scalp, neck, and back of the head, and neck stiffness with pathologic x-ray findings. Similar findings have been noted in other retrospective cohort studies, with a lack of association between longitudinal plain film changes and the presence or severity of pain 10 years after the onset of neck pain. The lack of association between facet joint osteoarthritis on cervical spine radiographs with reports of neck pain has been reported in larger population studies of women and men aged 20–65 years. More recently, a retrospective cohort study confirmed these earlier findings of the lack of association between facet or uncinate process hypertrophy and pain intensity, headaches, referred shoulder/hand pain, radiculopathy, or numbness. More high-quality prospective research is needed to understand the relationship between cervical spine x-ray findings and facet-mediated pain.

CT represents a more sensitive imaging modality for the assessment of cervical facet pathology and may yield abnormalities in asymptomatic individuals, with one study finding a 33% prevalence of cervical facet arthritis in patients who underwent CT scans for non-spinal pain. Morishita et al performed a retrospective study in 215 patients with cervical spine degenerative disease. Although the authors reported a significant association between hypertrophic changes on CT studies and the presence of neck pain, the statistical analysis was flawed in that it failed to control for important covariates such as age and gender known to affect the prevalence of facet degenerative changes and neck pain. Similar cross-sectional studies with limited numbers of patients report a weak association, but the lack of statistical power in describing small cohorts of patients represents a serious limitation.
The prognostic value of this imaging modality. Given that CT of asymptomatic cervical facet osteoarthritis (33%) decreases limitation. CT can demonstrate osteophytes and hyperostosis, but not changes in articular cartilage, which presents limitations in identifying painful facet joints. The high prevalence of asymptomatic cervical facet osteoarthritis (33%) decreases the prognostic value of this imaging modality. Given that CT evidence of cervical facet arthrosis is common among older patients with neck pain at the C2–6 levels, additional imaging techniques may need to be incorporated to differentiate the characteristics of painful cervical facet joints from those that are asymptomatic. At present, the limited research that has examined the association of CT findings with cervical facetogenic pain is inconclusive.

### Table 5: Studies examining history (including referral patterns) and physical examination signs for patients with cervical facetogenic pain

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patients</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
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<tbody>
<tr>
<td>Dwyer et al 1990</td>
<td>4 asymptomatic volunteers and 1 patient with neck pain whose cervical facet joint capsules were ’stimulated’ using 1 mL IA contrast</td>
<td>Prospective cohort study</td>
<td>Pain referral maps produced for C2–3 (lower head, upper neck), C3–4 (upper neck), C4–5 (well localized to mid-neck below C3–4), C5–6 (top of scapula and shoulder above the scapular spine) and C6–7 (lower neck to inferior angle of scapula) joints</td>
<td>Pain produced by injection in 9 out of 11 joints</td>
</tr>
<tr>
<td>Aprill et al 1991</td>
<td>10 pts with neck pain received MBB with LA and steroid</td>
<td>Prospective cohort study</td>
<td>Concordance between painful joint level(s) predicted based on clinical evaluation and response to diagnostic blocks</td>
<td>4 pts had undergone anterior cervical fusions. 3 pts had negative discography results for cervical discogenic pain</td>
</tr>
<tr>
<td>Barnsley and Bogduk, 1993</td>
<td>16 pts with chronic neck pain, with or without referred pain in the head or shoulder after MVC, received controlled MBB with LA</td>
<td>Prospective study</td>
<td>11 of 16 pts had complete relief of neck pain with restoration of neck movements after cervical MBB; 4 of the remaining 5 pts had a positive cervical MBB at non-predicted levels</td>
<td>No control group. Levels for cervical MBB chosen based on pain maps and sites of maximal tenderness. No patient had radiculopathy. Normal imaging studies. The 25 MBB performed were highly specific</td>
</tr>
<tr>
<td>Lord et al 1994</td>
<td>100 pts with chronic neck pain after whiplash received double diagnostic MBB with LA</td>
<td>Prospective study</td>
<td>C2–3 joint was responsible for headaches in 27% of pts confirmed by diagnostic TON block. Tenderness over C2–3 joint on examination predicted positive block</td>
<td>No control group. C2–3 joint responsible for headaches in 53% of pts when headache was main symptom</td>
</tr>
<tr>
<td>Lord et al 1996</td>
<td>24 pts with chronic neck pain after MVC with Quebec task Force WAD grade I–IV selected by double diagnostic MBB with LA and placebo injection who underwent medial branch RFA</td>
<td>Prospective RCT</td>
<td>44% of screened pts had headache and neck pain from cervical facet joints</td>
<td>Sham medial branch RFA group included C2–3 facet joint pain in 33% of pts</td>
</tr>
<tr>
<td>Fukui et al 1996</td>
<td>61 pts with neck pain from the cervical facet joints confirmed by IA capsular stimulation or electrical stimulation of dorsal rami C3–7</td>
<td>Prospective cohort study</td>
<td>Pain region and source (joint and/or DR): Occipital cervical region: C2–3 and C3 DR Upper posterolateral cervical region: C1–2, and C2–3 Upper posterior cervical region: C2–3, C3–4, and C3 DR Middle posterior cervical region: C3–4, C4–5, and C4 DR Lower posterior cervical region: C4–5, C5–6, and C5 DR Supracervical region: C4–5, C5–6, and C4 DR Superior angle of scapula: C6–7, C6, and C7 DR Mid–scapular region: C7/T1 and C7 DR</td>
<td>Internal controls were asymptomatic joints. 100% sensitivity and specificity of physical examination to predict block response. Incidence of cervical facet joints as the cause of neck pain was 75%</td>
</tr>
<tr>
<td>Jull et al 1998</td>
<td>20 pts with neck pain who had complete pain relief with dual MBB. Assessed the diagnostic accuracy of physical examination</td>
<td>Observational study</td>
<td>15 of 15 (100%) pts with cervical MBB-proven facet joint pain (and no CMBB-negative pts) were correctly identified based on physical examination. The correct segmental level was identified in all pts</td>
<td>Pain patterns of adjacent segments overlapped</td>
</tr>
<tr>
<td>Cooper et al 2007</td>
<td>194 pts with neck pain who underwent dual comparative MBB</td>
<td>Prospective observational study</td>
<td>Segmental patterns of pain arising from cervical facet joints identified: Suboccipital: C1–2, C2–3 Posterior lateral neck: C3–4 Neck to shoulder girdle: C4–5 Lower neck to upper limb girdle: C5–6, C6–7</td>
<td>Internal controls were asymptomatic joints. 100% sensitivity and specificity of physical examination to predict block response. Incidence of cervical facet joints as the cause of neck pain was 75%</td>
</tr>
<tr>
<td>Cohen et al 2007</td>
<td>92 pts who underwent cervical medial branch RFA</td>
<td>Retrospective study to determine factors associated with successful MBB</td>
<td>Paraspinal tenderness associated with successful outcome</td>
<td>Radiation of pain to head, opioid use, and pain exacerbated by neck extension and/or rotation associated with failure</td>
</tr>
<tr>
<td>King et al 2007</td>
<td>173 pts with suspected cervical facet joint pain based on physical examination studied with MBB</td>
<td>Observational study</td>
<td>Physical examination lacked validity, refuting results of a previous study with overlapping authors. Examination had a high sensitivity (88%) but low specificity (39%)</td>
<td>Pts with previous cervical spine surgery and those with negative physical examination signs were excluded</td>
</tr>
<tr>
<td>Smith et al 2013</td>
<td>90 subjects with WAD &gt;6 months duration post-MVC who received IA injections and MBB; 30 healthy controls</td>
<td>Cross-sectional design comparing physical and psychological examination in responders and non-responders with WAD to control pts</td>
<td>58 of 90 (84%) achieved at least 50% pain relief with IA or MBB. No difference in objective sensory testing, muscle activity or ROM between facet block responders and non-responders, but all were abnormal compared with controls. Facet non-responders had greater medication use and catastrophizing scores compared with responders</td>
<td>Large proportion of participants were lost to follow-up</td>
</tr>
<tr>
<td>Schneider et al 2014</td>
<td>125 pts with neck pain in whom a clinical examination protocol was validated against positive dual cervical MBB outcome (80% reduction of pain)</td>
<td>Prospective cohort study</td>
<td>A protocol consisting of MSE, PST, and ER test had a specificity of 84% (95% CI 77% to 90%) and a positive likelihood ratio of 4.94 (95% CI 2.8 to 8.2) for cervical facet joints being the source of neck pain</td>
<td>Sensitivity of PST and MBB were 94% (95% CI 90% to 98%) and 92% (95% CI 88% to 97%), respectively. Any single test was insufficient for diagnosis</td>
</tr>
</tbody>
</table>

**Notes:**
- DR, dorsal ramus; ER, extension rotation; ITT, intention to treat; LA, local anesthetic; LR, likelihood ratio; MBB, medial branch block; MSE, manual spinal examination; MVC, motor vehicle collision; PP, per protocol; PST, palpation for segmental tenderness; pts, patients; QTF, Quebec Task Force; RFA, radiofrequency ablation; ROM, range of motion; TON, third occipital nerve; WAD, whiplash associated disorders.
Box 1  Proposed protocols for identifying painful cervical facet joints

Cervical spine range of motion (ROM)
Measurements of cervical ROM for flexion and extension in the sagittal plane, left- and right-sided lateral flexion, and rotation are taken with the patient seated. The patient is asked to report any pain response and these responses are categorized as increased, decreased, or resulting in no change in baseline cervical spine pain.

Extension-rotation (ER) testing
Patients are seated and asked to fully extend their head, followed by rotation to both sides. Subjects report any pain at the end of motion. A positive test for pain arising from the cervical facet joints is provocation of baseline cervical spine pain.

Manual spinal examination (MSE)
The patient is positioned prone with the cervical spine in a neutral position. The assessor applies a posterioranterior directed force over the articular pillars from C2–3 to C6–7 on each side. The subject reports any pain provocation, whereby a positive test is defined as worsening baseline or referred pain when the assessor perceives moderate or marked resistance to motion.

Palpation for segmental tenderness (PST)
PST is performed with the subject in the prone position. The assessor palpates the segmental muscles overlying the facet joints (C2–3 to C6–7) bilaterally. These muscles have the same nerve supply as the painful joint(s) and elicit tenderness and spasm. The test is considered positive if the patient reports an increase in baseline pain, either localized or referred. Paraspinal tenderness was reported to be predictive of a positive response to cervical medial branch RFA in one study.20

MRI represents an imaging modality that can identify the presence of edema in a degenerated facet joint. In a retrospective study composed of 173 patients, Nevalainen et al15 found a significant correlation between the presence (vs absence) of neck pain and the presence of ipsilateral cervical facet bone marrow edema. However, the severity of neck pain did not significantly increase with the severity of bone marrow edema, raising questions regarding the utility of this finding for characterizing facet-mediated pain severity. Future research to confirm the presence of facet-mediated pain through prognostic blocks would build on these study findings.

Radionuclide bone scintigraphy with single-photon emission computed tomography (SPECT) provides functional imaging to assess microcalcification resulting from increased osteoblastic activity. This increased activity may reflect areas of mechanical stress and degenerative changes. SPECT alone as a diagnostic tool is limited by imprecise localization of affected spinal segments and low spatial resolution. The SPECT/CT modality combines the high sensitivity of SPECT with the anatomic localization of CT.89 The addition of CT corrects for soft tissue attenuation, thereby improving scan sensitivity. CT also increases specificity by demonstrating structural pathology that is causing increased tracer activity. Two small retrospective studies have examined the association between SPECT/CT findings and outcomes of cervical facet joint blocks. Neither study found a correlation, and each noted a large discrepancy between facet joint SPECT activity and the location of the cervical facet joint injection or MBB.29

There are more robust data investigating the use of SPECT to identify levels in the lumbar spine. Moderate evidence supports the use of SPECT for the identification of painful lumbar facet joints prior to MBB, and weak evidence supports the use of SPECT to identify painful lumbar facet joints prior to IA joint injections.29 Future research extending into these combined imaging modalities may elucidate a connection between radiological findings and facet-mediated pain (tables 6–9).

Radionuclide bone scintigraphy with positron emission tomography (PET) provides real-time information on abnormal biological processes. It can demonstrate foci of hypermetabolism in several inflammatory and infectious disease processes. Intense F-fluorodeoxyglucose (F-FDG) activity has been noted in regions of facet joint arthropathy.91 Combining F-FDG PET with MRI allows for further accurate anatomic localization of metabolic information demonstrated through PET. Benefits of this technique over F-FDG PET/CT include lower radiation exposure. In a small case–control study performed in 10 patients with clinically diagnosed cervical facet syndrome, F-FDG PET/MRI localized CT-guided MBB resulted in significantly greater pain relief for up to 3 months compared with landmark-guided injections in patients with negative PET/MRI.92 However, the MBB technique was non-standard due

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<tr>
<td>Rudy et al 201520</td>
<td>322 pts with neck stiffness, shoulder pain, arm pain and/or headache attending chiropractic teaching clinics with cervical radiographs</td>
<td>Retrospective cross-sectional study</td>
<td>Symptoms of neck pain, headaches, referral of pain to the shoulder, and upper extremity radiculopathy did not correlate with cervical facet joint degeneration</td>
<td>Convenience sample with no asymptomatic comparison group. Small association between neck stiffness and facet hypertrophy</td>
</tr>
<tr>
<td>Van der Donk et al 199117</td>
<td>5440 volunteers 20–65 years of age enrolled in a national survey study stratified by presence of neck pain</td>
<td>Cross-sectional survey study</td>
<td>Osteoarthritis of the facet joints noted on cervical radiographs was not associated with neck pain</td>
<td>Mean age of participants was 46 years. Age was positively associated with neck pain</td>
</tr>
<tr>
<td>Gore et al 198692</td>
<td>205 pts with neck pain &gt;10 years duration</td>
<td>Retrospective</td>
<td>Presence or severity of pain was not associated with the presence of degenerative changes including facet arthropathy, sagittal diameter of the spinal canal, or degree of cervical lordosis on initial or final cervical x-ray</td>
<td>68 of 205 (33%) were involved in litigation. Mean age at onset of neck pain was 43 years; mean age at final evaluation was 58 years.</td>
</tr>
<tr>
<td>Heller et al 198645</td>
<td>653 case pts were referred for cervical spine x-ray. 365 control pts who were referred for barium studies received cervical spine x-rays</td>
<td>Retrospective case–control</td>
<td>No significant difference in the presence of cervical spondylisis between groups. There were no significant associations between neck, arm, or occipital pain, and neck stiffness with x-ray findings</td>
<td>Cervical spondylisis was positively correlated with age</td>
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to its high volume (3 mL) and inclusion of steroids (dose unmentioned). To date, conventional MRI, plain CT, dynamic flexion films, and radionuclide bone scanning have not demonstrated reliable diagnostic utility for identifying suspected cervical facet-mediated pain generators.39

**Radiological findings associated with whiplash injury**

Imaging findings immediately after whiplash illustrate the extent of injury to cervical facet joints. In a prospective study by Rydman et al40 conducted in 121 patients presenting to the ED after MVC who underwent cervical CT scans within 10 days of admission, the authors found that mean pain intensity 6 months after MVC was significantly associated with baseline CT findings of facet joint degeneration. Overall, the prevalence of cervical facet joint degeneration was 45.5%, and those patients with a moderate degree of facet joint degeneration were significantly more likely (OR 6.7, 95% CI 1.9 to 24.3) to self-report absence of recovery at 6 months. Facet joint degeneration on CT was graded by the presence of joint space narrowing, osteophytes, and irregularities of the articular surface. However, any specific correlation between the affected facet joints on CT and the suspected levels of pain was not analysed. In a longitudinal study by Daimon et al41 comparing MRIs of the cervical spine obtained 2 weeks and 20 years after a whiplash injury, changes in clinical symptoms (eg, neck pain, shoulder stiffness, dizziness, and tinnitus) were not associated with the progression of degenerative changes on MRI. In another study by Gore et al83 the presence or severity of neck pain was not related to the presence of degenerative changes on radiographs. However, postmortem studies performed in victims of fatal MVCs have identified lesions and small fractures undetectable on plain radiographs, which raises the possibility that more sensitive radiological studies may also fail to detect clinically significant injuries.71,95 More data are needed to understand the link between radiological findings and pain after a whiplash injury.

**Radiological findings and outcomes after prognostic blocks or RFA**

The association of radiological findings with outcomes of diagnostic cervical facet joint blocks has rarely been examined. Among 37 patients presenting for single, unilateral or bilateral, one-level CT-guided cervical facet joint blocks, no significant difference in pain relief was noted based on the grading of cervical facet osteoarthritis.96 In a prospective observational study conducted in 121 patients referred for CT-guided cervical IA facet injections with steroid, a greater proportion of patients referred based on pain palpation compared with imaging (CT or MRI) reported improvement for up to 1 month.97 Minimal research has examined the association between radiological findings and RFA outcomes. Cohen et al20 performed a retrospective study evaluating factors associated with outcomes in 92 patients who underwent cervical medial branch RFA after positive diagnostic blocks. Although facet pathology was found

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**Table 7** Studies evaluating the association between CT imaging pathology and facet pain

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<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
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<tbody>
<tr>
<td>Tiwari et al 2020</td>
<td>10 pts were referred for cervical spine CT for reasons unrelated to spinal pain</td>
<td>Cross-sectional</td>
<td>Facet joint arthritis on CT was negatively associated with patient-reported neck pain</td>
<td>Subgroup analysis of only 10 pts</td>
</tr>
<tr>
<td>Kim et al 2019</td>
<td>50 pts who received CT scans for non-spinal pathologies. Pts with neck pain were excluded</td>
<td>Retrospective</td>
<td>33.4% prevalence of asymptomatic cervical facet arthritis</td>
<td>C5–7 joint was most likely to demonstrate arthritic changes with findings more common among older (≥40 years) pts</td>
</tr>
<tr>
<td>Rydman et al 2019</td>
<td>121 pts present to the emergency department for neck pain after MVC with a cervical CT scan performed at admission</td>
<td>Prospective longitudinal</td>
<td>Moderate facet joint degeneration, but not disc degeneration, was associated with persistent pain after 6 months</td>
<td>Regions of mild and severe facet degeneration were not associated with recovery</td>
</tr>
<tr>
<td>Le Clec'h et al 2016</td>
<td>121 pts who underwent cervical IA facet injections based on MRI or CT imaging findings (91 pts) vs palpation for pain (30 pts)</td>
<td>Prospective observational</td>
<td>A greater proportion of pts referred for injections based on pain palpation reported relief for up to 1 month</td>
<td>Cervical facet joint injections were completed under CT guidance</td>
</tr>
<tr>
<td>Morishita et al 2008</td>
<td>215 pts with degenerative disease of the cervical spine</td>
<td>Retrospective</td>
<td>Neck pain was more common among pts with hypertrophic changes in facet joints</td>
<td>Did not control for confounding variables</td>
</tr>
<tr>
<td>Hechelhammer et al 2007</td>
<td>37 pts who underwent 50 cervical IA facet joint injections</td>
<td>Retrospective</td>
<td>No statistically significant difference in pain relief from cervical facet joint blocks based on osteoarthritic grade</td>
<td>IA injections performed under CT guidance. 56% of injections were peri-articular, 40% were peri- and IA, and 4% were IA</td>
</tr>
</tbody>
</table>

CT, computed tomography; IA, intra-articular; MRI, magnetic resonance imaging; MVC, motor vehicle collision; pts, patients.

**Table 8** Studies evaluating the association between MRI imaging pathology and facet pain

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
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<tbody>
<tr>
<td>Daimon et al 2019</td>
<td>81 pts presenting immediately and 20 years after whiplash injury for MRI</td>
<td>Prospective longitudinal</td>
<td>Progression of degeneration on MRI was observed in 95% of subjects. Changes in neck pain severity was not associated with progression of degenerative changes on MRI</td>
<td>C4–5 and C5–6 levels most frequently exhibited degeneration</td>
</tr>
<tr>
<td>Nevalainen et al 2016</td>
<td>173 pts with MRI studies demonstrating cervical facet edema</td>
<td>Retrospective</td>
<td>Significant correlation between neck pain and/or unilateral radiculopathy and ipsilateral bone marrow edema. No correlation between pain intensity and severity of edema</td>
<td>9% prevalence of cervical facet edema, most commonly at C3–4, C4–5, and C2–3. The study did not confirm the presence of facet-mediated pain through diagnostic blocks</td>
</tr>
<tr>
<td>Cohen et al 2007</td>
<td>92 pts who underwent cervical facet RFA after positive MBB</td>
<td>Retrospective</td>
<td>The only clinical variable associated with positive response to cervical medial branch RFA was paraspinal tenderness</td>
<td>Facet pathology was noted on cervical spine MRI in 48% of pts but was not predictive of treatment outcome</td>
</tr>
</tbody>
</table>

MBB, medial branch block; MRI, magnetic resonance imaging; pts, patients; RFA, radiofrequency ablation.
on cervical MRI in almost half the patients, these findings were not predictive of treatment outcomes.

Recommendations
We conclude that the current evidence is insufficient to assess the balance of harms and benefits of radiological imaging modalities for the diagnosis of cervical facetogenic pain and as a prognostic indicator for the success of cervical facet blocks or RFA; Grade 1 recommendation. However, for the purpose of procedural planning, radiological imaging should be strongly considered when indicated; Grade C recommendation, low level of certainty.

QUESTION 4: SHOULD PHYSICAL THERAPY AND/OR CONSERVATIVE TREATMENT BE A PREREQUISITE BEFORE PROGNOSTIC FACET BLOCKS? IF SO, FOR HOW LONG SHOULD THEY BE CONTINUED?

Conservative management of cervical facet joint pain typically involves a trial of analgesic and anti-inflammatory medications, physiotherapy (also known as physical therapy), and various other modalities (heat and/or ice, massage, transcutaneous electric nerve stimulation, traction, and spinal mobilization). Although supported by little evidence, these conservative treatments are frequently applied before consideration for interventional treatments.88 Many clinical studies8 89 100 evaluating cervical facet injections or radiofrequency (RF) neurotomy have required a course of conservative treatment, while others have not.101–103

Although not well supported in the literature, the rationale behind the de facto use of conservative management is that it may assist the recovery process. The use of conservative management prior to prognostic facet blocks is based on pragmatism and to some extent insurance requirements, not empiric data. As with the majority of musculoskeletal conditions, neck pain generally is self-limiting. However, the clinical course of neck pain in the absence of formal treatment is not well-documented. One prospective cohort study describes the natural course of acute neck and low back pain (LBP) in the general population of Norway.104 The authors found that the course of pain declined rapidly within 1–2 months of onset in most subjects, with small changes over the follow-up year. These findings provide a general timeframe for the use of conservative management for most patients with acute, but not chronic, neck pain.

The efficacy of physiotherapy for acute neck pain was examined in a prospective cohort study by Vos et al105 in which 187 patients with acute neck pain (mean duration at baseline was 16 days) were followed for 1 year. During that period, 118 patients were referred to a physiotherapist with 74% (87/118) reporting recovery at 1-year follow-up. Interestingly, the authors found that 79% (55/69) of control patients reported similar recovery at 1-year follow-up without any physiotherapy intervention. This again implies that most cases of acute neck pain resolve spontaneously without the need for further work-up and treatment. An RCT performed on 156 patients with neck pain found that the use of a multimodal approach containing self-management with coping skill training was more effective than individualized physical therapy over a 2-year follow-up.106 However, in another study, manual physical therapy and exercise were shown to be a more effective treatment strategy than advice on motion exercises for chronic mechanical neck pain.107 It is important to note that neck pain does not necessarily equate to cervical facet joint pain, as there are other causes of neck pain including myofascial or discogenic neck pain. However, cervical facet joint pain is known to make up a substantial portion of the patient population with neck pain, with a reported prevalence in a pain clinic population approaching 60%.6

The use of conservative treatments (which are often advocated for non-specific symptoms) prior to prognostic blocks may also be related to the absence of pathognomonic physical examination or radiological findings for facet joint pain. In the absence of any reliable means of clinically diagnosing facet joint pain, the treatment of mechanical or neuropathic neck pain often starts with less invasive treatments. The response to conservative treatments may prevent the need for further work-up and interventions. Of note, there is no evidence that conservative treatment guarantees functional improvement or pain reduction, nor does lack of response to conservative measures predict success or failure of procedural interventions. Confounding things further, responders and non-responders to prognostic facet blocks were found in one study to demonstrate similar presentation of sensory disturbances, motor dysfunction, and psychological distress.108

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### Table 9: Studies evaluating the association between SPECT and PET imaging modalities and facet pain

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sawicki et al 201749</td>
<td>10 pts with suspected cervical facet joint pain</td>
<td>Retrospective case–control</td>
<td>F-FDG PET/MRI was used to determine the location of MBB in 6 pts. Landmarks were used in 4 PET-negative pts. The PET-positive pts had significantly less pain up to 3 months after MBB</td>
<td>CT-guided MBB done with 3 mL of LA and steroid. Pain did not decrease in PET-negative pts</td>
</tr>
<tr>
<td>Lehman et al 201460</td>
<td>74 pts with SPECT/CT scan of the cervical spine who underwent IA facet joint injection or dual comparative MBB</td>
<td>Retrospective</td>
<td>18 pts received cervical IA facet joint injections and 1 received cervical MBB. 52 pts (70%) had at least one discrepancy between facet joint activity on SPECT/CT and clinical treatment</td>
<td>103 of 195 (53%) active facet joint(s) observed on SPECT/CT did not correlate with clinical findings</td>
</tr>
<tr>
<td>Matar et al 201359</td>
<td>72 pts with clinically suspected facet-mediated neck and back pain and non-conclusive MRI/CT findings</td>
<td>Retrospective</td>
<td>Among the 24 cervical SPECT-CT scans, 13 (52%) had evidence of active cervical facet joint arthropathy and 10 (36%) demonstrated other pathology</td>
<td>No correlation with outcomes from IA facet joint blocks</td>
</tr>
<tr>
<td>Perez-Roman et al 2020104</td>
<td>190 pts with axial neck (n=25) or back pain underwent high-resolution SPECT/CT scan</td>
<td>Retrospective</td>
<td>A total of 202 hypermetabolic facet joints in 85 pts (48%) were identified. Lumbar facet joints were most commonly affected (69%), followed by cervical (24%) and thoracic regions (6%). C1–2 and C2–3 (22% each) were the most commonly affected in the neck. In the 37 pts who reported axial neck pain, 16 (43.2%) were found to have cervical facet hypermetabolism</td>
<td>Diagnostic facet blocks were not performed. Injection techniques were not described</td>
</tr>
</tbody>
</table>

CT, computed tomography; F-FDG, F-fluorodeoxyglucose; MBB, medial branch block; MRI, magnetic resonance imaging; PET, positron emission tomography; pts, patients; SPECT, single photon emission CT.
In a Cochrane Database systematic review of physical therapy for the treatment of non-specific chronic neck pain, there was moderate evidence supporting cervico-scapulothoracic and upper extremity strength training, endurance training, strengthening and stretching exercises, mindfulness exercise, and stabilization exercises to improve pain and function based on moderate-quality evidence. A meta-analysis evaluating physical therapy techniques found that therapeutic exercise had significant short-term and intermediate-term effects, but no long-term benefit on pain. Physical therapy did not provide significant short-term, intermediate-term, or long-term effects on disability. In a systematic review evaluating exercise programs for chronic non-specific neck pain, the authors found strong evidence for the effectiveness of muscle strengthening and endurance exercises. Moderate evidence supported the use of muscle endurance exercise in reducing disability attributed to neck pain. However, no physical therapy efficacy studies were found in the literature that included patients with MBB-proven cervical facet joint pain.

Medications have been recommended as part of a conservative treatment regimen for patients with cervical facet-related pain, despite there being a scarce number of high-quality studies evaluating pharmacotherapy for chronic neck pain. Accurate extrapolation is even more challenging since most studies included individuals with non-specific neck pain. As noted in a review by Cohen, systemic non-steroidal anti-inflammatory drugs have been found to be beneficial for spinal pain in general, but not specifically for neck pain. The use of acetaminophen, topical and oral non-steroidal anti-inflammatory drugs, and intermediate doses of the muscle relaxant (cyclobenzaprine) were found to be useful in the treatment of acute and subacute neck pain symptoms.

Third occipital neuralgia/cervicogenic headaches

As with cervical facet joint-mediated pain, third occipital neuralgia and cervicogenic headaches can only be reliably diagnosed with IA injection or MBB. As per revised criteria of the International Headache Society (IHS), evidence of a cervical source of pain is required for the diagnosis of cervicogenic headache. However, the IHS notes that clinical features historically thought to be related to cervicogenic headaches are not unique and “they do not necessarily define causal relationships”. In a review by Bogduk and Govind, the authors concluded that diagnostic blocks are the only means of reliably establishing this diagnosis.

There have been several moderate quality studies exploring the use of conservative treatments including therapeutic exercises for third occipital neuralgia and cervicogenic headache. These studies have reported conflicting evidence regarding the effects of manipulative therapy on cervicogenic headaches. However, study results must be interpreted with caution since the diagnosis of cervicogenic headache was made clinically instead of by diagnostic blocks. In the only RCT that investigated the effects of exercise in the treatment of cervicogenic headache, Jull et al found that either exercise or spinal manipulation provided statistically significant improvements relative to a control group through 12 weeks, with the combination treatment group faring no better than stand-alone treatments. For chronic cervicogenic headache, moderate-quality evidence supports static-dynamic cervico-scapulothoracic strengthening/endurance exercises including pressure biofeedback at long-term follow-up. In a review by Bogduk and Govind, the authors concluded that manual therapy (including physiotherapy) was no more effective than exercise alone. The authors further proposed a ‘pragmatic clinical approach’ involving exercises with or without manual therapy for clinically suspected cervicogenic headache, with the efficacy of most other treatments (eg, medications, transcutaneous electrical stimulation) being speculative at best.

Recommendations

Due to a generally favorable natural history of acute neck pain symptoms, our recommendation is for a 6-week trial of conservative management prior to prognostic cervical facet blocks to prevent unnecessary invasive procedures and associated healthcare costs. The use of conservative measures may prevent the need for prognostic blocks (or further interventions) but does not preclude the use of blocks for those patients who have failed conservative treatments. Grade B recommendation, moderate level of certainty for a requirement of conservative management before prognostic blocks in patients with at least 3 months of neck pain; Grade C recommendation, low level of certainty for at least a 6-week trial of conservative therapy which may vary based on a personalized medicine paradigm; grade I recommendation for concomitant use of conservative measures to accompany prognostic blocks.

QUESTION 5: IS IMAGE GUIDANCE NECESSARY FOR CERVICAL FACET BLOCKS AND RFA?

Guidance versus no guidance: accuracy and safety

Whereas no specific imaging modality has been identified as the reference standard, image guidance for cervical spine interventions has become an essential component in minimizing patient harm and optimizing results. For cervical facet procedures including IA injections, MBB and medial branch RFA, fluoroscopy and, to a much lesser extent, CT and ultrasound (US) are commonly used. Cervical procedures may pose a higher risk than analogous procedures in the lumbar region; therefore, the use of advanced imaging including US or CT may be more common and useful. Similar to the lumbar region, the use of imaging allows accurate needle placement to ensure the lowest volume of anesthetic is administered, thereby reducing spread to surrounding tissues which may lead to false-positive test results. Image guidance also improves safety through direct visualization of bony elements of the neuraxis, thus avoiding proximal structures including pleura, neural foramina, and vascular supply. In the USA, the current procedural terminology (CPT) code 77003 (fluoroscopic guidance and localization of needle or catheter tip for spine or paraspinal diagnostic or therapeutic injection procedures) should not be used for facet blocks or RF as imaging is considered an integral part of the procedures. When US guidance is used, the category III codes 0213T–0218T should be reported.

Manchikanti et al examined procedural risks of fluoroscopically-guided cervical facet procedures in a prospective observational study in which 3370 cervical MBBs were performed. They found no instances of nerve damage, spinal cord injury, infection, or epidural hematoma; however, cervical procedures had a higher risk of intravascular adverse events (eg, oozing, intravascular penetration) compared with thoracic and lumbar regions. The lack of moderate to severe adverse events or a difference in incidence between cervical and lumbar spine interventions when image guidance is used is unsurprising given the rarity of moderate to severe complications associated with either region. Neither this nor other studies examined the relative risk of performing cervical joint procedures with and without image guidance. This type of empiric study is unlikely to be
designed or performed, as the scientific community has encouraged image guidance as a general harm reduction strategy. Heckman and colleagues reported a case of transient tetraplegia following cervical facet IA injection in which no image guidance was used, and closed claims analyses have revealed at least two other cases involving facet injections in which the use of imaging was not noted. Cervical joint procedures performed without image guidance are likely to result in at least as many complications and poor outcomes as unguided lumbar paravertebral or facet injections.

### Existing guidelines and insurance coverage

The scientific question related to the accuracy and safety of image-versus non-image-guided procedures has not been adequately evaluated in clinical trials. The ASA’s 2010 practice guidelines are referenced in some insurance company determinations, although their language more generally references ‘appropriate image guidance’ and does not limit recommendations to specific imaging modalities. The SIS guidelines recommend the use of fluoroscopic imaging with multiple views using the lowest amount of radiation but do not mention the use of CT, US, and imaging modality combinations. For MBBs, the nerve is not directly injected, and closed claims analyses have revealed 7% using fluoroscopic guidance with 20% incidence of intravascular penetration. For cervical MBB, 20% incidence of extravascular penetration.

<table>
<thead>
<tr>
<th>Table 10</th>
<th>Studies comparing imaging modalities for cervical facet injections</th>
</tr>
</thead>
<tbody>
<tr>
<td>Author, year</td>
<td>Patient population</td>
</tr>
<tr>
<td>Park et al 2017</td>
<td>Pts with injection-confirmed chronic cervical facet joint pain who received US-guided (n=68) or fluoroscopically-guided cervical MBB (n=58)</td>
</tr>
<tr>
<td>Finlayson et al 2013</td>
<td>40 pts undergoing TON block were randomized to fluoroscopic or US guidance</td>
</tr>
<tr>
<td>Finlayson et al 2015</td>
<td>50 pts undergoing C7 MBB under US or fluoroscopic guidance</td>
</tr>
<tr>
<td>Manchikanti et al 2012</td>
<td>7500 episodes of 43,000 facet joint nerve blocks with 3370 episodes in the cervical region. All facet joint nerve blocks were performed under fluoroscopic guidance</td>
</tr>
<tr>
<td>Zhou et al 2010</td>
<td>31 pts with refractory cervicogenic headache who underwent fluoroscopically-guided AA and C2–3 facet joint injections and C2 and 3 dorsal rami blocks</td>
</tr>
<tr>
<td>Obernauer et al 2013</td>
<td>40 pts (54 joints) with subacute axial neck pain were randomly assigned to US- or CT-guided IA facet injections</td>
</tr>
</tbody>
</table>

AA, atlanto–axial; CT, computed tomography; IA, intra-articular; MBB, medial branch block; pts, patients; TON, third occipital nerve; US, ultrasound.

Use of fluoroscopy

Fluoroscopy is the reference standard for prognostic interventions of the cervical spine including TON block and MBB. A randomized study of the cervical spine found the incidence of ‘missed nerves’ to be 7% using fluoroscopic guidance with 0.25–0.5 mL of injectate, and an earlier study showed that a 0.5 mL injection reliably encompassed the target nerve.
superior benefit immediately post-procedure and at 1 month for US-guided single-level injections, with shorter procedure duration. For two-level injections, the benefit favoring US-guided injections fell shy of statistical significance. Eichenberger et al.\(^{144}\) achieved cutaneous analgesia in the distribution of the TON after a US-guided TON block in nine of 10 injections in normal volunteers. It should be noted that the authors used a large volume (0.9 mL) of injectate which will spread well beyond the margins of the TON.\(^{74}\) The C2–3 joint was correctly identified in 27 of 28 cases, and in 23 of 28 injections the needle fell within 0.5 mm of the target nerve.\(^{144}\) These findings were confirmed in a subsequent volunteer study by a group with overlapping authors.\(^{357}\)

Finlayson et al.\(^{136}\) performed a randomized study in 40 patients undergoing TON block to determine the comparative effectiveness of fluoroscopic versus US guidance. Their study found comparable effectiveness (19 of 20 patients received successful TON hypoesthesia) with US guidance, which required fewer needle adjustments than fluoroscopically-guided interventions. The TON was directly identified in 16 of 20 US procedures and vascular penetration was observed in zero patients in the US group versus one in 20 in the fluoroscopy group.

These findings were replicated in a retrospective comparative study by Park et al.\(^{145}\) conducted in 126 patients undergoing cervical MBB by either fluoroscopic or US guidance. Their results demonstrated similar accuracy rates, but reduced procedure time and needle adjustments using US. Paredes et al.\(^{146}\) conducted a systematic review and meta-analysis showing that using US for cervical prognostic interventions including TON block and cervical MBB was non-inferior to fluoroscopic guidance, albeit with a lower incidence of vascular penetration and no radiation exposure. US may also offer the additional benefit of real-time imaging of the cervical spine.

Whereas US may provide comparable accuracy and confer some advantages over fluoroscopy, several studies have revealed diminished accuracy rates for C7, which may also be more challenging to block with fluoroscopy.\(^{137,138}\) In one randomized study involving 50 patients, US and fluoroscopy were found to have similar accuracy rates (92–96%) and to provide comparable post-block pain relief, although the former was associated with shorter performance time and less intravascular contrast spread.\(^{147}\) It is important to recognize that even an imaging modality that permits direct visualization of neurovascular structures is not devoid of risks, with Park et al.\(^{148}\) reporting a case of permanent spinal cord injury after a C7 MBB was performed under US guidance, which reinforces the challenges at this cervical level (table 10).\(^{137}\)

### Imaging for therapeutic interventions (IA steroid and RFA)

**Use of fluoroscopy, CT, and US**

In the lumbar anatomic region, recent multi-society guidelines recommended fluoroscopy as the preferred imaging modality for IA injections and lumbar medial branch RFA.\(^{29}\) The use of fluoroscopic guidance is well-established for TON through C8 medial branch RFA procedures.\(^{21,68,101,149}\) However, the use of CT and US guidance for cervical interventions including IA facet injections with steroid is still in its infancy and cadaveric evaluation has yet to definitively establish comparative effectiveness and safety relative to the fluoroscopic approach.\(^{143,150}\) A prospective clinical trial comparing CT versus US-guided IA facet injections in 40 adults demonstrated equivalent accuracy and effectiveness.\(^{143}\) However, the US-guided procedures were faster to perform, with no radiation exposure. A single RCT has been performed evaluating cervical medial branch RFA using US. Siegenthaler et al.\(^{151}\) examined the effect of US-positioned and fluoroscopically-confirmed placement to refine cannula

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**Table 11** Clinical studies evaluating AO joint injections

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Busch and Wilson, 1989(^{146})</td>
<td>25 pts with head and neck pain of which 2 pts received LA and steroid AO and AA joint injections</td>
<td>Retrospective</td>
<td>Pain relief in both cases ranged from 3 weeks to 4 months (1 patient received serial injections)</td>
<td>No reported adverse events other than transient ataxia and worsening pain for 24–48 hours. Fluoroscopically-guided AO and AA injections performed concurrently</td>
</tr>
<tr>
<td>Dreyfuss et al 1994(^{149})</td>
<td>3 pts who received fluoroscopically-guided AO injections</td>
<td>Case series</td>
<td>Pain relief in all 3 pts ranging from 6 to 12 months</td>
<td>No reported adverse events. Multiple concurrent injections limits generalization</td>
</tr>
<tr>
<td>Lee et al 2015(^{169})</td>
<td>29 pts with refractory headache and neck pain and findings suggestive of AO joint pain. Pts with ≥50% pain relief after diagnostic AO block underwent fluoroscopically-guided AO joint injections with LA and steroid</td>
<td>Prospective observational study</td>
<td>20 of 24 (83%) pts had a positive diagnostic block. Pain scores and function improved from baseline at 2-month follow-up</td>
<td>Pts received two AO joint injections 1 week apart. No reported adverse events</td>
</tr>
<tr>
<td>Centeno et al 2018(^{158})</td>
<td>10 pts received bilateral AO joint injections</td>
<td>Case series</td>
<td>All injections reported as ‘successful’ with no adverse events</td>
<td>Head flexed to open up joint and rotated ipsilaterally to displace vertebral artery medially. Injectate composition not reported. Effectiveness outcome measures and follow-up period not reported</td>
</tr>
<tr>
<td>Shin et al 2018(^{151})</td>
<td>23 pts with chronic upper cervical pain, pain score ≥23/10, and ≥50% pain relief after diagnostic AO joint block received either AO joint injection with LA and steroid (n=11) or pulsed RF of the AO joint (n=12)</td>
<td>Randomized comparative-effectiveness trial</td>
<td>Pain score improved from baseline with sustained relief at 6 months with no differences between groups</td>
<td>Between 60% and 70% of pts achieved ≥50% pain relief through 6-month follow-up. Study not blinded. No adverse events reported</td>
</tr>
<tr>
<td>El Abd et al 2008(^{146})</td>
<td>Single patient with right-sided neck pain and headache due to congenital fusion of AO joints bilaterally. Two injections done with LA and steroid</td>
<td>Case report</td>
<td>2 weeks after second therapeutic injection, patient reported 75% improvement. At 6 and 12 month follow-ups, pain reduction persisted</td>
<td>Authors reported no noticeable improvement in ROM</td>
</tr>
</tbody>
</table>

AA, atlanto–axial; AO, atlanto–occipital; LA, local anesthetic; pts, patients; RF, radiofrequency; ROM, range of motion.
Positioning for cervical medial branch RFA in a cohort study involving 15 patients with an average body mass index of 26. The authors demonstrated that the target nerve was visible under US guidance in all patients at all levels and that long-term effectiveness was comparable to the fluoroscopic interventional literature. However, the authors cautioned against performing US-guided cervical medial branch RFA without fluoroscopic guidance.

**Limitations of fluoroscopy, CT, and US**

The use of fluoroscopy is limited by radiation exposure and an inability to directly visualize the nerve and its trajectory. Additionally, the upfront costs including the C-arm and monitor, radiology technician, and fluoroscopic table represent a barrier. For MBB, CT precludes the use of real-time contrast injection or DSA to detect intravascular uptake. Regarding cervical medial branch RFA, the imaging constraints imposed by fluoroscopy recommendations are present but less substantial than in the lumbar spine, enabling parallel or near-parallel placement of electrodes. Yet, the widespread use of CT remains limited because of substantial equipment costs, radiation exposure, lack of real-time vascular imaging, and the need for different patient positions during procedures performed at the upper and lower facet joints.

The use of US may provide an alternative imaging modality for performance of MBB but it is not ideal for cervical IA injections or medial branch RFA that require a set trajectory. Although US is portable, can be used in pregnancy, and does not require the use of protective garments, there are significant disadvantages of using US guidance for cervical spine interventions. US cannot visualize the entire field including adjacent levels, thereby increasing the risk of incorrect level identification. Widespread adoption may also be limited by anatomic difficulties associated with specific anatomic levels in the neck.

### Table 12  Clinical studies evaluating AA joint injections

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bogduk and Marsland, 1988</td>
<td>24 consecutive pts who underwent cervical injections for head and neck pain, 4 of whom received AA injections with LA and steroid</td>
<td>Retrospective study</td>
<td>1 of 4 (25%) pts obtained pain relief from AA joint injection for 2 months</td>
<td>Used a fluoroscopically-guided posterior approach. Pts also trialed other cervical injections. No reported adverse events</td>
</tr>
<tr>
<td>Lamer, 1991</td>
<td>2 pts with cervical spine OA and ear pain provoked with head turning received AA joint injection with LA and steroid</td>
<td>Case series</td>
<td>Both pts had pain relief injection</td>
<td>Used a fluoroscopically-guided posterior approach. Injectate volumes not noted. Duration of pain relief not reported</td>
</tr>
<tr>
<td>Chevrot et al 1995</td>
<td>100 pts (osteoarthritis, rheumatoid arthritis, anklyosing spondylarthritis, and diverse conditions) who received AA joint injections with LA and steroid</td>
<td>Retrospective study</td>
<td>18 (60%) of first 30 pts showed clinical improvement (duration of follow-up 6 months to 3 years)</td>
<td>Used a fluoroscopically-guided posterior oblique approach. One accidental vertebral artery puncture without clinically significant sequelae</td>
</tr>
<tr>
<td>Glemarac et al 2000</td>
<td>26 pts with either mechanical (n=16) or inflammatory disorders (n=10) who received AA joint injections with steroid (no LA)</td>
<td>Retrospective study</td>
<td>69.3% responder rate with mean pain score reduction of 52.3% and mean duration of pain relief 8.1 months</td>
<td>Used a fluoroscopically-guided posterior approach. Those with inflammatory conditions responded better than those with mechanical disorders. One case of moderately severe hypertension following injection</td>
</tr>
<tr>
<td>Aprill et al 2002</td>
<td>34 pts with occipital pain and clinical features suggestive of AA joint origin received AA injection with LA and steroid</td>
<td>Prospective observational study</td>
<td>21 of 34 (62%) pts obtained complete pain relief for at least the duration of action of LA</td>
<td>Used a fluoroscopically-guided posterior approach. Duration of response not assessed. Clinical features did not predict positive response</td>
</tr>
<tr>
<td>Narouze et al 2007</td>
<td>32 pts with clinical features suggestive of AA joint pain who received AA injection with LA and steroid</td>
<td>Retrospective study</td>
<td>15 of 32 (47%) pts had complete pain relief for the duration of action of LA, 26 of 32 (81%) had ≥50% improvement sustained at 3 months</td>
<td>Used a fluoroscopically-guided posterior approach. 5 of 32 (16%) pts stopped opioid use and 3 of 32 (9%) pts had complete pain relief sustained at 9 months</td>
</tr>
<tr>
<td>Zhou et al 2010</td>
<td>31 pts with suspected cervicogenic headache who received AA joint, C2 and C3 dorsal rami, and C2–3 facet joint injection with LA and steroid</td>
<td>Retrospective study</td>
<td>28 of 31 (90%) pts had &gt;50% pain relief with an average duration of 21.7 days</td>
<td>Used fluoroscopically-guided posterior approach. Pts also experienced decreased headache frequency and duration. Non-responders diagnosed with temporomandibular disorder (n=2) and migraine (n=1)</td>
</tr>
<tr>
<td>Aiudi et al 2017</td>
<td>72 pts who received AA joint injections with LA and steroid</td>
<td>Retrospective study</td>
<td>Adverse event rate was 25 of 135 (18.5%) injections with 13 procedural events (vascular uptake/paresthesia) and 12 post-procedural events (increased pain/neurologic symptoms)</td>
<td>Used a fluoroscopically-guided posterior approach. No serious adverse events noted and all post-procedural adverse events resolved within 3 months. Pain outcomes not reported</td>
</tr>
<tr>
<td>Hetta et al 2019</td>
<td>60 pts with rheumatoid arthritis and AA joint pain received AA joint injection with LA and steroid or LA and saline</td>
<td>Randomized controlled</td>
<td>LA+steroid injection &gt; LA-only injection through 3-month follow-up for pain and function, with improvement in imaging findings not observed in LA-only group</td>
<td>Fluoroscopically-guided posterior approach. Permitted continued use of disease-modifying agents and oral NSAIIDs for breakthrough pain</td>
</tr>
<tr>
<td>Kuklo et al 2006</td>
<td>14 pts with AA joint pain received AA joint injection ‘via a standard technique’ with LA+steroids – no specifics provided</td>
<td>Retrospective study</td>
<td>11 of 14 (79%) pts treated ‘successfully’ with 1–4 injections over the study period with significant pain relief. 3 refused injections, 3 had no relief from multiple injections, 3 had temporary relief from injections and went on to C1–2 fusion surgery</td>
<td>No discussion on how ‘significant’ relief was defined</td>
</tr>
</tbody>
</table>

AA, atlanto–axial; LA, local anesthetic; NSAIDs, non-steroidal anti-inflammatory drugs; OA, osteoarthritis.
especially involving C7. Of note, the US-guided approach to the cervical medial branch is less commonly taught in residency, fellowship, and postgraduate courses; therefore, widespread adoption would require additional physician training. Although US enables direct visualization of nearby vessels, it does not easily detect inadvertent vascular uptake, which can be reliably detected using real-time contrast injection or DSA. The limitations in the lumbar spine related to decreased needle visibility due to body habitus and depth to target are present, but may be less of a barrier in the cervical spine.

Recommendations

We recommend that fluoroscopy or (in providers with expertise) US be used for cervical MBB. US can be useful in patients in whom radiation exposure may be associated with potential harm; however, the lack of training may limit widespread adoption; Grade A recommendation, moderate level of certainty. For IA injections, we recommend the use of fluoroscopic imaging as the additional radiation exposure from CT compared with fluoroscopy precludes any theoretical benefit; Grade C recommendation, low level of certainty. For cervical medial branch RFA, we recommend that fluoroscopy be used as the additional radiation exposure from CT compared with fluoroscopy precludes any theoretical benefit. Whereas CT-fluoroscopy is associated with less radiation than CT alone, it is not widely available and adds significant upfront equipment costs and radiation exposure; Grade A recommendation, high level of certainty for the use of imaging, Grade B recommendation, moderate level of certainty for the use of fluoroscopy instead of other imaging modalities.

Question 6: What is the optimal technique for injection into the AA and AO joints? Should steroids be used and, if so, what type of steroids? What are the most common complications and how can they be minimized?

Image guidance and patient positioning

The use of image guidance is essential when performing AA and AO joint injections (tables 11 and 12). In clinical practice, fluoroscopy is typically used. Although CT guidance has been anecdotally reported, no studies describe this technique or outcomes in the literature. The feasibility of an US-guided approach for AO joint injection has been described in cadavers, but no clinical studies have been published to demonstrate safety or efficacy. However, a combined approach using fluoroscopy with US assistance to identify the vertebral artery has been advocated. For both AO and AA joint injections, patients are typically placed in the prone position with a pillow or cushion under the chest to allow for flexion of the neck. A fluoroscopically-guided posterior (also known as posterior parasagittal or posterior sagittal) approach is typically employed (table 11). Some advocate rotating the head 30 degrees ipsilateral to the side of injection to displace the vertebral artery to a more medial location. However, earlier descriptions of AO injections had patients placed in a lateral decubitus position with the head rotated contralaterally to the side of the injection. Unlike the AA joint in which the vertebral artery is generally situated lateral to the joint margin, the artery traverses the AO joint space (figure 3). To avoid inadvertent vertebral artery injury or injection, the most superior and lateral portion of the joint is targeted. The joint may be accessed either directly in a coaxial view or after contacting the periosteum and redirecting into the joint. After confirmation of IA needle placement using low-volume contrast injected under real-time fluoroscopy or DSA, approximately 1 mL of injectate is administered (table 11).

Atlanto–occipital (C0–1) joint injection

A fluoroscopically-guided posterior approach is most commonly used to access the AA joint. A fluoroscopic image of the posterior cervical spine in the AP and lateral views is needed to locate the C0–1 joint. Once the joint is located, the needle is advanced in a posterior-inferior direction to access the joint. The needle is then advanced in a posterior-superior direction to inject the joint. The optimal target point is the junction of the lateral one-third and medial two-thirds of the AA joint to minimize the risk of vertebral artery injury (which is generally lateral to the joint line), contacting the C2 nerve root, or dural puncture with intrathecal spread of injectate. The joint may be accessed either via a straight coaxial trajectory or after first making contact with the periosteum along the joint margin to establish depth. After accessing the joint, the needle is advanced into the joint space and injectate is injected. The injection is repeated until a total of 2-3 mL of injectate has been administered.
confirmation of IA needle placement in posteroanterior and lateral views, and with a very small volume of contrast injected under DSA or real-time fluoroscopy, <0.5 mL of injectate is typically used, as is illustrated in all but one of the clinical studies where the volume of injectate was described (table 12). The use of higher contrast volumes is discouraged given the relatively small capacity (≤1 mL) of the joint.164

Intra-articular steroids for AO and AA joint injections

AO and AA IA steroid injections have been reported to be therapeutic interventions for pain emanating from these joints since the late 1980s. Subsequently, evidence supporting this modality has come primarily from case reports and series and retrospective studies. Very few of these studies reported administration of a separate diagnostic IA injection with LA prior to the administration of a therapeutic injection with steroid.165 166 More recently, prospective observational studies and RCTs have been performed to identify whether IA steroid injections have superior efficacy to non-steroid (LA or saline) IA injections. The current body of available literature has provided modest evidence that is generally supportive of the use and effectiveness of IA steroid AA and AO injections in the treatment of a variety of different patient populations including: cervicogenic and occipital headache,5 165-167 chronic neck and head pain,8 62 159 163 168-171 and pain due to inflammatory disease of the AO and/or AA joints.160 172 173 No studies or review articles have been published regarding which type of steroid (short-acting, long-acting, particulate vs non-particulate) should be used for AO and AA joint injections. (tables 11 and 12)

Only two studies have used a prospective randomized design with a comparative/control arm to determine potential efficacy differences between AO or AA injections with and without steroids. In the study by Hetta et al,173 patients with rheumatoid arthritis and AA inflammation and pain were randomized to either AA injections with LA and steroid or LA and normal saline. All patients were maintained on a standardized regimen of oral steroids and immunosuppressive therapy during the study. The authors reported that patients who received LA and steroid AA injections experienced statistically significantly greater reductions in numerical pain scores and improvements in physical functioning as measured by the Neck Disability Index at 3-month follow-up. They determined that the LA and steroid group had MRI-confirmed resolution of the inflammation observed pre-procedure compared with the LA and saline group. Shin et al171 performed a randomized prospective study investigating the comparative effectiveness of AO joint LA and steroid injection versus AO joint pulsed RF. The findings in this study showed no superiority of one modality compared with the other; however, both groups experienced significant within-group reductions in numerical pain score ratings over 6 months.

A systematic review and meta-analysis has recently been published highlighting that, in the broad scope of non-cancer interventional injections, there is little statistical increase in the effect sizes seen with the addition of steroid to LA and/or saline for IA injections and other procedures.174 The authors concluded that the use of steroids in interventional pain procedures may not be justified in all, or even most cases. They recommended that an in-depth evaluation of the risks, benefits, and safety of using steroids should be prioritized when performing interventional pain procedures for patients with non-cancer pain.

Complications of AO and AA joint injections and risk mitigation

The risk of adverse events associated with AA joint injections was found to be 18.5% (25 of 72 patients) in a retrospective observational study.163 In this cohort, no serious adverse events were reported and the most common side effects were dizziness, paresthesia, and/or increased pain. Vascular uptake on contrast injection (not differentiated between arterial or venous) was noted on real-time imaging or DSA during five of the injections that either resulted with needle repositioning or resulted in aborting the procedure due to safety concerns. One patient was noted to have blood return on aspiration with needle insertion which resulted in cessation of the procedure. Whereas no serious adverse events were reported in this study, the potential for serious adverse events such as inadvertent intrathecal injection, vertebral artery injury or injection, and C2 dorsal root ganglion injury (with AA injection) exist.175 The risk of adverse events is reduced with optimal needle placement. However, the presence of anatomic variations could result in adverse events. For example, anatomic studies have shown that in 0.72–1% of patients the vertebral artery is present along the needle trajectory for AA joint injections, and in 1.64% of patients the dural sac is vulnerable.162 176 These anatomic variations are the basis for obtaining advanced imaging (CT/MRI of the cervical spine) prior to performing AO and AA injections.176

The type of steroid used is also important to minimize the risk of complications. In a preclinical study performed in 11 pigs, the injection of particulate steroid in the vertebral artery resulted in all four pigs failing to regain consciousness and requiring ventilatory support, while the seven pigs injected with non-particulate steroid all recovered.177 A case of posterior circulation stroke resulting in a coma with the withdrawal of care following AA joint injection with a particulate steroid has been reported.178 It is unknown whether pre-procedural advanced imaging was obtained (as is generally recommended) or what type of approach was used, since the only image saved from the injection was in a lateral view.179 180 The use of real-time fluoroscopy and/or DSA has been advocated to prevent intravascular injection and has been mandated in guidelines for transforaminal lumbar epidural steroid injections, giving its greater sensitivity for detecting intravascular uptake and hence preventing catastrophic neurological complications.156 157 181 182 Another case report described the development of AA joint pyogenic osteomyelitis requiring debridement and joint arthrodesis that remained unrecognized for 4 months after an AA joint injection.183 In this report, little information was provided to draw any conclusions regarding risk mitigation and a history of diabetes mellitus placed the patient at higher risk for infection. Although no reported serious complications have been identified with AA joint injections, the theoretical risks are higher than for AA joint injections given the exposed location of the vertebral artery and the closer proximity to the brainstem.177

Recommendations

Pre-procedural advanced imaging of the cervical spine with either CT or MRI should be obtained prior to performing AO and AA joint injections to ascertain pathology and help guide needle trajectory; Grade C recommendation, low level of certainty. When performing AO and AA joint injections, we recommend a posterior approach with confirmation of IA spread using real-time fluoroscopy or DSA in both anteroposterior and lateral views; Grade B recommendation, moderate level of certainty. There is insufficient evidence regarding the use of CT guidance...
or US guidance without fluoroscopy when performing AO and AA injections; Grade I recommendation. There is a small body of evidence that the use of steroids in AO and AA joint injections may be beneficial in selected populations; however, the magnitude of benefit is small; Grade C recommendation, low level of certainty. Based on indirect evidence, we recommend that, if steroids are administered, <1 mL of non-particulate steroids be administered; Grade C recommendation, low-to-moderate level of certainty.

QUESTION 7: WHAT IS THE MOST APPROPRIATE APPROACH TO CERVICAL MBB? WHAT ARE THE RISKS/BENEFITS OF THE DIFFERENT APPROACHES?

There is no published consensus regarding the ‘correct’ approach to performing cervical MBBS. The SIS MBB safety practices publication states, “The ultimate choice of approach or technique to use should be made by the treating physician by balancing potential risks and benefits with each technique for each patient”. It is generally thought that a lateral approach to the TON and the C3 through C7 medial branch nerves is most optimal, while a posterior approach to the C8 medial branch nerve (target is the superior lateral aspect of T1 transverse process) is most ideal. This is, in part, related to the fact that in most patients these respective approaches allow for the target injection site to be accessed by traversing the least amount of tissue from the skin entry point, which may decrease procedure time as well as improve patient comfort and satisfaction. However, critics of this viewpoint argue that a posterior approach best simulates the natural nerve trajectory and that, when physicians use the injection approach with which they are most comfortable, the best results are obtained.

A lateral needle approach can be performed with the patient in the prone, lateral decubitus, and supine position, and there are no studies comparing these approaches. Of note, the benefits purported by a lateral approach remain largely theoretical with minimal literature comparing approaches. Furthermore, exceptions exist and are often dependent on unique anatomic considerations (eg, body habitus, neck length and thickness, neurovascular anatomy). Regardless, lateral and posterior approaches during the performance of these nerve blocks are recommended in established clinical practice guidelines based on foundational literature establishing safety and accuracy when these approaches are employed. Recent studies have introduced the feasibility of US-guided cervical MBBS.

However, the safety and diagnostic characteristics of the US technique, in which contrast spread cannot be observed, are not yet well-defined, and fluoroscopic guidance remains the clinical standard (as discussed in Question 5). In order to appropriately assess the optimal approach to fluoroscopic TON and cervical MBBS, both the accuracy and safety of each approach must be considered.

Accuracy

There is currently no direct comparative evidence indicating that a specific approach is associated with greater target specificity during TON block or cervical MBB. One RCT demonstrated no difference in target specificity of C3–C7 MBB when comparing a lateral approach in the decubitus position to a posterior approach. Another study assessed target specificity of C4–C6 MBB comparing different injectate volumes, but not stratified by different (lateral vs posterior) approaches. No study has reported on the accuracy of the lateral versus posterior approach for TON or C8 MBB. Notably, the foundational studies that established the accuracy and diagnostic value of TON and C3–C7 MBB have generally used a lateral approach (table 13).\textsuperscript{12,18,188,169}

Safety

A lateral approach to TON and C3–C7 MBB and a posterior approach to C8 MBB may confer practical (a lateral approach may be faster to perform and allows for easier use of the single-needle technique)\textsuperscript{190} clinical (less procedure-related pain from less tissue penetration leading to more accurate post-block pain assessment and in some cases a decreased need for sedation), and/or safety benefits, although clear images with a lateral approach may be challenging in heavy-set people and those with broad

<table>
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<tr>
<th>Author et al.</th>
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<tr>
<td>Cohen et al 2010 \textsuperscript{18}</td>
<td>24 pts with axial neck pain. Evaluated the accuracy and safety of posterior vs lateral approach for cervical MBBS</td>
<td>RCT, radiologist blinded</td>
<td>No difference in target specificity of TON and C3–C7 MBBS when comparing lateral to posterior approach</td>
<td>Only RCT to directly compare posterior vs lateral approach for TON and cervical MBBS</td>
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<tr>
<td>SIS guidelines \textsuperscript{18}</td>
<td>N/A</td>
<td>Guidelines</td>
<td>Lateral approach recommended for TON and C3–C7 MBBS</td>
<td>Expert consensus regarding cervical MBB approach based on literature review</td>
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<tr>
<td>Finlayson et al 2013 \textsuperscript{188}</td>
<td>40 pts undergoing TON block with US vs fluoroscopy using a posterior approach</td>
<td>RCT</td>
<td>IA spread of contrast and vascular breach occurred in 15% and 10% of participants, respectively, with the posterior fluoroscopic approach vs 0% with US</td>
<td>No comparison of posterior vs lateral approach</td>
</tr>
<tr>
<td>Wahezi et al 2019 \textsuperscript{189}</td>
<td>C4–6 cervical MBBS using a posterior approach with 0.25 or 0.50 mL of injectate. Post-injection (CT) imaging and gross dissection performed to assess injectate spread</td>
<td>Cadaveric study (n=6, 18 MBBS)</td>
<td>0.25 mL volume was target-specific</td>
<td>No comparison of posterior vs lateral approach</td>
</tr>
<tr>
<td>Verrills et al 2008 \textsuperscript{18}</td>
<td>4134 cervical MBBS (number of pts not noted)</td>
<td>Retrospective cohort study</td>
<td>3.9% rate of vascular uptake associated with the lateral approach</td>
<td>No comparison of posterior vs lateral approach. Posterior approach noted to be used ‘at times on lower cervical joints’</td>
</tr>
<tr>
<td>Jeon et al 2015 \textsuperscript{18}</td>
<td>178 cervical MBBS in 72 pts</td>
<td>Prospective cohort study</td>
<td>10.7% rate of vascular uptake associated with the lateral approach (as detected by DSA)</td>
<td>No comparison of posterior vs lateral approach</td>
</tr>
<tr>
<td>Elgueta et al 2018 \textsuperscript{18}</td>
<td>500 pts with CT angiograms of the head and neck</td>
<td>Retrospective cohort study</td>
<td>Vertebral artery loop located in the typical location of a TON block in 5–8% of individuals</td>
<td>Safety implications with anterior needle trespass during TON blocks</td>
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CT, computed tomography; DSA, digital subtraction angiography; MBB, medial branch block; RCT, randomized controlled trial; TON, third occipital nerve.
shoulders. These respective approaches allow the advancement of the needle to the target location using a coaxial trajectory with the bony articular pillar functioning as a safety backstop. The lateral approach to TON block is the singular exception to this principle, as the needle tip may need to be directed slightly superficial to the lateral aspect of the C2–3 facet joint in order to provide a block that accounts for the cephalo-caudal variability of the TON. 191 The possibility of inadvertent IA penetration or through-and-through trespass into the central canal with a lateral approach must be acknowledged and mitigated. Regardless of the level, this is possible if a true lateral view is not obtained and a long needle is used. The lateral approach also presents the possibility of trespass into the neuroforamen with possible encounter of the exiting spinal nerve root and/or puncture of the dura or spinal cord, or anterior to the neuroforamen and into the vascular prevertebral space. This may occur if the needle is advanced anterior to the lateral pillar. The safety and feasibility of a lateral approach to TON block is supported by a prospective clinical study,136 and while a posterior MBB approach best simulates the recommended needle trajectory for RFA, similar investigations using a posterior approach have not been reported. Further, large cohort studies of cervical MBB procedures performed according to clinical practice guidelines (lateral approach to TON and C3–C7 MBB and posterior approach to C8 MBB) have been associated with an excellent safety profile.192 Alternative methods have been associated with various severe and minor complications, although it is unclear what role the use of a lateral versus posterior approach played as authors did not describe this element of the procedures.126 193 Since a posterior approach is associated with a longer insertion distance and more tissue trauma, an effect which may be magnified in individuals with thick necks, the need for sedation may be higher in some patients. Finally, there is no evidence that the lateral versus posterior approach is associated with a lower likelihood of intravascular injection including inadvertent arterial injection (ie, cervical radicomedullary and vertebral arteries).78 194 Depending on the cervical level, the rate of inadvertent vascular injection appears to vary from 2% to >10% when DSA is used78 134 194; however, the level of confidence in this incidence rate is low due to small sample sizes in the representative primary literature. In one large study evaluating 4134 cervical MBBs, the authors reported a rate of cervical intravascular injection of 3.9%, with a lateral approach used for most blocks and a posterior approach occasionally used for lower cervical levels.194

Considering anatomic variations unique to each patient

When determining the ideal approach to TON and cervical MBB, anatomic factors unique to each patient must be considered. Unique anatomic variations might impact the possibility of: (1) penetration of the C2–3 joint, through and through, with subsequent trespass into the dura or spinal cord (unique to TON targeting from a lateral approach); (2) breach of the vertebral artery; (3) trespass into the neuroforamen with possible encounter of the exiting spinal nerve root and/or puncture of the dura or spinal cord; or (4) needle trespass posterior to the articular pillar and into the dura or spinal cord. In most patients, a needle shorter than 3.5 inches (6.35–8.9 cm) is adequate to reach the target using a lateral approach, thereby reducing the risk of inadvertently reaching the dura or spinal cord as described above. A small diameter (eg, 25-gauge) short needle may reduce the risk of trauma to the vertebral artery if punctured, although there are no studies to support this supposition. Specific to TON block, a CT-angiography study demonstrates that a loop of the vertebral artery may be located in the typical location of a TON block (midpoint of the lateral C2–3 facet joint margin) in 5–8% of individuals.162

Recommendations

For logistical reasons that vary by level and patient, and to optimize safety, we recommend consideration of a fluoroscopically-guided lateral approach for TON and C3–C7 MBB, but a fluoroscopically-guided posterior or posterior oblique approach for C8 MBB. However, physician comfort with these approaches and unique anatomy should be a primary consideration. Primary data comparing the safety and accuracy of the two approaches are limited to one prospective study. Notably, given the unique anatomic considerations in each patient, the approach at a given level should ultimately be at the discretion of the treating physician. For needle size, we recommend consideration of a short 25-gauge needle to reduce the risk of arterial trauma or trespass into the dura or spinal canal when a lateral approach is used, although again physician judgment should be prioritized (ie, longer needles might be needed in obese patients); grade I recommendation.

QUESTION 8: WHAT IS THE IDEAL VOLUME FOR PROGNOSTIC MBB AND IA INJECTIONS?

Factors affecting injectate spread and rationale for considering injectate volume

TON and MBB are considered both diagnostic for facet joint pain and prognostic for nerve ablation. Both functions are contingent on precisely targeting the nerve with minimal or no spread to the surrounding areas.79 76 78 195 The distribution of fluid after injection into anatomic spaces can be affected by several factors including fluid viscosity, injection velocity, direction of the bevel tip, and fluid volume, all of which may interact with one another. Injection velocity has not been found to be a significant factor in neuraxial injection and the effect of bevel orientation has demonstrated inconsistent results.196–198 Regarding composition, most LA agents have similar viscosities, which is lower than that of contrast media and liposomal formulations. This could result in a wider spread to the surrounding structures when using LA alone compared with solutions mixed with contrast or contrast media alone, resulting in false-positive results.187 199

In addition to injectate properties,187 the anatomy of the cervical spine200 and needle trajectory (ie, fluoroscopic lateral or posterior) can theoretically influence injectate spread.78 LA can spread to adjacent pain-generating or transmitting structures besides the targeted nerve(s) to include the lateral branches innervating paraspinal musculature, neural foramen and spinal nerve root, the facet joint capsule, adjacent levels, muscles and ligaments, and into the cervical epidural space. Spread to any of these structures could compromise the specificity, reliability, and positive predictive value of the MBB due to false-positive results.29 Although studies have found no significant difference in the accuracy of cervical MBB and other injections based on needle trajectory and size, there is a theoretical underpinning for technical factors to affect injectate spread.78 201

The anatomy of the cervical spine is such that the vertebral and medial branches are smaller and closer together than at adjacent lumbar levels.200 The largest of the cervical medial branches, the TON has a mean diameter of 1.5 mm,202 with the other medial branches ranging from 0.6 to 1.2 mm in diameter, being slightly smaller at more caudal levels.202 203 The C4–C8 medial branches vary in their courses in relation to the rostral-caudal location of
the waist of the articular pillar between the periosteum and the tendon of the semispinalis capitis muscle. Barring the C5 mediastinal branch which traverses the center of the transverse process, the medial branches at C3–C6, and C7 course higher on their articular pillars. It has been shown that about one-quarter of individuals have a dual medial branch at C4, with a smaller percentage having two nerves in close proximity at more caudal levels. The target points for the medial branches are closer to the spinal nerve roots than in the lumbar spine, which suggests the need for lower volumes. The horizontally aligned articular pillars (figure 3) are closer to the spinal nerve roots than in the lumbar spine, which makes them more specific to the cervical spine.

Medial branch (MBB)
Indirect evidence from lumbar MBB and other injections
The most influential factor that affects the validity and specificity of blocks is the area of the spread of the injectate. It has been shown from studies involving selective nerve root blocks, lumbar MBB, and sacroiliac joint injections that lower volumes increase the accuracy and specificity of blocks. Studies have shown that volumes as low as 0.3 mL for cervical MBB could result in false positives. However, even 0.3 mL spreads to an area greater than the volume of a thermal RF lesion created by an 18-gauge cannula with a 10 mm active tip.

Several studies carried out in the lumbar spine indicate that the use of smaller MBB volumes may enhance specificity. Volumes used for prognostic lumbar MBB have varied from 0.3 mL to 1.0 mL, with no obvious effect on medial branch RFA clinical outcomes based on indirect comparisons. In the study by Tekin et al. which used a single block with 0.3 mL of LA as a diagnostic test, conventional lumbar medial branch RFA was found to be superior to pulsed RF for up to 1 year. For therapeutic lumbar and cervical MBB, the volumes have ranged from 0.5 to 2 mL in clinical trials. A CT study performed in the lumbar spine clearly demonstrated that fluoroscopically-guided MBB with 0.5 mL was sufficient to anesthetize the lumbar medial branch in all 120 blocks, which suggests that lower volumes may enhance specificity. In a case report published in abstract form, a total of 4 mL of LA given before right-sided C3–5 mediastinal branch RFA resulted in temporary hoarseness, dysphagia, and difficulty coughing. For the smaller cervical mediastinal branch contained within a more confined area, it is difficult to justify the use of higher volumes.

Direct evidence from cervical MBB injections
Cohen et al. conducted an RCT evaluating the accuracy and specificity of different cervical MBB injectate volumes. The authors randomized 24 subjects to receive either 0.5 mL or 0.25 mL of LA injectate mixed with contrast. Subjects were suballocated to receive blocks using either a posterior or lateral approach. The accuracy of the block and the incidence of aberrant spread of the injectate were then evaluated by CT scan. The study found that both 0.25 mL and 0.5 mL volumes of injectate enveloped the mediastinal branch in 93% of the 86 injections, with no statistically significant difference in analgesic benefit, although aberrant spread to adjacent levels was over twice as likely (38% vs 16%) with the higher volume. At C3, adjacent spread to the TON occurred in 57% of blocks irrespective of volume. Foraminl spread was noted in five instances with 0.5 mL and two instances with 0.25 mL. There were no differences in accuracy rate or specificity between the posterior or lateral approach.

In a small (n=6) cadaveric study that sought to determine the optimal cervical MBB injectate volume, Wahezi and colleagues found that 0.25 and 0.5 mL of contrast and methylene blue fully incorporated the targeted C4–6 medial branch using a fluoroscopically-guided posterior oblique approach. However, the lower injectate volume spread to fewer adjacent structures including intermediate and superficial spinal muscles, surrounding fascia, and terminal branches of the medial branch nerve. They concluded that smaller volumes are more specific and should be used for prognostic MBB before RFA. An earlier study by the same group of authors performed in the lumbar spine found 0.25 mL lumbar MBB to be more specific than 0.5 mL blocks.

Wahezi and colleagues performed a similar study in five cadavers to determine optimal TON block volumes. Using landmarks for injections after partial dissection, the authors found that six of 10 TON blocks using 0.25 mL captured the greater occipital nerve, while increasing the volume to 0.5 mL resulted in 100% of blocks inadvertently anesthetizing the greater occipital nerve. The use of US guidance when performing cervical MBB may enable the operator to visualize the spread of injectate in real time and allow the physician to limit the volume only to that necessary to incorporate the medial branch. Although some earlier US studies used volumes as high as 0.9 mL, subsequent studies have used much smaller volumes (e.g., 0.2–0.3 mL) to accurately target the medial branches from C3–C6.

Cervical IA facet joint injections
The cervical facet joint is a true synovial joint with a reported capacity of 1.0 mL of fluid. There is very limited evidence for short- and long-term analgesic benefits using IA facet joint injections in the cervical spine. In RCTs examining the efficacy of cervical IA injections, a wide range of volumes have been used from as little as 0.5 mL to as high as 2 mL. Volumes greater than 1.0 mL may result in rupture of the joint capsule leading to inadvertent spread to other potential pain generators, thereby undermining specificity.

Several studies have sought to determine the specificity of IA cervical facet injections. In an observational study examining the accuracy of 760 fluoroscopically-guided IA injections performed in 208 patients using 0.2 mL of contrast administered through a 25-gauge needle, Won et al. found that joint overflow occurred in 23.6% of injections, being highest at C5–6 (36.2%) and lowest at C3–4 (3.3%). In a retrospective study examining the spread patterns of 29 patients who underwent a single CT-fluoroscopy-guided cervical IA facet injection performed with 0.5–1 mL of contrast and 1 mL of steroid, Bureau et al. reported IA and retrodural spread in 62% of injections, IA and epidural spread in 7%, and extra-articular spread without IA contrast was observed in 21% of injections. In only 7% of injections was IA spread without contrast extravasation reported. Collectively, these studies suggest that even very low volumes injected into cervical facet joints lack specificity. Yet, using volumes that are too low may result in failure to achieve IA spread leading to a false-negative result, which may be more common in people with facet joint osteoarthritis.

Recommendations
We recommend that cervical MBB volumes be ≤0.3 mL, though slightly higher volumes may be considered if contrast spread fails to capture the most frequent patterns of medial branch innervation; grade C recommendation, low level of certainty. For cervical IA facet joint injection, a total volume not to exceed...
QUESTION 9: DO INTRA-ARTICULAR FACET BLOCKS OR MEDIAL BRANCH BLOCKS CONFERR THERAPEUTIC VALUE?

The cervical facet joints are innervated either by the TON or the medial branch of the dorsal rami that innervate the joints above and below. Medial branch RFA is considered to be a definitive durable analgesic treatment for patients with neck pain arising from the cervical facet joints. However, there is conflicting evidence in the literature about the therapeutic (durable) benefits of MBB or IA with LA and/or steroids.

Efficacy of TON and cervical MBB

Prolonged relief from diagnostic blocks

Diagnostic blocks of cervical medial branches and the TON involve an injection of a short-acting LA such as lidocaine with its effect lasting around 2 hours, or a long-acting LA such as bupivacaine that reliably provides pain relief for 3–8 hours in some studies, but not all. However, prolonged analgesic benefit with LA following these procedures in some patients has been reported, which may obviate the need for medial branch RFA. Bogduk and colleagues injected LA (bupivacaine 0.5%) to block the TON or other cervical medial branches in 24 patients with idiopathic neck pain. Seventeen patients reported pain relief for at least 2 hours with 15 undergoing a repeat block with LA to confirm their response. The two major groups of patients identified in this study were patients with neck pain and headache stemming from the C2–3 joint and those with neck pain and shoulder pain stemming from the C5–6 joint. One patient who had a TON block reported relief of headaches for over a month. In a study performed in 47 patients with neck pain who received dual diagnostic cervical MBB with lidocaine and bupivacaine, 13 patients reported pain relief for a period longer than the duration of action of the LA with one or both LA. Five patients had prolonged pain relief with lidocaine, three patients had prolonged relief with bupivacaine, and five patients reported prolonged benefit with both LA agents.

In one study, 50 patients received double comparative diagnostic blocks with LA and a third placebo injection for neck pain following an MVC. Investigators in the study classified patients based on the duration of pain relief. Among the 14 patients with a concordant response to LA (duration of pain relief consistent with the drug’s pharmacokinetics), three also experienced pain relief with placebo. Two patients had a discordant prolonged response (duration of pain relief prolonged for one or both LA with a longer response to the long-acting LA) and neither had a placebo response. Eleven patients had a discordant prolonged response (duration of pain relief prolonged for one or both of injected LA with a longer response to the short-acting LA), but four of these patients were also placebo responders. There were 23 patients with a discordant (duration of pain relief prolonged for the short-acting LA) or discrepant (pain relief with only one of the two LA injections) response, and 13 of these were placebo responders. It was concluded that a placebo response is more likely if patients who receive dual comparative diagnostic LA blocks have a more prolonged response with the shorter-acting LA or if they report pain relief with only one of the LA. The phenomenon of prolonged analgesic benefit in some patients has also been reported with other (non-cervical medial branch) nerve blocks and in one randomized trial evaluating lumbar medial branch RFA, and may be due to prolonged neural conduction blockade, a reduction in neural inflammation, or reversing central or peripheral sensitization.

Randomized trials evaluating long-term benefit

The impact of adding steroids to LA for TON and cervical MBB was addressed in an RCT that compared the analgesic benefit and duration of action of LA and sarapin to a combination of LA, sarapin and steroids in 120 patients with chronic neck pain who responded to comparative LA blocks. Blocks were repeated over a 2-year period when pain relief returned to more than 50% of baseline. Over 85% of patients obtained ≥50% pain relief over the study duration, with no significant differences between groups. The average duration of relief for each procedure (mean 5.7 blocks over 2 years) was over 4 months in both groups. However, the trial had serious methodological flaws including an enriched enrollment design, failure to control for concurrent interventions, repetitive blocks providing long-term relief without the need for RFA, a high percentage of patients on opioids, and an unclear methodology for evaluating outcomes. More recently, Hussain et al performed a randomized, double-blind study in 60 patients with non-radiating neck pain comparing two-level cervical MBB performed with LA and steroid to trigger point injections with the same solution. Through 12 weeks of follow-up, the MBB group had greater reductions in pain intensity and disability scores than those who received trigger point injections (table 14).

Double-blind placebo-controlled studies evaluating cervical RFA against MBB

There have been three double-blind trials that evaluated outcomes of cervical MBB in the context of a ‘sham-controlled’ study. Van Eerd and colleagues randomized 76 patients with presumptive facetogenic pain based on historical and physical examination findings to receive cervical MBB at three contiguous levels with 0.5 mL bupivacaine and sham RFA, or the same volume of bupivacaine and true RFA. In the RFA group, the mean average neck pain score decreased from 6.8 to 3.6 and 3.8 at 3- and 6-month follow-ups, respectively. In the bupivacaine/sham RFA group, these 3- and 6-month average neck pain scores were 4.3 and 4.5, respectively. The responder rates for ‘significant improvement’ in the RFA group were 57% and 50%, respectively, at the 3- and 6-month time points (vs 51% and 41% at these time points for the bupivacaine MBB/sham RFA group, with no significant difference between groups). The differences in pain reduction and functional improvement favoring the RFA group did not reach statistical significance through 6 months after the procedures, although the benefits statistically and clinically lasted longer (42 months vs 12 months for median benefit).

In a small randomized study by Stovner et al, 12 patients with cervicogenic headache and neck pain were allocated to TON and C3–6 MBB with 1 mL LA (specific LA not noted) plus sham-RFA on the symptomatic side or TON and MBB followed by real RFA. Although dual comparative LA blocks were performed, the results were not used for patient selection. For the primary outcome measure, days per week with moderate or severe pain, 4/6 in the treatment group experienced a positive outcome versus 2/6 in the control group at 3 months post-treatment, with no differences noted at later follow-ups.

In an earlier RCT, Lord and colleagues randomized 24 patients with chronic neck pain attributable to an MVC to receive cervical MBBs and RFA at one or two levels based on complete pain relief following dual comparative LA blocks and a
negative response to placebo injection (ie, 3 blocks), or cervical MBB and sham RFA. The MBBs were performed with 2 mL bupivacaine. The median time for >50% of the pain to return was 8 days in the bupivacaine-only group versus 263 days in the MBB plus RFA group. Among those who received only bupivacaine, three of 12 (25%) experienced at least 50% pain relief at 100 days after treatment and one continued to have pain relief after 200 days.

### Efficacy of cervical IA facet joint injections

Bogduk and Marsland injected LA and steroid into cervical facet joints in eight patients with idiopathic neck pain, finding a median duration of pain relief of 1 month (range 4 days to 12 months). However, repeat IA injection in three patients provided pain relief for only 4–7 days. Similar variability in the duration of relief, with mostly short-term benefit, has been reported with cervical IA facet injections in other studies. In a study that compared the physical and psychological characteristics of responders versus non-responders to cervical IA facet injections, LA and a steroid were injected into the joints of 90 patients. Fifty-eight patients who experienced pain relief following the IA injections and confirmatory MBB were classified as responders. The duration of pain relief in this population varied from 2 hours to just under 3 months. In a prospective cohort study performed in 51 patients (44 with 1-year follow-up) who had cervical facetogenic pain confirmed by dual diagnostic MBB, 24 (54.5%) patients reported analgesic benefit from IA facet injections at 1 year, with 11 patients requiring a repeat procedure within 2–52 weeks. None of the 24 patients required cervical medial branch RFA during the study. In an RCT conducted in 400 patients with myofascial neck and shoulder pain and a positive response to dual cervical IA or MBB, 155 patients with 1-year follow-up received bilateral C5–6 and C6–7 IA injections, while no injections were performed in the other 151 patients with 1-year follow-up. Both groups also received exercise therapy, medications, and trigger point injections. The treatment group reported a greater range of cervical motion, lower pain scores, and a reduction in the incidence of tension-type headaches at 1-year follow-up, although the variable use of co-interventions limits generalization. Finally, in an RCT

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Dory et al 1983</td>
<td>14 pts (22 joints) received IA injections with steroids for neck pain</td>
<td>Prospective cohort study</td>
<td>9 pts had pain relief for 3 days to 13 months</td>
<td>Distension of the joint capsule provoked pain in 50% of pts</td>
</tr>
<tr>
<td>Bogduk et al 1988</td>
<td>Patients with neck pain received cervical medial branch or TON block with LA (n=24) and IA injections with LA and steroids (n=8)</td>
<td>Prospective cohort study</td>
<td>17 of 24 (71%) pts had pain relief for at least 2 hours with TON or MBB. 7 of 7 pts had pain relief with IA injections varying from 4 days to 12 months (median 1 month)</td>
<td>1 pt had relief of headaches for over 1 month</td>
</tr>
<tr>
<td>Barnsley et al 1993</td>
<td>47 pts with chronic neck pain received dual comparative TON or cervical MBB with lidocaine and bupivacaine.</td>
<td>Prospective cohort study</td>
<td>13 of 47 (29%) pts had pain relief lasting longer than expected for either LA; 5 with lidocaine, 3 with bupivacaine, and 5 for both LA agents.</td>
<td>No long-term analgesic benefit in either group; median duration of 50% pain relief was 3 days in both groups</td>
</tr>
<tr>
<td>Barnsley et al 1994</td>
<td>41 pts with chronic neck pain following whiplash injury and a positive response to dual IA diagnostic cervical MBB received IA injections of either bupivacaine or bethemathasone</td>
<td>RCT</td>
<td>No-long-term analgesic benefit in either group; median duration of 50% pain relief was 3 days in both groups</td>
<td>15 of 21 pts in the steroid group and 13 of 20 pts in the LA group had ≥50% pain relief for ≤10 days</td>
</tr>
<tr>
<td>Lord et al 1995</td>
<td>50 pts with chronic neck pain after MVC who received triple comparative TON or cervical MBB with lidocaine, bupivacaine and saline</td>
<td>Prospective cohort study</td>
<td>2 (4%) pts had ‘concordant prolonged’ response (pain relief &gt;7 hours with lidocaine and/or &gt;24 hours with bupivacaine, but longer with bupivacaine)</td>
<td>None of the ‘concordant prolonged’ response pts were placebo responders</td>
</tr>
<tr>
<td>Manchikanti et al 2010</td>
<td>120 pts with neck pain who responded to comparative LA blocks were randomized to cervical MBB with LA and sarapin or with LA, sarapin and steroid</td>
<td>RCT</td>
<td>Similar responder rate (≥50% reduction in pain NRS scores) in both groups (85% for LA and sarapin, 93% for LA, sarapin and steroid)</td>
<td>Average number of treatments was 5.7 in 2 years. Co-interventions not controlled for</td>
</tr>
<tr>
<td>Park et al 2012</td>
<td>400 pts with chronic neck pain secondary to myofascial and facet joint pathology were randomized to receive bilateral C5–6 and C6–7 IA facet injections with LA+steroid and conservative treatment (155 with 1-year follow-up) or conservative treatment alone (151 pts with 1-year follow-up)</td>
<td>RCT</td>
<td>IA injection cohort had increased cervical ROM, greater pain relief, and fewer headaches during the 1-year follow-up</td>
<td>Analgesic medications, trigger point injections with LA and botulinum, and home exercises used variably in both cohorts</td>
</tr>
<tr>
<td>Smith et al 2013</td>
<td>90 pts with WAD grade II ≥6 months post-MVC who received dual IA facet injections and MBB (medications not noted); 30 healthy controls</td>
<td>Cross-sectional study</td>
<td>Similar level of sensory disturbance, motor dysfunction, psychological distress in both groups</td>
<td>No patient experienced pain relief for ≥3 months following IA facet injections</td>
</tr>
<tr>
<td>Lee et al 2018</td>
<td>51 pts (44 with 1-year follow-up) with neck pain and positive dual diagnostic cervical MBB who received IA cervical facet injections with LA and steroid</td>
<td>Prospective cohort study</td>
<td>24 (54.5%) pts reported ≥2-point reduction in pain NRS or ≥50% overall improvement in pain; 9/44 (18%) underwent RFA</td>
<td>11 pts required repeat IA injections in the 1-year study period with a mean interval of 6 weeks between injections</td>
</tr>
<tr>
<td>Hussain et al 2020</td>
<td>60 pts with non-radiating neck pain received cervical MBB at two levels with LA and steroid or trigger point injections with LA and steroid</td>
<td>RCT</td>
<td>Cervical MBB pts had mean pain score of 2.0 at 12 weeks vs 6.96 in trigger point injection group. Functional improvement also greater in MBB group</td>
<td>Technique for MBB and number of trigger point injections not noted</td>
</tr>
</tbody>
</table>

IA, intra-articular injections; LA, local anesthetics; MBB, medial branch block; MVC, motor vehicle collision; NRS, numerical rating scale; pts, patients; RCT, randomized controlled trial; RFA, radiofrequency ablation; ROM, range of motion; TON, third occipital nerve; WAD, whiplash-associated disorder(s).
performed in 41 patients with chronic neck pain following whiplash injury who responded to dual comparative cervical MBB, Barnsley and colleagues compared IA injections of LA to steroids as stand-alone treatments. In both cohorts, a majority of patients (over 65%) experienced 50% or greater pain reduction lasting less than 10 days, with only 10% in each cohort reporting substantial pain relief lasting more than 3 months. In summary, it appears that some patients may have prolonged therapeutic benefit with cervical facet IA injections with LA and/or steroids, with stronger evidence for chronic neck pain in the absence of whiplash injury.

**Recommendations**

We recommend against the routine use of IA injections, although we acknowledge that in patients who may be at risk of adverse consequences from RFA (eg, young athletes, older individuals on anticoagulation therapy, or with implantable cardiac devices) in whom there is a strong likelihood of success (eg, individuals who obtained prolonged relief from previous diagnostic injections with or without steroids), and/or patients who do not have readily available access to cervical medial branch RFA, it may be reasonable to consider IA facet joint injections with steroid (non-particulate at C2–3) in the hope of deriving intermediate-term relief; grade C, low-to-moderate level of certainty. Given the lack of a pathophysiological basis for prolonged relief and the known risks of steroids, the routine use of steroids with cervical MBB should be avoided; grade D recommendation, moderate level of certainty.

**QUESTION 10: SHOULD BILATERAL CERVICAL MBB OR RFA BE PERFORMED DURING THE SAME VISIT? SHOULD THE NUMBER OF LEVELS BLOCKED OR DENERVATED BE LIMITED TO A CERTAIN NUMBER?**

**Clinical anatomy and function of the cervical medial branch nerves**

The cervical facet joints, C2–3 to C7–T1, are innervated by the medial branches derived from the dorsal rami of the cervical spinal nerve roots. These include the TON (to the C2–3 joint), the deep C3 medial branch (to the C3–4 joint), and the C4 to C8 medial branches (C3–4 to C7–T1 facet joints), all of which are targets for cervical MBB or RFA. In addition to giving off articular branches to the facet joints, medial branch nerves also innervate the semispinalis capitis (C2 and 3), multifidi and semispinalis cervicis muscles, and cutaneous areas. The semispinalis capitis is innervated by both medial and lateral branches of the C2 and C3 dorsal rami. Semispinalis capitis, semispinalis cervicis, and multifidus muscles are considered important posterior neck stabilizers. Muscle spindles are present in cervical muscles and the density is higher in the upper region of the neck. Mechanoreceptor endings have been identified in the human cervical facet joint capsules as well. The vestibular system incorporates input from the eyes and cervical proprioceptors when stabilizing head and body positions. Injury of afferent nerves that carry proprioceptive information from these receptors may impair one’s position sense. Compared with lumbar facet joint pain, cervical facet joint pain is more likely to be unilateral, perhaps because of the increased mobility in the neck and since cervical facet joint pain is more likely to result from trauma.

**Indirect evidence**

Data extracted from studies designed for other objectives

Table 15 shows studies denoting the number of facet joints and laterality of treated patients. More procedures were performed on one to two joints than on multiple joints, and a large majority of procedures were unilateral. No sustained complication or side effects were reported that can be attributed to bilateral and/or multi-level facet nerve blocks or RFA, even in cases in which the TON was ablated bilaterally alone or in combination with other medial branches.

Studies by the same group of investigators have sought to determine, through controlled blocks performed one level at a time, the number and distribution of affected joints in patients with chronic neck pain after whiplash. In the most comprehensive of these studies, 31 of 52 patients were diagnosed with cervical facet joint pain, with four patients having two-joint involvement and only one having three-joint involvement. Although elderly patients with advanced osteoarthritis often have multiple levels concomitantly affected, the clinical significance of this in the context of other degenerative changes (eg, uncovertebral joints, cervical discs) is unclear. Performing MBB at multiple segments may lead to the unnecessary treatment of unaffected levels, not only for the initial RFA but for subsequent procedures as well.

**Case reports**

There are two published case reports of dropped head syndrome after cervical medial branch RFA. In one instance, severe progressive cervical kyphosis with inability of active head raising developed after bilateral C2–C3, C3–C4, and C5–C6 facet joint medial branch RFA. The ablations were performed on each side separated by 1 week. Subsequent electromyography (EMG) showed active denervation of the cervical paraspinal muscles and MRI revealed paraspinal muscle atrophy. In the second case, dropped head syndrome developed 3 months after left-sided TON and C3–C4 facet joint medial branch RF denervation with MRI evidence of left semispinalis cervicis and splenius capitis atrophy/regeneration. In neither case was sensory or motor stimulation used. In both cases, passive head extension was not impaired. The authors postulated the etiologies as loss of collateral muscle innervation from bilateral and/or multi-level RFA. It is worth noting that bilateral multi-level facet nerve blocks were performed without reported adverse effects in both cases. Although unpublished, the authors (SPC, RWH) are aware of several cases of temporary ataxia and loss of balance when bilateral upper cervical MBBs were performed.

**Guidelines**

The SIS Practice Guidelines state that cervical MBB, including the TON, can be done bilaterally at the same visit for patients with bilateral neck pain and headache. However, the guidelines advocate staged facet nerve blocks such as blocking one side or only upper or lower segments in different visits in order to isolate the painful joint(s). The guidelines explicitly caution against performing medial branch RFA bilaterally and at numerous levels without careful judgment because of the possibility that treating more than one segment could compromise the function of the cervical musculature. They recommend that bilateral TON ablation be performed on separate occasions after test blocks to establish tolerance for bilateral denervation without causing debilitating ataxia and other untoward side effects. These conservative recommendations are generally consistent with US Medicare coverage determinations which recommend that no more than two levels—unilateral or bilateral—be treated.
Table 15  Studies reporting bilateral and/or multi-level cervical medial branch blocks or radiofrequency ablation

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Number treated</th>
<th>Bilateral treatments (cases)</th>
<th>Number of joints treated (cases)</th>
<th>Adverse events</th>
</tr>
</thead>
<tbody>
<tr>
<td>Lord et al 1996&lt;sup&gt;66&lt;/sup&gt;</td>
<td>Pts with chronic whiplash disorder treated with MBB and medial branch RFA (C2–3 joint was excluded from ablation)</td>
<td>RCT (C2–3 joint was excluded from ablation)</td>
<td>Blocks: 24 pts Ablations:12 pts</td>
<td>All blocks:</td>
<td>Unilateral blocks:</td>
<td>C2–3 (25) 1 (40) 2 (8)</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td></td>
<td>C2–3 plus C5–6 and ipsilateral C6–7 (1) Unilateral C2–3 and contralateral C5–6 (2)</td>
<td>(1) Bilateral blocks:</td>
<td>Vasovagal 2% Postoperative pain 97% Ataxia, special disorientation, unsteadiness 23% Cutaneous numbness C2–3 88% C3–4 80% Lower joints19% Dysesthesia C2–3 56% C3–4 30% Lower joints 17% Transient neuritis 2% Dermoid cyst 1% Köbner’s phenomenon 1%</td>
</tr>
<tr>
<td>Lord et al 1998&lt;sup&gt;66&lt;/sup&gt;</td>
<td>Cervical facet joint pain</td>
<td>Retrospective Ablations:</td>
<td>C2–3 joint 12 pts (25 procedures) Lower cervical facet joints: 28 pts (48 procedures)</td>
<td>Not reported</td>
<td>C2–3 (23) 1 (21) 2 (3)</td>
<td>Numbness 97% Ataxia 95% Dysesthesia 55% (all self-limiting/no intervention needed)</td>
</tr>
<tr>
<td>Govind et al 2003&lt;sup&gt;67&lt;/sup&gt;</td>
<td>Cervicogenic headache Prospective cohort Ablations:</td>
<td>C2–3 ablations (2), each side was treated on separate days</td>
<td>49 pts (51 procedures)</td>
<td>Unilateral C2–3 ablation (47)</td>
<td>8</td>
<td></td>
</tr>
<tr>
<td>Barnsley, 2005&lt;sup&gt;56&lt;/sup&gt;</td>
<td>Chronic neck pain</td>
<td>Retrospective Ablations:</td>
<td>35 pts (47 procedures)</td>
<td>2 joints were treated in 3 cases. It is unclear if they were ipsilateral or contralateral</td>
<td>C2–3 (23) 1 (21) 2 (3)</td>
<td>Nearly all had postoperative pain, lasting 1 week in most patients. 1 case of local wound infection</td>
</tr>
<tr>
<td>Manchikanti, 2006&lt;sup&gt;68&lt;/sup&gt;</td>
<td>Chronic neck pain</td>
<td>RCT Blocks:</td>
<td>60 pts</td>
<td>75% cases</td>
<td>2 (48%) 3 (50%) 4 (2%)</td>
<td>Not reported</td>
</tr>
<tr>
<td>Shin et al 2006&lt;sup&gt;65&lt;/sup&gt;</td>
<td>Chronic neck pain</td>
<td>Retrospective Ablations:</td>
<td>28 pts (6 excluded as only one medial branch nerve of a facet joint was recorded as treated)</td>
<td>8</td>
<td>Unilateral C2–3 (1) 1 (3) 2 (9) 3 (4) Bilateral 1 &amp; 1 (1) 1 &amp; 3 (1) 2 &amp; 1 (1) 2 &amp; 2 (1) 2 &amp; 3 (1) 3 &amp; 3 (1)</td>
<td>Muscle cramping pain and numbness lasting &lt;4 weeks, in a majority of cases, &lt;2 weeks</td>
</tr>
<tr>
<td>Manchikonda et al 2007&lt;sup&gt;68&lt;/sup&gt;</td>
<td>Chronic neck pain</td>
<td>Retrospective Blocks:</td>
<td>251 pts</td>
<td>180</td>
<td>2 (127) 3 (122) 4 (2)</td>
<td>None reported</td>
</tr>
<tr>
<td>Klessinger, 2010&lt;sup&gt;67&lt;/sup&gt;</td>
<td>Pts had previous anterior cervical operations</td>
<td>Retrospective Ablations:</td>
<td>32 pts</td>
<td>10</td>
<td>Unilateral C2–3 (1) 1 (4) 2 (15) 3 (3) Bilateral 1 &amp; 1 (2) 2 &amp; 2 (8)</td>
<td>Not reported</td>
</tr>
<tr>
<td>MacVicar et al 2012&lt;sup&gt;68&lt;/sup&gt;</td>
<td>Chronic neck pain</td>
<td>Prospective cohort Ablations:</td>
<td>104 pts</td>
<td>6</td>
<td>Unilateral C2–3 (29) 1 (54) 2 (16) 3 (5) Bilateral C2–3 (1) 1 &amp; 1 (3) 1 &amp; 2 (2)</td>
<td>Not reported</td>
</tr>
<tr>
<td>Hamer and Purath, 2014&lt;sup&gt;68&lt;/sup&gt;</td>
<td>Cervicogenic headache</td>
<td>Retrospective Ablations:</td>
<td>17 pts C2/3 joint ±C2 DRG</td>
<td>8</td>
<td>Unilateral C2–3 (4) C2–3 &amp; C2 DRG (5) Bilateral C2–3 (4)</td>
<td>Dizziness (1) and suboccipital hyperesthesia (2) (in one unilateral and one bilateral C2/3 ablation case)</td>
</tr>
</tbody>
</table>
per session, with “3-level procedures considered under unique circumstances and with sufficient documentation of medical necessity on appeal”. 139

Recommendations

In summary, indirect evidence suggests bilateral cervical MBB, including the TON, can be performed during the same visit. Although bilateral and multi-segment (>2 levels) RFA have been described, we recommend performing them at separate visits to maximize safety. Given the scant clinical evidence for treating multiple levels and lack of precision, performing MBB at more than two levels simultaneously should be routinely avoided in the absence of compelling clinical evidence to the contrary; grade C recommendation, low level of certainty.

QUESTION 11: ARE FACET, AO, AA BLOCKS ‘DIAGNOSTIC’, ‘PROGNOSTIC’, OR BOTH?

Premise of cervical facet, AO, and AA interventions

The terms ‘diagnostic’, ‘prognostic’, and ‘predictive’ are commonly used interchangeably in the literature on chronic LBP.29 although they are not the same.245 A similar misconception holds true for LA interventions into the cervical spine joints. Portions of the discussion below are substantially derived from our previous lumbar facet intervention working group consensus guidelines.29 The term ‘diagnosis’ refers to the ‘process of identifying a disease, condition, or injury from its signs and symptoms’.246 ‘Prognosis’ most commonly refers to the forecasting of the likely course of a disease (which may include the effects of treatment), while ‘predictive’ provides specific information about the likely effect of a therapeutic intervention. Whereas these terms may overlap in some scenarios, they refer to different concepts.

The patterns and location of the innervation of the cervical spine joints are more complicated than the thoracic and lumbar levels. The cervical spine can be divided into at least five distinct regions (AO, AA, C2–3, C3–4 through C6–7, and C7–T1 joints) based on their unique anatomy. These differences will be referenced where they impact the question of diagnosis and/or prognosis.

Diagnosis

Diagnostic injections can be used to isolate the anatomic structures that are the source of pain.76 They are a critical but potentially imperfect element in the practice of interventional pain and spine medicine. An indirect approach to diagnosis is relied on due to the lack of any pathognomonic historical or physical examination finding, diagnostic test, and/or spine imaging finding indicative of pain of facetogenic or joint origin (ie, a reference standard). Although the administration of LA into the joint itself or onto the nerves supplying the joint with diagnostic intent has face validity, it is based on the assumption that there are no other factors that will alter the pain, and relies on the patient’s report of pain relief, which is subject to bias and cannot be independently verified.247 Therefore, the possibility of false-positive or -negative reporting is an inherent risk with cervical IA or MBB injections.248–249 However, this potentially can be mitigated through placebo injections189 and a thorough understanding of cervical spine anatomy.28 79 187

Prognosis

Prognostic injections can be used in risk stratification and treatment planning. Prognosis is closely aligned in medicine to predictability. A patient may appropriately ask his or her clinician, “What are the chances that I will get 50% pain relief from this intervention?” Similarly, a patient may ask, “What is my prognosis or expectation if I get 70% relief from the diagnostic block and then undergo RF denervation?” These are not easily answered questions. Evidence-based medicine may provide outcome estimates from interventions, but this is not the same as providing a prognosis or prediction.

Limitations of diagnostic injections

The limitation of cervical joint interventions in providing diagnostic information is confounded by the placebo response, which is robust for pain and spine procedures.248 249 Although a placebo response may be observed, this should not be interpreted as a lack of pathophysiologic pain and is a factor in an unknown proportion of patients reporting concordant pain relief after diagnostic blocks.189 Placebo response is discussed at greater length in the lumbar facet guidelines29 and is similarly applicable to those of the cervical spine. Lord et al189 found that comparative diagnostic blockade was highly specific (88%), but only marginally sensitive (54%) in identifying painful cervical facet joints, resulting in potentially labeling 46% of the patients as ‘placebo’ responders and denying them access to RFA. This potentially very high false-negative rate might not be considered acceptable in light of the modest risks of the procedure and the certitude of continued pain and disability for the patient.

The accuracy of a diagnostic block is contingent on several technical and anatomic factors. First, it assumes the procedure

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patient population</th>
<th>Design</th>
<th>Number treated</th>
<th>Bilateral treatments (cases)</th>
<th>Number of joints treated (cases)</th>
<th>Adverse events</th>
</tr>
</thead>
<tbody>
<tr>
<td>Van Eerd et al 201428</td>
<td>Chronic neck pain (excluding whiplash and C2–3 joint involvement)</td>
<td>Retrospective</td>
<td>Ablations: 65 pts</td>
<td>0</td>
<td>2 (65)</td>
<td>Not reported</td>
</tr>
<tr>
<td>Hahn et al 201828</td>
<td>Chronic neck pain and vertigo</td>
<td>Retrospective</td>
<td>Blocks: 178 pts</td>
<td>142</td>
<td>Specific information lacking</td>
<td>Not reported</td>
</tr>
<tr>
<td>Van Eerd et al 202028</td>
<td>Chronic neck pain</td>
<td>RCT</td>
<td>Blocks (39)</td>
<td>76 pts</td>
<td>Unilateral 2 joints per patient (same session)</td>
<td>3 serious adverse events</td>
</tr>
</tbody>
</table>

DRG, dorsal root ganglion; MBB, medial branch block; RCT, randomized controlled trial; RFA, radiofrequency ablation.
is performed in a manner that results in anesthesia of the intended, but not unintended, structure(s). Similar to lumbar MBB, anesthetic injections over the TON and the C3 through C8 medial branches are unlikely to be specific due to the proximity of the cervical spine source of pain relief and restoration of function at 27 weeks following cervical spine intervention. Evaluation of the cervical spine facet joints for diagnostic or prognostic purposes has been well described for lumbar spine facet joints. Although cervical spine facet joint blocks are generally safe and efficacious, they are characterized by high technical failure rates; grade C recommendations are denoted as “limited.”

**Evidence for diagnostic and prognostic utility**

The prevalence of cervical spine joint pain has a range of 26–60% when using history, physical examination, and radiological imaging for diagnosis. In light of the lack of objective measures of cervical spine joint-mediated pain, IA and MBB injections with LA remain the most widely accepted approach to diagnosis and a surrogate measure for prognosis. An IA injection with LA can serve as a diagnostic tool for a subsequent therapeutic joint injection performed with a steroid into the AO and AA joints. In a small prospective observational study, 18 of 20 patients with a positive response to an IA AO injection with LA experienced >2-point reduction in their pain score on a 0–10 visual analog scale 2 months following a therapeutic IA injection with LA and steroid. In one arm of an RCT, seven of 11 patients who had a positive IA AO with LA experienced at least 50% reduction in their pain 6 months after an AO injection with steroid. There are a lack of high-quality studies to address this question for the AA joint; a single study provides an incomplete answer. In a retrospective study, 26 of 32 patients who underwent lateral AA joint injections experienced ≥50% pain reduction post-procedure, with 15 reporting no pain. Unfortunately, the results did not differentiate characteristics of the subgroup who received relief immediately after the injection from those who did not achieve relief in the longitudinal results. Other studies used unreliable screening criteria for AA therapeutic injections leading to inconclusive results.

Several studies have examined the prognostic utility of C3 through C7 medial branch interventions; however, high-quality evidence for the utility of C2–3 through C7–T1 IA injections is lacking. In the rigorous RCT by Lord et al using placebo-controlled MBB for patient selection, 58% (7/12) of the patients experienced complete pain relief and restoration of function at 27 weeks following cervical medial branch RFA. A follow-up to this study with additional patients reported a similar success rate. A larger RCT using a similarly rigorous protocol as Lord et al included the C2–3 joint along with the lower cervical facet joints. Twenty-one of 35 patients in this study received complete relief from RFA at 12 weeks.

**Recommendations**

C3 through C8 MBB meet most criteria as a diagnostic intervention for cervical joint-mediated pain, although the nerves that innervate the facet joints innervate other potential pain-generating structures. Technically sound IA joint injections theoretically meet criteria as a diagnostic intervention for cervical joint-mediated pain, although they are characterized by high technical failure rates; grade C recommendation, low-to-moderate level of certainty. IA injections are less predictive than MBB for response to medial branch RFA for the C2–3 through C7–T1 joints; grade C recommendation, low level of certainty. Although accuracy may be improved with CT guidance or arthrography, these tools are not well supported in peer-reviewed investigations. IA injections of the AO and AA joints with LA may be diagnostic and provide predictive information for IA steroid injections; grade C recommendation, low level of certainty.

**Question 12: What is the effect of sedation on the accuracy of diagnostic or prognostic intra-articular facet joint blocks and MBB?**

**Arguments for and against sedation during diagnostic blocks**

The general arguments regarding the use of sedation during diagnostic blocks were eloquently outlined in the lumbar facet guidelines. Proponents assert that sedation allays anxiety and reduces procedure-related pain (thereby minimizing the likelihood of a false-negative block), enhances patient satisfaction (thereby reducing the chance of subsequent no-shows), and may prevent movement, thereby facilitating performance. Anxiety

**Limitations of prognostic injections**

IA injections and MBB have been used to assess the probable response of medial branch RFA. The rationale is that, if an MBB relieves pain, then a treatment capable of interrupting conduction along the same nerve(s) should provide comparable relief, but last longer depending on the expected duration of disrupted signaling (ie, until nerve regeneration or reinnervation occurs). The use of these interventions as a surrogate measure for outcome prediction carries the same limitations as their use as a diagnostic tool. The accuracy of a prognostic test also depends on how success or failure is defined. For instance, the success of RFA might be expressed simply as pain relief at the point in time at which the LA is no longer active or it could be defined as alleviation of pain at some pre-designated distant time point (months) following the procedure. Depending on the time chosen, the prognostic power of the initial intervention will be different.
has also been shown to lower pain perception thresholds and tolerance in experimental studies. Critics argue that sedation increases the rate of false-positive blocks, increases risks and costs, and may interfere with physician–patient communication and engagement in post-procedure activities, which can interfere with the interpretation of post-block pain relief. Along with analgesics such as opioids and ketamine, benzodiazepines may also promote pain relief by virtue of their muscle relaxant properties, as a myofascial component is present in over 90% of patients with chronic axial neck pain. Estimates on the economic costs of sedation for interventional pain procedures exceed $300 million per year in the USA.

**Differences between the cervical spine and lumbar spine**

Unlike lumbar MBBs, cervical MBB procedures can be performed in the lateral or prone position. For cervical MBB, the lateral approach involves a shorter distance between skin insertion and the target medial branches and has been shown in a randomized trial to provide comparable accuracy and pain relief. Depending on the approach, cervical MBB may involve less tissue damage and consequent pain than lumbar MBB. However, a randomized trial comparing cervical and lumbar epidural steroid injections found a higher proportion of cervical patients requested additional LA, suggesting possible patho-anatomical differences. Compared with cervical epidural steroid injections (ESIs) whereby patient movement can result in neurological complications, MBBS are less subject to movement-related procedural complications.

Most studies have found a significantly higher incidence of vasovagal reactions with cervical injections compared with lumbar injections, and sedation has been shown to reduce the incidence in those who are predisposed. In a large retrospective study by Kennedy et al. that evaluated the incidence of vasovagal reactions during a host of spinal injections including cervical MBB and cervical medial branch RFA, the authors found that light sedation decreased the rate both in individuals without a prior vasovagal event compared with no sedation (0% vs 3%) and in those with a prior history of a vasovagal reaction (0% vs 23%). A history of an anxiety disorder is widely considered an indication for sedation, and there is a high co-prevalence rate in individuals with both neck and back pain, with one systematic review and meta-analysis reporting an anxiety risk ratio of 3.29 (95% CI 2.16 to 5.00) for neck pain. In a large cross-sectional study evaluating the incidence of anxiety in neck and back pain in 1580 adolescents, anxiety and depression scores on the Youth Self Report scale of the Child Behavior Check List were associated with a slightly higher association for neck pain than back pain with an OR of 1.43 (95% CI 1.20 to 1.70) versus 1.38 (95% CI 1.15 to 1.66) (table 16).

**Clinical trials**

**Cervical facet blocks**

A randomized study by Manchikanti et al. allocated 180 patients who had previously responded to comparative LA cervical MBB and were undergoing repeat procedures to receive midazolam 1–5 mg, fentanyl 50–250 μg, or up to 5 mL of saline titrated to effect. Pain scores were measured before medication administration and shortly thereafter, before performing the actual MBB. Using 80% pain relief as the threshold, the authors found that 5% of saline patients and 8% of both midazolam and fentanyl patients experienced significant pain relief before the MBB. Using 50% pain relief as the cut-off threshold for a positive response, the authors reported that 8%, 13%, and 27% of patients who received saline, midazolam, and fentanyl were responders, respectively (p<0.05 between 50% and 80% relief only for fentanyl). In a follow-up randomized study performed by the same group of investigators evaluating the effect of sedation on pain relief prior to cervical and lumbar MBB in 60 patients, 30 of whom had neck pain, the authors reported ≥50% pain relief in 5% of saline patients, and in 15% of midazolam and fentanyl patients after medication administration. Using an 80% cut-off threshold resulted in 5% saline responders and 10% midazolam and fentanyl responders. These studies did not address the question of the effect of sedation on the outcome of MBBS because, in each study, the authors evaluated the effect of medication administration before the MBB was performed. Flaws in these studies also include the high percentage of patients on opioids, the relatively high percentage with prior surgery, and that these blocks were being done for therapeutic and not diagnostic purposes.

**Extrapolated evidence on the positive rate from other prognostic blocks**

The most methodologically sound study to examine the effect of sedation was a randomized crossover study by Cohen et al. performed in 73 people who received two sympathetic or sacroiliac joint blocks, with sedation and no-sedation given in random order. Midazolam, with or without fentanyl, was titrated to effect by a board-certified anesthesiologist. In the main crossover analysis, procedures performed with light sedation were associated with a greater than two-fold increase in a positive block based...
on pain diary assessment using \( \geq 50% \) pain relief as the cut-off, and a three-fold increase using 80% as the cut-off threshold. Similar increases in the rate of positive diagnostic blocks were noted for the parallel group and omnibus (all sedation vs all no-sedation) analyses. Whereas sedation reduced pain from procedures, it did not affect satisfaction scores or 1 month outcomes. Paradoxically, statistically significant differences were noted between the no-sedation group and those who received low-dose midazolam only (\( \leq 4 \) mg) or light sedation with both midazolam and fentanyl (\( \leq 4 \) mL of midazolam 1 mg/mL and/or fentanyl 50 \( \mu \)g/mL), but not those who received heavy sedation (\( >4 \) mL of midazolam and/or fentanyl). Although not used diagnostically, a multicenter prospective study by Dreyfuss et al\(^{264}\) found no difference in immediate post-procedure pain scores after lumbar, thoracic, and cervical ESIs performed in 102 patients.

Effect of sedation during prognostic blocks on treatment outcomes

Several retrospective studies have evaluated the effectiveness of sedation during diagnostic blocks on the results of therapeutic interventions. A retrospective study that sought to identify predictive factors associated with celiac plexus neurolysis in 50 patients with cancer pain reported a 73% success rate in people who underwent prognostic blocks without sedation versus 39% in those who received sedation during celiac plexus blocks.\(^{265}\) However, in a large retrospective study evaluating outcome predictors in 265 patients who underwent genicular nerve RFA, the authors found no outcome differences stratified by whether or not sedation was used during the diagnostic injections.\(^{266}\)

Patient preference for sedation

Two studies performed in the same private practice setting reported disparate results on the necessity of sedation before lumbar, thoracic, and cervical ESIs and facet blocks.\(^{267,268}\) In the survey study by Cucuzzella et al\(^{267}\) the authors found that 17% of 500 patients requested oral sedation before their procedure, with 28% reporting they would request it before a second injection. Surprisingly, those who were sedated reported significantly higher procedure-related pain scores than those who were not sedated on the first injection (\( p=0.12 \)) and significantly higher pain scores on the second injection. This may be attributable to those patients with anxiety and anticipating higher procedure-related pain being more likely to request sedation. This is supported by the observation that a history of anxiety, depression, and higher procedure-related pain were associated with requesting sedation. In a prospective follow-up study in which 301 patients were offered oral or IV sedation, 58% requested sedation, with 90% of these individuals being satisfied with the anxiolytic effect.\(^{268}\) For patients not requesting sedation, 93% were satisfied.

Guidelines

Several guidelines have been published on the use of sedation for interventional procedures and have generally recommended against routine sedation for simple blocks such as facet blocks and ESIs based on the rationale that the risks outweigh the benefits. These include the ASA, SIS, ASRA, and the lumbar facet guidelines committee,\(^{269-271}\) with ASIPP recommending the avoidance of opioids for diagnostic facet blocks but allowing for the use of sedation with midazolam and opioids to alleviate pain and anxiety for therapeutic procedures.\(^{272}\)

Recommendations

We recommend performing cervical MBB without sedation, including using anxiolytics (benzodiazepines, propofol) or analgesics (opioids, ketamine) as there is evidence it may increase the false-positive rate. Patients in whom light sedation may be considered include those with pre-existing psychiatric conditions that may include anxiety and post-traumatic stress, and for procedures in which the blocks are expected to be particularly painful (eg, obesity, those with anatomic derangements); grade B recommendation, moderate degree of certainty.

QUESTION 13: WHAT SHOULD THE CUT-OFF BE (PERCENT RELIEF) FOR DESIGNATING A BLOCK AS ‘POSITIVE’ AND IS THERE ANY BENEFIT IN USING NON-PAIN SCORE OUTCOME MEASURES?

Guidelines and basis for recommendation

The multi-organizational lumbar facet guidelines advocate using 50% pain relief as the cut-off for selecting patients for RFA, citing the rarity of isolated lumbar facet joint pain, maximizing access to treatment, and the absence of reliable treatment alternatives as the main reasons.\(^{29}\) In their 2013 guidelines, SIS advocates using ‘complete relief of pain in the topographical region targeted’ as a selection criterion.\(^{18}\) In a 2012 systematic review, ASIPP noted there was stronger evidence for the use of \( \geq 75% \) pain relief with double blocks than there was for single blocks, or double blocks using lower thresholds.\(^{22}\) The recommendations by the lumbar facet guidelines group were based on numerous retrospective studies showing no difference in outcomes when cut-offs were stratified by 50% and 80% thresholds\(^{28}\) and a prospective study demonstrating no differences in 3-month outcomes between pain relief from single diagnostic blocks stratified by 10% increments.\(^{272}\) Two studies included in those guidelines did report non-statistically significant higher success rates for higher cut-off thresholds for lumbar medial branch RFA. Manchikanti et al\(^{273}\) reported 1-year success rates of 75% in individuals who obtained between 50% and 79% pain relief from prognostic blocks, and 93% in those who obtained at least 80% relief, with some patients receiving serial MBB and others undergoing RFA. A retrospective study by Derby and colleagues\(^{274}\) reported a 54% success rate at 6 months in patients who experienced between 50% and 79% relief on single or double lumbar MBB versus 84% in those who obtained at least 80% relief. However, the high RFA success rates in those who experienced less than 80% pain relief strongly support the consideration of using less rigorous cut-offs.

Differences between the lumbar and cervical spine

It is generally acknowledged that the prevalence of facet joint pain is higher in the neck than in the low back in individuals with axial complaints, which is a consequence of greater stress on the cervical than lumbar joints during spine movements, and the relatively greater size compared with the intervertebral discs.\(^{27-275}\) This should theoretically lead to greater pain relief during diagnostic facet blocks. But whereas radiological facet joint degeneration in the absence of disc degeneration is rarer in the lumbar spine than the cervical spine, the overall prevalence of degeneration is similar in the two regions.\(^{276}\) The proportion of individuals with cervical zygapophysial joint disease is slightly higher than those with cervical disc degeneration, with one study finding 53% of individuals with cervical facet degeneration versus 43% with disc degeneration.\(^{276}\) There is some evidence that the density of nociceptors in the facet joint capsule and bone and mechanoreceptor firing in response to stimuli may be higher in the cervical spine than the lumbar spine.\(^{239,277}\) In one observational study involving 56 patients with chronic neck pain, Bogduk and April found that 23 patients (41.1%)
exhibited both a positive MBB, defined as complete pain relief lasting for the duration of action of the local anesthetic, and a positive discogram at the same level, indicating that neither the cervical facet joints nor intervertebral discs were likely to be the only source of pain, or that one or both tests are characterized by a high false-positive rate. Studies have also found electromyographic differences between neck pain and control patients and a very high prevalence rate of trigger points in individuals with neck pain.

### Identifying meaningful cut-off thresholds

Studies vary regarding RFA outcomes based on MBB cut-off thresholds in the cervical spine. Commonly cited cut-off values are discordant in cervical facet literature and are sometimes inferred from lumbar and thoracic facet studies. Cut-off thresholds commonly cited include 50%, 75%, 80%, or 100% relief following cervical diagnostic MBB, with no consensus in the literature or pain community regarding which cut-off leads to the best outcomes. Initiative on Methods, Measurement, and Pain Assessment in Clinical Trials (IMMPACT) guidelines note that a 30% or a 2-point reduction in pain signifies clinically meaningful benefit, and the lumbar facet guidelines committee agreed with this threshold. Importantly, they also noted that quality of life changes should be considered as part of an individualized approach to identifying outcomes that individual patients consider important.

Two points or 30% pain relief is also commonly used to designate responders in federally sponsored studies and late phase clinical trials, including those evaluating interventions for cervical pain and RFA.

### Studies stratifying cervical RFA outcomes by prognostic block cut-off thresholds

One might argue that there should theoretically be some correlation between pain relief after cervical MBB and pain relief after RFA, but that numerous other factors might function to confound this correlation (eg, technique, number, and types of diagnostic blocks) and make it difficult to detect a signal. This is suggested by a recent systematic review. Despite the general recommendations for less than 50% pain relief to be considered clinically meaningful, many RFA studies maintain higher cut-off thresholds from MBB. Determination of a successful MBB can help the clinician decide which patients would most benefit from RFA; however, few studies have evaluated cervical medial branch RFA outcomes stratified by cut-off threshold. In the earliest study examining this question, Cohen et al evaluated factors associated with cervical medial branch RFA outcomes in 92 patients from three treatment centers. They reported a 56% success rate in those who obtained 50–79% pain relief versus 58% in those who experienced ≥80% relief. In this study, ≥50% pain relief lasting at least 6 months post-treatment was defined as a positive outcome. Other studies that examined cervical medial branch RFA outcomes have also failed to find a difference in results broken down by cut-off threshold.

In a cross-sectional study evaluating the effect of pain relief cut-off thresholds following dual MBB, Burnham et al found no difference in symptom relief (defined both as ≥50% pain reduction and 2-point or greater decrease on a 0–10 scale after RFA) between patients who had reported 80–99% symptom relief and those who experienced 100% symptom relief with dual MBBs. Holz and Sehgal evaluated lumbar and cervical medial branch RFA outcomes in 112 patients stratified by the results of diagnostic blocks, with a positive block designated as at least 70% pain relief. The authors found no correlation between the amount of pain relief after MBB and RFA outcomes. They did not perform a subgroup analysis based on the area of treatment. In both of these studies, the cut-off thresholds were determined based on payer requirements.

In an observational study that followed 28 patients with chronic neck pain who received dual comparative blocks using >50% pain relief as the cut-off threshold, Shin et al found no correlation between pain relief after prognostic blocks and cervical medial branch RFA outcomes at 6-, 9-, and 12-month follow-up. For the cervical MBB, patients were stratified based on 50%, 75%, 80%, and 100% relief cut-off thresholds. For RFA, percent pain relief was measured as a continuum. Nineteen (68%) patients achieved a positive RFA outcome (≥50% pain relief) at 6-month follow-up, with eight (29%) achieving complete pain relief (table 17).

### Reasons for discordance

Due to the distinct nature of MBB and RFA from a procedural standpoint, it should be acknowledged that pain relief is likely going to be greater for MBB than for RFA. In the cervical and lumbar spinal regions, respectively, Cohen et al and Dreyfuss et al compared pain relief after cervical MBB and cervical RFA at 6-, 9-, and 12-month follow-up. For cervical MBB, patients were stratified based on 50%, 75%, 80%, and 100% relief cut-off thresholds. For RFA, percent pain relief was measured as a continuum. Nineteen (68%) patients achieved a positive RFA outcome (≥50% pain relief) at 6-month follow-up, with eight (29%) achieving complete pain relief (table 17).

### Table 17: Cervical medial branch blocks or IA cut-off correlation with RFA outcomes

<table>
<thead>
<tr>
<th>Study</th>
<th>Patient population</th>
<th>Design</th>
<th>Results</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cohen et al 2007</td>
<td>92 pts with chronic neck pain from three treatment centers who underwent RFA after a positive response to diagnostic cervical LA blocks. Positive response was defined as ≥50% pain relief lasting at least 6 months</td>
<td>Retrospective study</td>
<td>56% of pts who had at least 50% but less than 80% relief from diagnostic blocks had a successful RFA procedure compared with 58% who experienced at least 80% relief from MBB</td>
<td>Multicenter study using single blocks, evaluating cervical facet RFA</td>
</tr>
<tr>
<td>Burnham et al 2019</td>
<td>50 pts who received 80–99% vs 100% pain relief after dual cervical MBB</td>
<td>Cross-sectional cohort study</td>
<td>54% of pts in both groups reported ≥50% pain reduction</td>
<td>87 records were screened and 50 pts were included in the study. Follow-up was conducted by phone call at various time intervals after 6 months</td>
</tr>
<tr>
<td>Holz and Sehgal 2016</td>
<td>112 pts with positive analgesic response to dual comparative MBB</td>
<td>Retrospective chart review</td>
<td>Percent pain relief after RFA was 48%. Overall analysis showed no correlation between percent or duration of pain relief after MBB and pain relief after RFA</td>
<td>28% underwent cervical facet interventions. Highest pain relief was in individuals who achieved 100% pain relief lasting &gt;8 hours with lidocaine, suggesting a strong placebo response</td>
</tr>
<tr>
<td>Shin et al 2006</td>
<td>28 pts with positive analgesic response to dual comparative MBB</td>
<td>Observational</td>
<td>No correlation between categorical pain relief on prognostic blocks (25%, 50%, 75%, 80% and 100%) and pain relief after RFA</td>
<td>5 pts had whiplash injury and 1 had penetrating neck trauma. Did not target TON</td>
</tr>
</tbody>
</table>

IA, intra-articular; RFA, radiofrequency ablation; MBB, medial branch block; TON, third occipital nerve.

Non-pain measures

Pain should always be evaluated in context, as a change in activity levels, anxiety, or analgesic usage can significantly affect pain intensity. In patients with chronic pain, improving function may be a more meaningful and realistic benchmark than pain relief with several high-impact spinal intervention studies using disability as their primary endpoint. For relief of acute pain, many experts advocate using the utilization of rescue analgesic medications as an appropriate benchmark for success. There is also an increased interest in pain biomarkers that may help guide diagnosis and response to treatment; however, the research and application of this information is still in its infancy. In one study performed in the lumbar spine, Cohen et al. found that while a decrease in diastolic blood pressure after MBB was significantly associated with a higher RFA success rate, the low sensitivity and negative predictive value precluded its use as a solitary screening tool. In clinical practice, the parameters assessed in validated disability instruments (eg, sleep, driving, activities of daily living, travel) preclude their use during the brief 2–6 hour diagnostic window after cervical facet blocks. The same is true for analgesic consumption, as long-acting medications cannot be tapered over a short period and withholding opioids may precipitate hyperalgesia. Future studies should explore whether ultra-long-acting formulations of LA (eg, liposomal delivery systems) provide better predictive value than plain LA.

Clinical trials evaluating cervical medial branch RFA after whiplash

This committee considered differences in patient populations with facet pain due to degenerative and arthritic changes as opposed to younger patients who may have facet pain secondary to an isolated mechanism of injury (ie, whiplash). Studies have found that the cervical facet joints are responsible for 25–65% of non-traumatic and 50–60% of trauma-induced cases of chronic neck pain. Given the higher prevalence, there should theoretically be a lower false-positive block rate in individuals with whiplash and older patients with osteoarthritis.

There are no direct comparisons of facet block pain relief cut-off thresholds in whiplash patients, but an indirect comparison of RFA outcomes does not reveal any significant differences between using 50%, 80%, and 100% pain relief cutoffs. Whereas the literature varies on recommendations for cut-offs in patients with whiplash-induced facet joint pain, a conjunctural argument might be made for a higher cut-off given the lower likelihood of confusing pain generators, the higher risk for neuritis at upper cervical levels, and the younger age (ie, greater impairment from muscle denervation) compared with those with facet joint osteoarthritis.

Recommendations

Given the strong evidence that <50% pain relief may be clinically meaningful and the absence of direct evidence that using higher MBB cut-off thresholds results in higher RFA success rates, this committee recommends that ≥50% reduction in pain be considered a positive prognostic block. There are theoretical underpinnings that in younger individuals (<45 years of age) with whiplash injuries who are having C2–3 and or C3–4 treated, higher cut-offs should be used. Although deductive reasoning indicates that greater pain relief with MBB should lead to greater pain relief after RFA, the literature on the cervical spine does not support this; grade C recommendation, low-to-moderate level of certainty. At present, non-pain measures such as activity level should not be used as the sole criterion to determine the success or failure of a prognostic block, but may be used in conjunction with pain assessment; grade B recommendation, moderate level of certainty.

QUESTION 14: HOW MANY PROGNOSTIC BLOCKS SHOULD ONE PERFORM BEFORE RFA?

The utility of screening blocks depends on numerous factors such as the positive and negative predictive value, the diagnostic validity, and the relative risks and costs compared with the definitive treatment. These factors are in turn contingent on myriad other variables such as patient selection (the predominant predictor of outcome), definitions (ie, threshold for a positive block, benchmark for measuring success), prevalence rate (ie, pre-test probability and false-positive rate), and technique (eg, large electrodes may increase the chance of nerve capture, lower block volumes increase specificity, and higher precision may reduce false-negative blocks). Screening tests have greater benefit when there is:

- High positive predictive value.
- A low prevalence rate for the index condition.
- High false-positive and low false-negative rates.
- The definitive procedure carries significant risks and costs compared with the screening test.

Cervical versus lumbar spine

The rationale for performing more than one diagnostic block is to reduce the false-positive rate and improve the success rate for medial branch RFA. In the cervical spine, the reported false-positive rates for uncontrolled MBB range from 36% to 55% based on dual block paradigms, although rates approaching 50% are unlikely to be accurate given the relative prevalence rate of 25–66% in individuals with axial neck pain (ie, the sum of true and false-positives, and true and false-negatives must equal 100%). In double-block studies, it is assumed that a negative second block means the first block was falsely positive, though an equally plausible interpretation in the cervical spine is that the second block was falsely negative. Among patients with pain following whiplash, one placebo-controlled trial that used three blocks reported a prevalence rate of 60%, with 50% of patients with headache having a symptomatic C2–3 joint. The prevalence (true positive rate) of cervical facet joint pain among individuals with axial neck pain is higher than the prevalence of facet joint pain in patients with chronic LBP owing to greater mobility in the neck, a possibly higher density of nociceptors, and the increased relative size of the cervical facet joints to discs; this should theoretically translate to a lower false-positive rate.

The effectiveness of cervical medial branch RFA has been less studied than in the lumbar spine, but there is no evidence...
Studies have also reported high success rates for cervical medial branch RFA without the use of prognostic blocks. In describing their results using a new posterior oblique approach for cervical facet RFA, van Eerd et al. reported that 51% of 65 patients were much or very much improved at 2 months. A later double-blind randomized trial performed in 76 patients with axial neck pain reported ≥50% success rates based on meaningful reduction in pain scores in both the RFA group (56%) and the LA block-sham RFA group (51%) at 6-month follow-up.68 Earlier, in a randomized trial by Stovner et al. that failed to enroll their intended sample size, 12 patients were allocated to true and sham denervation of the medial branches innervating the C2–6 facet joints. Although patients underwent medial branch and occipital nerve blocks before treatment, the results were not used to select participants. At 3 months post-treatment, four of six in the treatment groups experienced clinically meaningful benefit versus two of six in the sham group, with the differences disappearing at later time points. The reasons why cervical facet joint RFA may be more effective than lumbar facet joint RFA include a higher population prevalence/lower rate of false-negative blocks, increased ability to place electrodes parallel to the target nerve, less variability in nerve location, and possibly a lower co-prevalence rate of concomitant pathology.

False-negative blocks

The rationale for the lumbar facet guidelines to recommend single blocks was based in large part on the incidence of false-negative blocks of the lumbar medial branch nerves, which were estimated to be 47% in a retrospective study by Derby et al. using either 50% (46.7%) or 70% (47.1%) cut-off thresholds. In this study, among the eight patients with initial false-negative blocks who underwent lumbar medial branch RFA after a positive subsequent block, six (75%) experienced a positive outcome. However, for reasons outlined above, the extrapolation of studies performed in the lumbar spine to the cervical spine has drawbacks. In an elegant controlled study by Lord et al. in 50 patients with whiplash injury, the authors performed LA blocks with lidocaine, bupivacaine, and saline in random order. Based solely on the response to comparative LA blocks which required longer ‘complete’ or ‘profound’ relief with bupivacaine than lidocaine, 34 individuals would have been classified as negative. However, when the authors reclassified a positive response as reproducible relief with both active agents but no relief with placebo, 11 (32.4%) were considered to be false-negative. Finally, Cohen et al. performed low-volume (<0.5 mL) cervical MBB under fluoroscopy, after which patients were sent for CT scans to assess accuracy. In six of 86 blocks the contrast was noted to miss the medial branches despite appropriate needle placement, for a potential false-negative rate of 7.0% (table 18).

In clinical practice and to reduce the number of procedures patients must endure, multiple joints are typically blocked at the same time which can lead to the treatment of an inappropriate level(s) (eg, a positive response to C4–5 and C5–6 MBBs in a patient with only C5–6 pain) but is less likely to result in a false-negative response. Conversely, anesthetizing only a single joint at a time may place an unreasonable burden on patients and healthcare resources, and lead to patients withdrawing from treatment before painful joints are identified. The use of pain referral patterns and the elicitation of tenderness under fluoroscopic guidance have been used in clinical trials to select levels in patients with cervical facetogenic pain, but secondary hyperalgesia and overlap in referral patterns limit their utility as a reference standard.

**Studies comparing different numbers of blocks before RFA**

The lumbar facet guidelines recommended single blocks based on high false-negative rates for lumbar facet blocks, which was based predominantly on indirect comparisons of outcomes in studies using single or multiple blocks, and retrospective studies comparing outcomes stratified by the number of blocks.29 Whereas the former suggests that using multiple blocks increases the lumbar medial branch RFA success rate, retrospective studies comparing single versus double blocks before RFA have mostly shown no or small, non-statistically significant differences (63.2% success rate with mean relief of 76.7% in 38 patients who underwent a single block vs 84.6% success rate with mean relief of 71.1% in 13 patients who underwent double blocks)274 in outcomes. In the only RCT evaluating the effect of the number of prognostic blocks on lumbar medial branch RFA outcomes in 151 patients with suspected lumbar facetogenic pain, Cohen et al. found the highest overall success rate when proceeding straight to RFA without blocks, but the highest RFA success rate (ie, considering only those who proceeded to RFA) when double blocks were employed. This predictable outcome was attributed to the inclusion of all placebo responders in the 0-block group, and the exclusion of all potential false-negatives. In the cost-effectiveness analysis, the 0-block paradigm was associated with the lowest cost per successful treatment and also lowest overall costs. The RFA to MBB reimbursement ratio for most payers is comparable in the cervical and lumbar spine, enhancing generalizability. Similar to the comments on false-negative blocks, the extent to which studies performed in the lumbar spine can be generalized to the cervical spine is unclear.

**Dual blocks**

The lumbar facet guidelines acknowledge that enhancing selection rigor, including by requiring more positive blocks, would likely increase the success rate, although at the expense of witholding treatment from individuals who might potentially benefit from RFA.29 Although strong data are lacking and a higher prevalence rate (pre-block probability) of facet joint pain in the neck may attenuate this effect, this may also be the case for cervical medial branch RFA. MacVicar et al. reported a 66% success rate, defined as 100% pain relief, in 104 patients using dual blocks to select patients for RFA, but also employed multiple lesions with 16-gauge electrodes and required complete pain relief during diagnostic blocks as a selection criterion. The authors did not note how many patients were screened, but in a similar study using nearly identical selection criteria performed in the lumbar spine in which 106 patients were treated, the same authors estimated that 575 were evaluated for treatment. In a randomized placebo-controlled trial evaluating cervical medial branch RFA in 24 whiplash patients, Lord et al. reported ≥3 months pain relief in seven of 12 RFA patients, with the median duration of pain relief being 263 days. They defined success as complete relief of their index neck pain, although some patients continued to have headache. In addition to performing three diagnostic blocks including one with saline, the authors required 100% pain relief during the diagnostic blocks and created multiple lesions per level. They screened 54 patients to obtain their enrollment numbers. Although these results suggest that multiple blocks may result in higher RFA success rates, these
studies also employed more stringent diagnostic criteria and rigorous treatment parameters, making it difficult to separate out the effect of the dual blocks.

**Other randomized studies evaluating the utility of prognostic blocks**

In a randomized trial performed in 54 patients with knee osteoarthritis comparing genicular nerve RFA outcomes based on 0-blocks (ie, moving straight to RFA) to proceeding to RFA only after a positive prognostic block, McCormick et al. found no significant difference between the groups (64%) success rate in the 0-block group vs 59% in the single-block group. Extrapolation of this finding to the cervical spine is limited by the differences in anatomy (including the multiple nerves providing nociceptive innervation to the knee joint) and that the diagnosis of knee osteoarthritis does not require diagnostic blocks, or even x-rays.

**Third occipital nerve (TON) blocks**

Most, but not all studies evaluating cervical medial branch RFA have included the TON, so it is challenging to evaluate the evidence behind targeting the TON by itself. The pain referral pattern for TON also overlaps with that for upper cervical facet joints, making it difficult to distinguish the effect of TON RFA from that of the C3–4 and C4–5 joints in studies that evaluated multi-level RFA. In a retrospective study evaluating the effects of C2 dorsal root ganglion and/or TON RFA in 40 patients with refractory cervicogenic headache based on a single diagnostic block requiring at least 50% pain relief, 70% of patients reported at least 80% pain relief lasting a mean duration of 4.5 months.

**Table 18: Studies evaluating the false-positive and false-negative rates of cervical MBBs**

<table>
<thead>
<tr>
<th>Study</th>
<th>No of subjects</th>
<th>Methodology</th>
<th>Prevalence</th>
<th>False-positive/ negative rate</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Barnsley et al 1993</td>
<td>55 pts (60 joints)</td>
<td>Near-complete relief with lidocaine and bupivacaine MBB, with the duration of relief lasting longer with bupivacaine</td>
<td>73%</td>
<td>FP rate 27% (2 pts had relief with only 1 block while 14 had longer relief with lidocaine)</td>
<td>All pts had at least one positive block (no negatives). All pts had whiplash after an MVC.</td>
</tr>
<tr>
<td>Barnsley et al 1993</td>
<td>47 pts</td>
<td>Near-complete relief with lidocaine and bupivacaine MBB, with the duration of relief lasting longer with bupivacaine but not longer than the expected duration (7 hours for lidocaine, 24 for bupivacaine)</td>
<td>57%</td>
<td>FP rate 40% (1 of 45 pts had no relief after second block while 17 had discordant or prolonged relief)</td>
<td>All pts had whiplash after an MVC. Only 2 of 47 pts obtained insufficient relief from initial block.</td>
</tr>
<tr>
<td>Barnsley et al 1995</td>
<td>50 pts</td>
<td>Near-complete relief with lidocaine and bupivacaine MBB, with the duration of relief lasting longer with bupivacaine</td>
<td>54%</td>
<td>FP rate 26% (2 of 38 had pain relief with only 1 block while 8 had longer relief with lidocaine)</td>
<td>All pts had whiplash after an MVC. 7 pts withdrew and 5 only completed one block.</td>
</tr>
<tr>
<td>Lord et al 1995</td>
<td>50 pts</td>
<td>Near-complete relief with lidocaine and bupivacaine MBB, with the duration of relief lasting longer with bupivacaine</td>
<td>26%</td>
<td>FP rate 19% based on concordant response to LA blocks and negative response to saline</td>
<td>20 pts responded to saline injections, including 3 of 14 who had concordant response to lidocaine and bupivacaine.</td>
</tr>
<tr>
<td>Manchikanti et al 2002</td>
<td>120 pts with axial neck pain</td>
<td>≥80% pain relief after MBB with lidocaine followed by bupivacaine</td>
<td>67%</td>
<td>FP rate 24% based on initial positive blocks</td>
<td>Required at least 2 hours relief with lidocaine and 3 hours with bupivacaine. Mixed sarapin and steroids with LA. All pts sedated.</td>
</tr>
<tr>
<td>Manchikanti et al 2002</td>
<td>106 pts with neck pain or without arm pain</td>
<td>≥75% pain relief after MBB with lidocaine followed by bupivacaine</td>
<td>60%</td>
<td>FP rate 21% based on initial positive blocks</td>
<td>Required at least 90 min relief with lidocaine and 3 hours with bupivacaine. Some patients had radicular pain. All pts sedated.</td>
</tr>
<tr>
<td>Manchikanti et al 2004</td>
<td>255 pts with neck pain</td>
<td>≥80% relief during painful movements after MBB with lidocaine followed by bupivacaine</td>
<td>55%</td>
<td>FP rate 34% based on initial positive blocks</td>
<td>Required at least 2 hours relief with lidocaine and 3 hours with bupivacaine. All pts sedated.</td>
</tr>
<tr>
<td>Manchikanti et al 2008</td>
<td>251 pts with non-radiculard head pain with n=45 and without n=206 prior surgery</td>
<td>≥80% relief and ability to perform previously painful movements after MBB with lidocaine followed by bupivacaine</td>
<td>39% in non-surgical and 36% in post-surgical pts</td>
<td>FP rate 43% in non-surgical group, 50% in post-surgical group</td>
<td>Required at least 2 hours relief with lidocaine and 3 hours with bupivacaine. All pts sedated.</td>
</tr>
<tr>
<td>Manchukonda et al 2007</td>
<td>251 pts with non-radiculard neck pain</td>
<td>≥80% relief and ability to perform previously painful movements after MBB with lidocaine followed by bupivacaine</td>
<td>39%</td>
<td>FP rate 45%</td>
<td>Retrospective study. Required at least 2 hours relief with lidocaine and 3 hours with bupivacaine.</td>
</tr>
<tr>
<td>Yin and Bigdul, 2008</td>
<td>84 pts with non-radiculard neck pain</td>
<td>Complete pain relief after MBB lasting longer with bupivacaine than lidocaine. Those with discordant relief had to have negative block to placebo</td>
<td>55%</td>
<td>FP rate 15%</td>
<td>Retrospective study.</td>
</tr>
<tr>
<td>Spelderwilde et al 2001</td>
<td>97 pts with disabling axial neck pain</td>
<td>Near-complete pain relief (post-block pain score ≤1/10) after MBB done with lidocaine and bupivacaine</td>
<td>36% based only on confirmatory double blocks, 55% if 18 pts who did not receive a confirmatory block are included</td>
<td>FP rate 0%, though 18 of the 53 pts with a (+) initial block did not return for a confirmatory block</td>
<td>Retrospective study. Most pts had post-traumatic pain.</td>
</tr>
<tr>
<td>Cohen et al 2010</td>
<td>24 pts with axial neck pain</td>
<td>≥50% pain relief lasting ≥3 hours after a single bupivacaine block. Pts underwent CT after MBB</td>
<td>38%</td>
<td>FN rate 7% (6 of 84 blocks failed to capture target nerve)</td>
<td>Randomized trial evaluating effect of block volume on outcomes (0.25 vs 0.5 mL), with lower volumes found to be more specific.</td>
</tr>
</tbody>
</table>

CT, computed tomography; FN, false-negative; FP, false-positive; LA, local anesthetic; MBB, medial branch blocks; MVC, motor vehicle collision; pts, patients.
of 22.4 weeks. The results were not stratified by type of RFA, but bilateral TON-only and left-sided TON-only accounted for 17% of cases. In an observational study by Govind et al using comparative LA blocks to treat third occipital headache, 49 out of 120 patients responded positively and underwent RFA. Among these, 88% obtained a positive outcome, lasting a mean duration of 297 days.

Predictive modeling
The NIH Federal Pain Research Strategy designated identifying individuals likely to respond (or fare poorly) to treatment as a major priority, which may facilitate a precision or personalized medicine approach. In theory, this might include considering the results of genetic testing in the future, clinical findings, and environmental and cultural considerations. In practice, this might entail tailoring the approach based on radiological imaging if available, physical examination findings, pre-test probability (eg, a younger person with non-traumatic pain in whom the false-positive rate of a single block might be higher than in an elderly patient with osteoarthritis and paraspinal tenderness), the results of an initial block (ie, considering dual blocks in an individual with a borderline positive block) and unique patient circumstances (eg, needle aversion or the need for sedation, travel difficulties, concomitant anticoagulant therapy in high-risk individuals).

Recommendations
To maximize access to care, we recommend a single block before cervical medial branch RFA. We acknowledge that using dual blocks may increase the RFA success rate, but the evidence is clear that using double blocks will result in a significant proportion of false-negative procedures and a decreased overall success rate; grade B recommendation, low-to-moderate level of certainty.

QUESTION 15: SHOULD ELECTRODES BE POSITIONED IN A CERTAIN ORIENTATION OR LOCATION AND, IF SO, WHAT IS THAT ORIENTATION? DOES THE ORIENTATION DIFFER WITH PRIOR SURGERY?
There is no primary peer-reviewed evidence directly comparing electrode location or orientation for the technical performance of cervical medial branch RFA, with the only direct comparative information coming from a non-peer-reviewed abstract. In this retrospective study that examined the medical records of 82 patients, individuals who received a lateral (ie, perpendicular) approach experienced greater pain reduction at 6 months and 1 year than those who underwent RFA via a posterior (ie, parallel) approach.

Conceptually, it stands to reason that treatment is unlikely to be successful beyond a placebo effect if the target nerve is not captured by the lesion. There is an increased likelihood of ablating targeted nerves with larger lesions created along the known course of the nerves. Evidence can be gleaned regarding optimal electrode location and orientation by considering the anatomy of the cervical medial branches and TON (figure 3), and the physics of lesions created using traditional monopolar RF electrodes. As a corollary, the existing body of evidence on the effectiveness and efficacy of cervical medial branch RFA may also be used to address this question.

Anatomy
The anatomy of the cervical medial branches of the dorsal rami was first described in detail by Bogduk in 1982. This seminal work remains one of the best descriptions of the relevant anatomy to cervical medial branch RFA. In general, the cervical medial branches course around the ipsilateral articular pillars along the anterolateral, lateral, and posterolateral portions of the pillar. Using C5 as the reference, the general archetypal relationship between the nerve and bone consists of the nerve running transversely across the centroid of the articular pillar from a lateral view, and running in a cephalad to caudal direction at roughly the same angle as the facet joint itself. From a posterior anterior view, the medial branch courses just lateral to the concavity of the articular pillar. Slight variations exist at each segment, with the nerve location being slightly higher on their respective articular pillars at levels distal to C5. This results in the C7 medial branch nerve being located along the superior portion of the articular process itself, and the deep C3 medial branch being located so high along the C3 articular pillar that at times it may cross the lower portion of the C2–3 joint line itself.

The innervation of the C2–3 joint differs in that it is innervated exclusively in most people by the TON. The location of this nerve is more variable than the locations of other cervical medial branches. The TON crosses the C2–3 joint anywhere from the apex of the C3 superior articular process to opposite the bottom of the C2–3 intervertebral foramen. This difference in anatomy must be accounted for when lesioning the TON.

Physics
Using a traditional monopolar electrode, coagulation occurs primarily in a radial direction perpendicular to the long axis of the electrode so that the lesion generated is prolate spheroid in shape. The lesion extends minimally beyond the tip of the electrode. With a 10 mm active tip, the lesion is just longer than 10 mm in a longitudinal section, and slightly wider than the width (gauge) of the cannula in a transverse plane.

Based on the known anatomy of the cervical medial branches and lesion shape created by a conventional unipolar RF electrode, using a posterior-to-anterior approach or a posterior oblique approach seems most likely to result in the active tip of the electrode being situated somewhere along the course of the nerve and generating a lesion that captures the medial branch (figure 4). This may be optimized by placing the electrodes in a plane parallel to the posteroanterior orientation of the articular pillar (ie, a posterior approach) in a trajectory that transects the nerves, or by creating multiple lesions in an orientation near-parallel to the target nerve(s). When choosing a posterior oblique approach, placing the cannula at the anterior aspect of the facet column has been recommended. No anatomic, physical, or conceptual evidence supports the idea that placing an electrode in a plane perpendicular to the articular pillar (ie, using a lateral approach) will result in an equal or greater likelihood of a technically successful nerve ablation. For the TON, which innervates the C2–3 joint, the variable location of the nerve relative to other cervical medial branches suggests that multiple lesions that extend from the bottom of the C2–3 foramen to the apex of the C3 superior articular process may be required to reliably capture the nerve.

Efficacy and effectiveness literature
One of the most recent comprehensive systematic reviews of cervical medial branch RFA included both comparative and cohort studies. There were no exclusion criteria pertinent to electrode orientation, although four studies were excluded since the technique was deemed outdated and inconsistent.
with current practice. Twenty-one manuscripts were considered, all of which used some variation of parallel electrode placement. One randomized sham-controlled study demonstrating the efficacy of cervical medial branch RFA for the treatment of neck pain described a rigorous technique including both a posterior pass and an anterior oblique pass of the electrode. Two other sham-controlled trials that yielded equivocal results (ie, some benefit favoring RFA but failure to meet the primary endpoint) described posterior oblique approaches. No published peer-reviewed studies supporting the use of cervical medial branch RFA for the treatment of neck pain have described using a perpendicular approach. As noted above, a single non-peer-reviewed abstract retrospectively comparing the perpendicular versus parallel approach in 82 patients concluded that “a statistically significant interaction between RFA treatment and post-procedural time indicated that the lateral RFA group experienced higher mean percent pain reduction were not statistically significant at 1 and 3 months (p=0.05) and 0.06, respectively”. No actual data were reported, including the type of electrode used, the magnitude of relief, or the responder rate.

Literature specific to RFA of the TON is much less robust. This may be in part due to early findings that outcomes when targeting the C2–3 joint were inferior compared with RFA targeting other cervical facet joints, suggesting that technical advances were necessary to reliably lesion the TON. A 2003 study recommended a new technique using a parallel approach that created three successive lesions with the electrodes positioned no further than two electrode diameters apart. This accounted for the greater variability in the location of the TON, with outcomes mirroring the success rates of RFA applied to other cervical levels. This has since been replicated in other studies. There are no studies reporting RFA outcomes targeting the TON using a perpendicular approach.

Performing RFA at operated levels

There are no clear-cut indications for cervical medial branch RFA at the level of a fused cervical segment, which is a common treatment for a painful diarthrodial synovial joint. Outside of the presence of non-union, pseudoarthrosis, or hardware failure, it is less likely that a cervical facet joint is the predominant pain generator at a fused segment than a non-fused segment.

Therefore, other causes of axial pain (eg, myofascial) must be ruled out. When performing plating or fusion of the articular pillars, some surgeons purposefully or inadvertently sever the medial branches. However, spondylotic disease and facet arthropathy may occur at increased rates at levels adjacent to fused cervical segments, which is a phenomenon known as adjacent segment disease. A cadaveric study performed in the lumbar spine found that fixation resulted in reduced facet joint capsular strain at the level of fixation, but increased strain at adjacent levels. In the cervical spine, another cadaveric study found increased facet joint forces at the treated level after arthroplasty during extension, and decreased forces at fused levels but increased forces at adjacent segments following arthrodesis.

Several studies have examined facet block and RFA outcomes in patients who have previously undergone cervical spine fusion. Manchikanti et al performed double blocks in 251 patients with chronic neck pain, 45 of whom underwent prior surgery. They found no significant differences in prevalence rates between the surgical and non-surgical cohorts. However, the authors did not note whether the cervical MBB was performed at the operated or non-operated levels. Lang and Buchfelder reported a mean 176 days of pain relief (return to 50% of baseline) in 21 patients with cervicogenic headache who underwent C2–3 or C3–4 cervical facet RFA after pseudoarthrosis. A study by Cohen et al reported a 59% success rate in 29 patients who underwent cervical medial branch RFA after fusion, which was not significantly different from RFA outcomes in surgically-naïve patients. The authors in this study also failed to note whether the procedures were performed at the operated levels. Finally, a study by Klessinger et al evaluating cervical medial branch RFA outcomes in post-surgical patients reported a 59% success rate, with no differences stratified by the type of operation (disc replacement, cage placement, surgical plate). Although statistical analysis was not performed, the outcomes did not appear to be different when RFA was performed on operated levels versus adjacent or non-operated levels. When performing RFA at operated levels, it is important to avoid placing the electrode in contact with hardware because of theoretical concerns for heat injuries. Yet, no study to date has reported such an injury. Because hardware for anterior discectomy and cervical fusion is placed in areas distant to the site of cervical medial branch RFA, the presence of hardware should not influence the approach used. For hardware that involves the articular pillars, cervical medial branch RFA would not be ideal.
should either be avoided or, in cases of non-union, a modified (eg, posterior oblique or lateral) approach performed under CT may be necessary.

**Safet y**

Different safety considerations exist for a posterior (parallel) and lateral (perpendicular) approach, both of which can be mitigated by meticulous use of fluoroscopy in multiple views. With a posterior approach, care must be made to avoid placing the electrode too anteriorly and risk violating the vertebral artery or cervical nerve root. With a lateral approach, care must be made to avoid advancing the electrode too medially into the spinal canal itself.

**Recommendations**

Current available basic science and outcome literature suggests a near-parallel (posterior or slight posterior oblique) approach should be used when performing cervical medial branch RFA for all cervical segments. This recommendation includes both anterior operations and non-operated spines but does not apply to the adaptation of other electrode technology, such as cooled RF; grade B recommendation, low-to-moderate level of certainty. For surgeries involving the articular pillars, a modified approach (posterior oblique with greater angulation or lateral) using advanced imaging may be necessary, with multiple lesions created if necessary; grade C recommendation, low level of certainty.

**QUESTION 16: SHOULD SENSORY AND/OR MOTOR STIMULATION BE PERFORMED BEFORE RFA?**

In addition to properly identifying patients whose pain is likely mediated via the medial branches, the RF cannula needs to be in close enough proximity to the intended target to adequately coagulate the medial branches. Care should also be made to avoid complications from lesioning the ventral rami, spinal nerves, or other unintended structures. Although appropriate needle placement for the intended anatomic target can be achieved based on fluoroscopic landmarks, physiologic feedback via sensory and motor stimulation can provide additional input guiding optimal needle placement in the presence of advanced degenerative changes that limit visualization of fluoroscopic landmarks, and anatomic variations in the locations of non-medial branch and medial branch nerves, and multiple articular branches emanating from each nerve. And when performing the stimulation, the patient is asked to inform the treating physician if they feel paresthesia (eg, tingling, buzzing, vibration, pain) and to identify the location of the sensory change to ensure concordance with established axial referral patterns. In one study, Marks found that pain evoked from medial branch stimulation often extended distal to generally recognized facet referral patterns as outlined by other means.

Ideally, the sensory threshold can be used to determine optimal RF cannula placement whereby the placement of the cannula can be adjusted in all three dimensions (anterior–posterior, cephalad–caudal, and medial–lateral) to achieve the relative lowest stimulation threshold. Local muscle stimulation and a perpendicular approach can ostensibly lead to adequate sensory stimulation (threshold ≤0.5 V) but result in a situation where the subsequent lesion is insufficient for complete coagulation of the nerve and relief of pain.

There are scant data on the relationship between sensory stimulation threshold and the distance to a nerve, although there is clearly a direct, but imperfect, correlation. In rodent and computer simulation studies, nerve activation with anodic stimulation at 0.5 milliamperes occurs at a distance of around 4 mm, while cathodic stimulation at 0.5 milliamperes can activate nerves at a distance of 7 mm. For regional anesthesia nerve blocks, the elicitation of a motor response at or below 0.5 milliamperes is considered to be a common aim for successful neural blockade.

Only one study has examined the correlation between medial branch RFA outcomes in humans. In a prospective study performed in 61 individuals who underwent lumbar medial branch RFA with light sedation after a positive diagnostic block, Cohen et al found no significant correlation between the average absolute sensory threshold and treatment results. However, for each patient sensory testing was used to optimize cannula placement so the lack of correlation between the average sensory thresholds and treatment outcomes could be related to other factors (eg, sedation, baseline algieses, comorbidities, age, and genetics) that affect sensory perception. Compared with light sedation (ie, the patient is relaxed but responsive to verbal stimuli), deep sedation could preclude the utility or even use of sensory stimulation to guide RF cannula placement. Sensory stimulation may be particularly important in the upper cervical levels (eg, TON, C3 medial branch, and C4 medial branch) wherein motor stimulation does not always result in discernable distal muscle contraction as seen in lower cervical levels where the spinal nerves innervate the arm.

**Rationale for motor stimulation and evidence**

Motor stimulation is typically performed at 2 Hz. The presence of multifidus or other paraspinal muscle stimulation can be a marker of proper cannula placement. In a small study performed in the lumbar spine, when multifidus stimulation was used without sensory stimulation, a success rate of 87% was observed at 12 months post lumbar medial branch RFA. In a retrospective review of 68 patients by Koh et al patients who underwent lumbar medial branch RFA were stratified into three groups: complete twitching (paravertebral muscle contraction at all stimulated levels), partial twitching (paravertebral muscle contraction at one or two levels), and no twitching (no paravertebral contraction observed at any level). At 6 months, the proportion of successful procedures was statistically greater in the group with paravertebral muscle contraction at all stimulated levels compared with the group where no paravertebral muscle contraction was observed at any level. The mean duration of benefit was 4.6, 5.8, and 7.0 months for the ‘no-twitch’, ‘partial twitch’, and ‘complete twitch’ groups, respectively. In a small observational study by Dreyfuss et al all patients treated with unilateral lumbar medial branch RFA had multifidus denervation observed on EMG 6 weeks following denervation and were noted to have sustained pain relief at 12 months, with no discernable segmental multifidus atrophy noted on MRI in long-term follow-up.

**Enhanced safety**

Eliciting multifidus and other paraspinal muscle contraction may provide false confidence in cannula placement when the motor nerve that is being stimulated does not innervate the facet joint (eg, cases of aberrant non-medial branch innervation, stimulation of other branches of the dorsal ramus). To enhance safety, practitioners typically increase the voltage to three times the sensory stimulation threshold (1.5–2.0 V) during motor testing. However, this threshold is based on typical sensory thresholds and anatomic...
C2–3 facet radiofrequency denervation

Many of the initial studies investigating the efficacy of medial branch RFA did not use motor or sensory stimulation to help guide RF cannula placement, but rather created multiple (eg, 3–6) empirical lesions per level based on anatomic landmarks using fluoroscopic imaging.\textsuperscript{68, 211, 312} Due to the high failure rate and ataxia observed in the pilot study, the C2–3 level was omitted from the incipient comparison with a prior report of Govind et al.\textsuperscript{244} When sensorimotor testing was used with TON systems, the efficacy of cervical RFA for the treatment of whiplash-induced neck pain.\textsuperscript{68} Subsequent studies with a revised technique and/or use of sensorimotor stimulation for TON RF neurotomy have demonstrated more sustained and reproducible responses along with fewer and more tolerable side effects after C2–3 facet denervation.\textsuperscript{206, 36, 91, 101, 307} One case of dropped head syndrome has been reported after denervation of the left C2–4 facet joints when motor stimulation was not performed.\textsuperscript{244} When sensorimotor testing was used with TON RFA in a retrospective review by Gazelka et al,\textsuperscript{293} the reported rate of RFA-induced third occipital neuralgia was 19%, which favorably compared with a prior report of Govind et al\textsuperscript{307} that found a 55% rate of dysesthesia and 15% rate of hypersensitivity when needles were placed solely by anatomic landmarks. When motor testing is used for levels above C5, one should look for contractions in muscles other than the arm to confirm the electrode is a safe distance from the spinal nerve or ventral rami (eg, trapezius, sternocleidomastoid, scalenes, shoulder, diaphragm).

Recommendations

We recommend strong consideration of the use of sensory stimulation when single lesions are anticipated and/or with C2–3 facet denervation; grade C recommendation, low level of certainty. When multiple lesions are planned, the evidence for sensory stimulation is inconclusive; grade I recommendation. For motor stimulation, we find that it may be beneficial for both safety and effectiveness purposes; grade B recommendation, low-to-moderate level of certainty.

<table>
<thead>
<tr>
<th>Table 19</th>
<th>Cervical medial branch diameters\textsuperscript{202, 203}</th>
</tr>
</thead>
<tbody>
<tr>
<td>Medial branch</td>
<td>Mean±SD diameter (mm) *</td>
</tr>
<tr>
<td>C3 superficial medial branch (TON)</td>
<td>1.5±0.4</td>
</tr>
<tr>
<td>C3 deep medial branch</td>
<td>0.5±0.2</td>
</tr>
<tr>
<td>C4 medial branch</td>
<td>0.9±0.3</td>
</tr>
<tr>
<td>C5 medial branch</td>
<td>0.9±0.3</td>
</tr>
<tr>
<td>C6 medial branch</td>
<td>0.7±0.1</td>
</tr>
<tr>
<td>C7 medial branch</td>
<td>0.6±0.2</td>
</tr>
</tbody>
</table>

\*Adopted from Lord et al\textsuperscript{202}  
†Adopted from Kweon et al\textsuperscript{293}  

QUESTION 17: IS THERE EVIDENCE FOR LARGER LESIONS TO IMPROVE OUTCOME MEASURES FOR RFA? IF SO, HOW CAN LESION SIZE BE INCREASED?

Anatomic considerations and challenges

Multiple anatomic constraints and challenges provide a logical basis for modulating lesion size when targeting the cervical medial branches including the TON. The cervical medial branches are small and associated with variable anatomy.\textsuperscript{300, 306, 307, 200, 202, 203} First, all the cervical medial branches except for the TON (ie, C3 superficial medial branch) have mean diameters ≤1 mm (table 19) and are often displaced from the bone approximately 1–2 mm by areolar tissue.\textsuperscript{300, 302, 203} Second, the cervical medial branches have variable courses relative to bony anatomic targets.\textsuperscript{200, 202, 307} Specifically, the C3, C6, and C7 medial branches exhibit significant anatomic variation. The C5 medial branch position has a more consistent relationship and typically courses transversely across the center of the articular pillar. The C3, C4, and C6 medial branches are typically located higher on the respective articular pillar. Third, the cervical dorsal rami at C4, C5, and C6 often give off two medial branches. In one anatomic analysis performed in 20 cervical cadaver specimens, two cervical medial branches were found 27%, 15%, and 2% of the time at the C4, C5, and C6 dorsal rami, respectively.\textsuperscript{203} Fourth, the articular pillar size and the proximity of the cervical nerve root at each level impose careful safety considerations when contemplating lesion size modification.\textsuperscript{200, 331–333}

Specifically for the TON, a high technical failure rate was noted when using small-gauge electrodes, with an early return of pain.\textsuperscript{307} When larger gauge electrodes were employed, an 88% initial success rate was documented with a median duration of complete pain relief in patients with a successful outcome of 297 days.\textsuperscript{307} The TON has the largest mean diameter (table 19). A single application of thermal RFA with a small-gauge cannula creating a commensurately small lesion may therefore inadequately coagulate the nerve.

Technical limitations and challenges

In addition to the anatomic challenges associated with cervical medial branch RFA, technical challenges exist. The clinical cervical medial branch RFA data to date have examined the use of traditional thermal ablation; therefore, this question will limit discussion of RFA with internally-cooled electrodes. Thermal RFA involves a high-frequency alternating current that flows from the uninsulated active tip of the cannula/electrode apparatus into the tissue, causing ionic agitation.\textsuperscript{334} The ionic agitation then results in tissue frictional heating adjacent to the cannula and the tissue becomes the primary heat source. Three main factors determine tissue heat generation: the distance from the active tip, RFA current intensity, and duration of RFA application.\textsuperscript{335} Furthermore, the tissue including the bone–muscle interface and the fluid environment surrounding the RFA zone modulate lesion size.\textsuperscript{335–341}

The lesion sizes associated with traditional thermal temperature-controlled RFA set-ups are small and therefore necessitate cannula placement extremely close or directly on top of the medial branch to ensure the created coagulation zone encompasses the full thickness of the targeted nerve.\textsuperscript{310} Although ex vivo studies examining RFA lesion dimensions can underestimate size, they do provide insight into lesion development.\textsuperscript{340, 342} Traditional thermal RFA lesions develop horizontally with minimal ablation occurring distal to the tip.\textsuperscript{310, 335, 343} When examining small-gauge cannulae (eg, 22-gauge) with small active tips (≤5 mm), lesion size is limited and the minimum relationships in the lumbar spine,\textsuperscript{329} which differ in the cervical spine.\textsuperscript{285} One case report described a patient who suffered L5 sensory radiculopathy following L3–5 dorsal ramus denervation when motor testing was not conducted above 1 V.\textsuperscript{230} Following motor stimulation, if no muscle contraction in the appropriate myotomal distribution is identified, most practitioners believe it is safe to proceed. Some RF systems also require the electrode to be removed during LA administration which can lead to inadvertent needle movement. With these older systems, best practices would entail obtaining comparative images before and after anesthetic injection to ensure that no movement of the RF cannula has occurred. Many newer RF systems contain a separate port to allow for LA injection without significant manipulation of the cannula.
Neuropathological correlation: RFA physiological and functional responses

In addition to limiting technical failures, increasing the energy delivered and hence lesion size may result in physiological and functional responses advantageous to prolonging pain relief with RFA. Thermal RFA affects both myelinated and unmyelinated fibers and results in a third-degree peripheral nerve injury with extensive Wallerian degeneration. In a third-degree peripheral nerve injury, the myelin, axon, and endoneurium are disrupted while the fascicular arrangement, epineurium, and perineurium are maintained. Following thermal RFA, nerve regeneration may be initiated as early as 30 min post-lesion creation through three mechanisms: remyelination, collateral sprouting from preserved axons, and regeneration.

Recently, preclinical work suggests that RFA technology, lesion size, and energy deposited may result in different structural and functional changes in peripheral nerves. A preclinical study comparing internally-cooled electrodes versus traditional RFA demonstrated that the bigger lesions associated with greater energy delivery generated from internally-cooled electrodes resulted in larger areas of thermal damage, with edematous/inflammatory zones persisting longer. In addition, there was greater attenuation with an extended loss of nerve function when assessed by EMG with internally-cooled electrodes. Future research is needed to determine if the greater interruption in axons and electrical conduction with internally-cooled electrodes is specific to the technology or solely dependent on the amount of energy delivered during RFA application. The addition of hypertonic saline to traditional thermal RFA also significantly increases peak power and total energy delivered during RFA.

Methods to enhance lesion size

Multiple methods exist to increase lesion size (ie, length and width) with traditional thermal RFA. The primary methods include adjustments in active tip length, cannula size, temperature, and time settings. For example, increasing the cannula diameter from a 22-gauge to a 16-gauge increases average lesion width by 58–65% (3–4 mm) with an RFA setting of 80°C for 2 min. When temperature is increased from 60°C to 90°C for 2 min, lesion width increases by 108–152%. Lesion growth occurs most prominently during the initial portion of the lesion cycle, with approximately 40% of the mean maximum surface area occurring once the set temperature is achieved and 87% of the maximal lesion surface area occurring by 90 s after the set temperature is reached. However, extending lesion time beyond 90 s is still advantageous because of a further increase in lesion size in many patients and a reduction in lesion variability. Extending lesion time should particularly be considered when only one lesion cycle is being performed. Other methods to increase lesion size include modification of the local tissue environment through fluid pre-injection (ie, saline pre-injection) and the use of bipolar or internally-cooled electrodes. However, caution is advised and future research focused on safety is warranted before routine deployment of these modalities in the cervical spine.

Risk mitigation

The creation of larger lesions with greater energy deposition must be performed with care in the cervical spine. Caution is heightened in the cervical spine because of cervical anatomy. The articular pillars are narrow in the cervical spine with critical structures in close proximity including the cervical nerve roots, deep cervical artery, radicular arteries, and vertebral artery. The CT-measured anterior–posterior (horizontal) diameters of the C3 through C6 vertebral body articular pillars range from 6.4 mm to 11.5 mm, with a mean of 9.1 ±1.2 mm. The CT-measured oblique anterior–posterior diameters of the articular pillars range from 11.4 mm to 19.3 mm, with a mean of 14.5 ±1.7 mm. The CT-measured average height of the articular pillars varies from 16.9 mm to 25.6 mm, with a mean of 21.0 ±1.9 mm. Therefore, depending on the RFA approach to the cervical medial branches, using a large active tip (ie, 10 mm) may create an extended lesion that is longer than the articular pillar anterior–posterior diameter.

Practitioners must appreciate the proximity of the cervical nerve roots to the cervical articular pillars. For the C3 through C7 vertebrae, the average distance from the nerve roots to the superficial posterior center of the inferior lateral mass is 5.5±0.8 mm, and 5.7±1.5 mm from the nerve roots to the superi- rior lateral mass. Based on in vivo work, the lesion width (ie, horizontal diameter 10.7 mm) created by an 18-gauge 10 mm active tip electrode heated for 120 s may reach and exceed these boundaries, placing the cervical nerve root at risk of coagulation. Other technical factors must be considered when optimizing safety in the cervical spine. Traditional thermal RFA needles are associated with limited projection distal to the active tip (<1 mm). Internally-cooled electrodes project 40% of the lesion distal to the active tip. Therefore, if these needles are placed via the posterior parasagittal and oblique approaches, the lesion may project towards the cervical nerve root. Fluid pre-injection with hypertonic saline also alters lesion dimensions, with the maximum width of the lesion shifted more towards the distal end of the active tip. Currently, to optimize cervical medial branch RFA, it is recommended that the cannula lies on the anterior side of the facet column. Therefore, any additional forward projection of the lesion from the distal end of the cannulae may be in close proximity to the cervical nerve root. The presence of bone and posterior hardware also alters lesion geometry and energy and heat transmission. When placing a cannula against bone, the maximal effective radius away from bone approximately doubles. Posterior hardware not only...
obstructs needle placement but also allows for the transmission of heat distally towards vulnerable structures such as cervical nerve roots and the vertebral artery.

**Recommendations**

Based on the current limitations of traditional thermal RFA and the small size of the targeted structures (ie, cervical medial branches), creating larger lesions with reduced lesion variability may increase the likelihood of lesioning the targeted structures. However, anatomic constraints limit the extent to which larger lesions can be employed in the cervical spine.

If larger lesions are used, care should be taken to limit damage to untargeted structures including vascular (eg, vertebral artery and deep cervical artery) and neurologic structures (cervical nerve roots). Although improved outcomes have been demonstrated with larger gauge electrodes when targeting the TON, caution must be emphasized. When there is a concern for lesioning non-targeted tissue, the need persists to optimize lesion size, a multi-lesion protocol with a smaller gauge and/or shorter active tip cannula may be considered. Sequential cannula placement for each targeted medial branch should be no more than 1 or 1.5 electrode widths apart. Multiple lesions are required when targeting the TON secondary to its relatively large and variable course. Before routine deployment of new cannula designs, bipolar or internally-cooled electrodes, and fluid modulation (besides pre-injection of LA for patient comfort), safety studies are required. Grade C recommendation with low-to-moderate level certainty for using larger lesions to improve the ability to capture the targeted nerves. Grade C with a low level of certainty for the ability of larger lesions to increase the duration of pain relief.

**QUESTION 18: WHAT ARE THE MOST COMMON COMPLICATIONS OF CERVICAL FACET INTERVENTIONS AND HOW CAN THEY BE MINIMIZED?**

**Common complications of procedures on cervical facet joints**

Cervical facet joint interventions (IA facet joint injections, MBB, and RFA of the innervation to the cervical facet joints) performed with appropriate use of image guidance rarely result in serious complications. However, both minor and major complications have been reported. Post-procedural pain and soreness at the procedure site, bruising, light-headedness, flushing, sweating, and nausea are some of the common but transient adverse effects following procedures on the cervical facet joints. A recent comparative study reported an incidence of 4% for exacerbation of pain lasting 2 weeks, syncope, and headaches following US or fluoroscopy-guided cervical MBB, with no difference between the two imaging modalities. Transient disequilibrium on rising from the supine, lateral, or prone position after the procedure and presyncope during the procedure is common with cervical MBB, especially in procedures on upper cervical facet joints. Headaches have been reported in 4% of patients after CT-guided IA cervical facet steroid injections and there is one report of a postural headache after an MBB performed in the supine position. Incorrect position of the needle or use of large volumes of LA for procedures on cervical facet joints can result in temporary sensorimotor neurological deficits in the back of the head, neck and the upper limb due to blockade of branches of the cervical plexuses or the roots of the brachial plexus that are close to the injection site. Intravascular needle placement and/or injection with bleeding, and hematoma formation (2.3% of patients), infection of the superficial tissues (2.9% of patients) and complications of LA such as temporary weakness and numbness (0.15% of patients) have been reported in patients who received cervical facet joint procedures.

**Major complications of procedures on cervical facet joints**

The most common cervical procedure-related serious adverse outcomes resulting in permanent neurologic injury or death are due to direct trauma from the needle to a nerve or spinal cord and infarction of the spinal cord or brainstem from arterial disruption or injection, usually of a particulate steroid into a feeder artery supplying the spinal cord or brain. Other causes of serious adverse outcomes from procedures performed on the cervical spine include hematomas causing cord compression, dural puncture, and high or total spinal block. Rathmell and colleagues acknowledged in their closed claims review that most adverse outcomes reported in the literature have been associated with epidural (interlaminar or transforaminal) injections on the cervical spine. The authors found that the use of general anesthesia or deep sedation was higher in cervical procedure claims associated with spinal cord injuries than for claims that were not associated with spinal cord injuries (67% vs 19%). The rate of non-responsive outcomes and vascular complications associated with spinal cord injuries was 5-fold higher than in cervical procedures not associated with these injuries. A previous closed claims analysis by Fitzgibbon et al reported one case of spinal cord injury after a ‘cervical facet injection’. An evaluation of trends in malpractice claims for pain medicine from 2010 to 2014 found that claims related to cervical procedures were disproportionate to the frequency with which they were performed. A more recent published analysis of 126 closed claims related to interventional pain procedures reported that eight (6.3%) were related to cervical facet joint procedures (four with cervical MBB, another four with RFA) while cervical interlaminar and transforaminal injections accounted for 32 (25%) (table 20).

**Vascular**

In addition to the anteriorly located carotid artery, the proximity of the cervical facet joints to other important vascular structures such as the vertebral artery, the deep cervical artery, the ascending cervical artery, and the transverse cervical artery exposes patients to the risk of vascular complications. Catastrophic consequences related to the penetration of these vascular structures include ischemic injury to the spinal cord and the brain, but these are rare and have mostly been reported with cervical transforaminal ESIs. Another aspect to consider regarding the potential for vascular penetration during cervical facet procedures is the practice of injecting particulate steroids following cervical medial branch RFA. This has been suggested as a strategy to reduce post-RFA neuritis following lumbar facet RFA, but the evidence for this is conflicting, and it is unclear if the results of studies on lumbar facet RFA can be extrapolated to cervical facet RFA procedures. Given the higher probability of encountering important feeder vessels to the spinal cord during cervical medial branch RFA, it may be appropriate to consider the use of non-particulate steroids if there is a concern about post-lesion neuritis.

A recent study in 275 patients (36 of whom received cervical facet interventions) found that continuing the use of antiplatelet and anticoagulant medications in patients undergoing procedures on the cervical facet joints does not increase the risk of bleeding and hematoma. An online survey of 325 members of ASIPP also found that thromboembolic events were more severe and common when anticoagulants were stopped prior to performing fluoroscopically-guided interventional pain procedures on the cervical spine. 38


spine, while there was no difference in the reported occurrence of bleeding complications based on whether or not anticoagu-
ulants were continued.160 However, cervical MBB and RFA are
currently classified as ‘intermediate-risk’ procedures in the multi-
organizational institutional guidelines for patients on antico-
agulant and antiplatelet medications, and the recommendations
regarding stopping these medications in the guidelines should be
followed unless extenuating circumstances dictate otherwise.161
Recent studies that used one or more of the following imaging
modalities—fluoroscopy with injection of contrast and DSA, US,
and CT—have provided information about the risk of vascular
injury with cervical facet joint interventions. In a study on the
anatomic variations of the vertebral artery evaluated on CT-an-
giography imaging, a loop of the vertebral artery originating
between C1 and C2 coursed over the anterolateral aspect of the
caudal quarter of the articular pillar of C2 in 0.2% of subjects,
over the cephalad quarter of C3 in 7.9%, and over the C2–3
joint line in up to 5.5% of persons.162 This suggests that vascular
injury and catastrophic complications can occur during proce-
dures performed on the TON where it is amenable to treatment
on the surface of the C2–3 facet joint. A recent study reported
that intravascular spread was detected in 10.7% of DSA images
versus 1.7% from static images during cervical MBB. The authors
reported a higher incidence of intravascular injections during
MBB at the C4, C5, C6 levels but did not perform blocks on the
TON.134 Another study reported that 12% of fluoroscopically-
guided—but no US-guided—cervical MBBs were associated with
intravascular placement of the needle tip. In the US-guided
group, vascular structures were found to overlie the C2–3 joint
9% of the time, and the articular pillars of C3, C4, C5, C6, and
C7 in 16%, 16%, 12%, 32%, and 46% of cases, respectively.145
However, DSA is the reference standard for detecting intravas-
cular uptake during facet procedures, with real-time
fluoroscopy, preferably with DSA, prior to performing MBB; grade B recommendation, moderate level of certainty. Reviewing advanced imaging (MRI or CT scan)

<table>
<thead>
<tr>
<th>Table 20 Complications of cervical facet joint procedures</th>
</tr>
</thead>
<tbody>
<tr>
<td>Author, year</td>
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<tr>
<td>--------------</td>
</tr>
<tr>
<td>Barnsley et al 1993</td>
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<tr>
<td>Lord et al 1995</td>
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<td>Lord et al 1996</td>
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<td>Govind et al 2003</td>
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<td>Gazelka et al 2014</td>
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MVC, motor vehicle collision; pts, patients; RFA, radiofrequency ablation; TON, third occipital nerve.

The potential risk for vascular injury and bleeding must be weighed against the risk of serious and catastrophic events when antiocoagulation is held for interventional pain procedures. Two large cohort studies indicate there is a risk of between 0.2% and 0.4% for major thromboembolic events when anticoagulants are held before pain procedures.164,165 In a systematic review designed to determine the benefits of heparin bridging before invasive procedures or surgery, the risk of a thromboembolic event was 0.9% in bridged patients and 0.5% in non-bridged patients, which fell shy of statistical significance (RR 1.26, 95% CI 0.61 to 2.58; p=0.53).166 A recent review by the Spine Intervention Society Patient Safety Committee found no cases of serious complications related to bleeding from cervical facet procedures, suggesting that the risk of discontinuing anticoagu-
ulants before cervical medial branch RFA might outweigh the benefits, and concluded that the decision to withhold anticoag-
ulants should be made on a case-by-case basis.167 The risk of a thromboembolic event depends on several factors including indication, demographic and clinical factors (age, sex, genetics, past history, lifestyle, obesity, hormonal levels), and the type of anticoagulation and length of discontinuation.168

Recommendations

The committee recommends checking for intravascular placement of the needle tip by aspirating and visualizing the spread of contrast using real-time fluoroscopy, preferably with DSA, prior to performing MBB; grade B recommendation, moderate level of certainty. Reviewing advanced imaging (MRI or CT scan)
of the cervical spine for any aberrations in vascular anatomy around the facet articular pillars and/or a ‘scout’ US scan prior to performing the procedure may reduce the risk of vascular complications with cervical facet procedures; grade C recommendation, low level of certainty. Positioning the tip of the RF cannula in the posterior two-thirds of the C2/3 facet joint and avoiding the anterior part of the inferior C2 facet pillar may avoid penetration of an aberrant loop of the vertebral artery; grade C recommendation, low level of certainty.

The issue of continuing or stopping anticoagulation is complex, and readers are referred to existing guidelines. The risk: benefit ratio between stopping or continuing anticoagulation depends on a multitude of factors including the procedure, approach, the use of imaging, needle size and levels, indication for anticoagulation, clinical risk assessment, alternatives to facet interventions, and physician judgment. Healthcare providers considering discontinuation of anticoagulants prior to cervical joint blocks or RFA procedures should consult with the physician prescribing these medications, weigh the risks and benefits for the individual patient (personalized medicine), and discuss these recommendations with the patient (ie, a shared decision model) prior to making any changes. If anticoagulation is continued, the risks of vascular trauma in these circumstances may be attenuated by adjustments in needle and electrode size, insertion trajectory, and the strategic use of pre- and peri-procedural imaging. Ultimately, the decision as to whether or not to continue anticoagulation for cervical MBB and medial branch RFA should be made on a case-by-case basis after carefully reviewing all relevant clinical factors; grade I recommendation.

Complications of RFA of the cervical medial branches

RFA of the nerve supply to cervical facet joints may be associated with more significant complications than MBB or IA injections because the needle diameter is often larger and tissue damage can occur from the thermal lesion. A systematic review of publications on fluoroscopically-guided RFA of cervical medial branches reported only minor and temporary adverse effects, with no serious complications. Common (incidence above 40% in some reports) adverse effects of this procedure include post-procedural pain, cutaneous numbness, dysesthesias, dizziness, and ataxia lasting a few days to weeks. Pruritus, vasovagal syncope, and transient neuritis occur in less than 10% of cases and are usually self-limited. Koebner’s phenomenon (rash at the site of insertion of RF cannula in a patient with a history of skin rashes) is a rare adverse effect of RFA.

Third occipital nerve RFA

A variable period of numbness in 60–97% of patients who undergo RFA of the TON in the cutaneous distribution of this nerve has been reported. The numbness usually lasts for 1–3 weeks and is often replaced by dysesthesia and pruritus followed by a return of normal cutaneous sensation. This is likely due to denervation of its cutaneous branches. Ataxia is also a common adverse effect of TON neurotomy with a variable time period required by patients to adjust. A prospective observational study on neuropathic pain 1 month to 1 year after TON RFA in 64 patients reported symptoms in 12 patients, for an incidence of 19%. Seven of the 12 patients had persistent symptoms for an average of 2.6 months after the ablation procedure (range 1–6 months). In four of the 12 patients the duration of symptoms could not be established because of a lack of follow-up. One patient had persistent pain at 1 year after TON RFA. Some studies on lumbar facet RFA procedures have sought to identify ways to prevent post-procedural pain and numbness. In a randomized trial performed in 45 patients, a significant decrease in local tenderness and post-lumbar medial branch RFA soreness was reported in patients who received either pentoxifylline 10 mg or methylprednisolone 10 mg after lesioning, but not in those receiving saline. However, a more recent retrospective study performed in 164 patients who underwent lumbar medial branch RFA found no difference in the incidence of post-procedure neuritis between individuals who received post-neurotomy particulate steroids (6.4%) compared with those who did not (6.9%). If steroids are injected, given the proximity to radiculomедullary arteries feeding the spinal cord and the vertebral artery and the increased risks associated with the inadvertent injection of particulate steroids into these vessels, the use of non-particulate steroids, which have a faster onset, should be considered.

Several studies have sought to determine whether adjuvants or non-steroidal anti-inflammatory drugs can prevent post-RFA neuritis, with mixed results. A retrospective study evaluating the incidence of neuritis in 215 patients after lumbar and cervical medial branch RFA found a lower incidence of neuritis in individuals taking gabapentin for at least 2 weeks before the procedure (7.1% vs 13.2%) but the difference was not statistically significant (OR 0.51, 95% CI 0.15 to 1.54). Another retrospective study by Singh et al performed in 164 patients found no difference in post-lumbar MBB RFA neuritis stratified by whether or not patients were receiving anti-neuropathic pain medications (membrane stabilizers and antidepressants). In an earlier randomized placebo-controlled trial by Ma et al performed in 66 patients who underwent lumbar medial branch RFA, the authors found that 3- and 7-day treatment regimens with diclofenac significantly reduced post-procedure pain for up to 7 days compared with placebo. A flaw in all of these preemptive analgesic studies is the failure to adequately discriminate between neuritis and procedure-related pain from electrode insertion.

Recommendations

The committee recommends that physicians inform patients about the common adverse effects of RFA including pain, dysesthesias, numbness, dizziness, and ataxia lasting from a few days to a few weeks following cervical facet joint denervation. The use of a short 3-day course of non-steroidal anti-inflammatory drugs commenced immediately after RFA and injection of steroids through the cannula after ablation but prior to its removal may reduce pain and discomfort following RFA. Given the relatively high incidence of critical arteries in the vicinity of an appropriately placed RF cannula, non-particulate steroids should be injected if there is heightened concern about the occurrence of post-RFA lesion neuritis (eg, younger individuals undergoing upper cervical medial branch RFA); grade C recommendation, low level of certainty. There is inconsistent evidence based on low-grade trials for the use of a 2-week course of gabapentin to prevent post-RFA neuritis; grade I recommendation. As noted above, there is evidence that a short course of non-steroidal anti-inflammatory drugs may decrease the severity of post-RFA pain, including neuritis; grade C recommendation, low level of certainty.

Direct trauma to neural structures

Cervical interlaminar and transforaminal epidural injections have been associated with rare catastrophic neurologic injury.
In a closed claims analysis of the type of cervical interventional pain procedure associated with neurologic injury, only 3% (2 out of 64) of procedures were done on the facet joints (both were IA injections). However, damage to the spinal cord from a cannula that deviated medially and was advanced too far anteriorly during a TON RFA procedure has been reported from a cannula that deviated medially and was advanced too far anteriorly during a TON RFA procedure has been reported by authors performing a medicolegal review of complications of procedures on the cervical spine. Anteroposterior views on fluoroscopy showed the RF cannula in the C3–4 intervertebral foramen where it was postulated to have injured a reconfirming radicular artery resulting in spinal cord infarction. In a second case reported by the same authors, a patient developed Brown–Sequard syndrome following an attempted neurotomy of the nerve supply to the C3–4 facet joint. The fluoroscopic images showed the tip of the RF cannula had been placed medial to the facet joint resulting in penetration of the spinal cord. Anterior misplacement of the RF electrode into the neuroforamen can result in the active tip being situated near the cervical nerve root or ventral ramus. Although no such complications associated with cervical facet procedures have been reported in the literature, a case report described new sensory symptoms in the dermatomal distribution of the fifth lumbar nerve following RFA of the third to fifth lumbar medial branches and dorsal rami. Sensory and motor stimulation to reduce the probability of proximity of the RF cannula tip to the spinal nerve root traversing the foramen has been advocated to prevent injury to spinal nerves, but the evidence is inconclusive. Subarachnoid injection of LA has also been described following a fluoroscopically-guided IA injection of the C6–7 facet joint in a patient with a prior anterior fusion. A non-image-guided injection of 4 mL LA into the C5–6 facet joint resulted in transient tetraplegia that lasted 30 min. The authors postulated that the LA was injected into a radicular artery feeding into the anterior spinal artery at that level. Although the use of US may reduce the incidence of vascular penetration, injury to neural structures may occur with misplaced needles (irrespective of the imaging modality) as evidenced by a case of spinal cord injury during a cervical MBB at C7.

Recommendations

The committee recommends the use of true anteroposterior and lateral views on fluoroscopy during placement of RFA cannulas or needles to ensure that the tips are positioned outside the neural foramina. A failure to obtain a true lateral view (ie, a complete overlap of the facet articular pillars) can result in inadvertent needle placement in the intervertebral foramen or the spinal canal. In addition to this view, a contralateral oblique view can be obtained to confirm the position of the tip of the RF cannula. The absence of sensorimotor responses in a radicular distribution in response to test stimulation prior to RFA may also reduce the probability of injury to the spinal cord and spinal nerve roots; grade B recommendation, low-to-moderate level of certainty.

RFA-related degeneration of spinal anatomy and musculature

The main cervical paraspinal muscles include the semispinalis cervicis and capitis, multifidus, splenius cervicis and capitis, trapezius, and levator scapulae muscles. The semispinalis and multifidus muscles are responsible for rotation of the head and extension of the cervical spine. The cervical dorsal rami provide motor innervation to the semispinalis muscles. Edema in the cervical paraspinal muscles was found on imaging following RFA of the upper three cervical facet joints in a patient who developed severe neck pain 1 week after the procedure, possibly reflecting denervation of these muscles. This denervation may expose susceptible patients to the risk of cervical instability and ataxia. RFA of the innervation to two adjacent unilateral cervical facet joints (C2–3, C3–4) has been associated with the development of kyphosis (‘dropped head’ syndrome) due to loss of cervical extensor muscle action that required instrumented posterior fusion for correction. MRI of the cervical spine showed degeneration and atrophy of the semispinalis cervicis and splenius capitis muscles in this patient. Closed claims analysis revealed that this patient may have had unrecognized anterocollis. There is also a similar case reported of RFA of the TON and innervation to the C3–C4 and C5–C6 facet joints on one side followed by RFA at the same levels on the other side 1 week later in a patient who then developed an inability to actively extend his head. The electromyogram in this patient demonstrated active denervation of the cervical paraspinal musculature and the patient eventually required anterior and posterior fusion of the cervical spine from the C3 to C6 levels. The potential causes for this complication include incorrectly positioned RF cannulas resulting in ablation of the dorsal rami, and aberrant anatomy. However, it is also possible that some damage occurs to the innervation of paraspinal muscles despite optimal RF cannula positioning. Cervical medial branch RFA at two adjacent levels denervates only one complete joint and a little more than 20% of the muscles acting on that segment. However, bilateral procedures and/or those done at multiple levels can result in the denervation of a significant mass of cervical musculature. This does not usually result in motor deficits because of collateral innervation to the posterior cervical muscles from adjacent levels. This ‘safety mechanism’ may be lost if RFA is performed at several levels.

Impact of RFA on existing implanted devices

Monopolar RFA of innervation to cervical facet joints involves the use of electrical current that emerges from the tip of the RF cannula and flows through the body before exiting through the dispersive electrode (also known as the grounding pad). Electromagnetic fields created from the use of short-wave diathermy can result in interference with the functioning of implanted electric devices such as deep brain or spinal cord stimulators. There is some evidence that bipolar RFA may be safer than monopolar RFA in patients with implanted devices, with a report of its safe use for cervical facet RFA. However, the aggressive lesions and additional tissue trauma entailed with the use of bipolar lesioning may outweigh any theoretical benefits for cervical medial branch RFA. Moreover, the safe use of monopolar RFA has also been reported in a patient with deep brain stimulators, with one of the implanted pulse generators located
in the anterior abdominal wall.\textsuperscript{377} Unlike lumbar facet RFA procedures for which the grounding pad is usually placed on the lower extremity, cervical RFA theoretically carries a greater risk of damage to implanted devices because the current will likely pass through the torso before exiting at the grounding pad. It is recommended that the grounding electrode be placed at least 15 cm away from pacing leads for both permanent pacemakers and implantable cardiac defibrillators.\textsuperscript{378} If using monopolar RF, placing the grounding pad close to the neurotomy site will reduce the size of the induced electromagnetic field. This will minimize the risk of heating the neurostimulator battery and electrodes. However, placing the grounding pad too close to the neurotomy site can increase the risk of tissue burns, particularly when using high current, long activation times, and the use of conductive fluid, since the energy has less tissue through which to dissipate.\textsuperscript{29}

A study from 1995 reported on the pacemaker activity in 25 patients with 13 different devices, most of whom underwent RFA with monopolar electrodes. The authors observed sensing failures in eight (32.0%) and pacing failures in four (16.0%) patients.\textsuperscript{379} Prolonged pauses and induction of tachyarrhythmias were also detected. Although no pacemaker damage was reported, permanent damage has been described by other investigators in non-peer-reviewed literature.\textsuperscript{380} In a recent position paper by SIS,\textsuperscript{381} the authors recommended consultation with the device manufacturer and cardiologist or electrophysiologist, with consideration of the following: (1) having on-site support for interrogation of the cardiac device during the procedure; (2) placing a magnet over the device to inhibit triggering the device by RFA; and (3) removing the magnet or use external defibrillator/pacing electrodes should cardiac arrhythmias occur during the RFA procedure.

**Recommendations**

Guidelines should be reviewed,\textsuperscript{382} and healthcare teams responsible for managing the implanted devices (eg, neurology, cardiology/electrophysiology, pain medicine) should be consulted regarding the planned cervical RFA procedure. If RFA is performed, in the absence of evidence to the contrary, implanted electrical devices such as neurostimulators should be programmed to an output of zero volts and turned off before the procedure, and the risks of device damage discussed with the patient. For pacemakers and defibrillators, we recommend a shared-decision model, with the cardiology team and device manufacturer being consulted prior to medial branch RFA and their recommendations followed (eg, program pacemaker to asynchronous mode). Given the inherent risks of using bipolar electrodes (eg, additional trauma, aggressive, less predictable lesions) in the cervical spine, we cannot routinely recommend it at this time. Using no or judicious sedation will allow the physician to communicate effectively with the patient and detect any potential injury to the nervous system or cardiovascular decompensation at an early stage. A deactivated neurostimulator should be turned back on following the RFA procedure and reprogrammed to pre-procedural settings; grade C recommendation, low level of certainty.

**Tissue burns**

The musculature and skin overlying the cervical facet joints is usually thinner than around the lumbar spine. Skin burns from equipment malfunction (eg, incorrect application of the grounding pad) or extension of the RFA lesion into the dermis in areas such as the knee where there is less tissue between the target nerve and the skin have been reported.\textsuperscript{383–385} Given the reduced soft tissue in the cervical region, there is a possibility this could occur during cervical medial branch RFA. Extension of the RFA lesion into cervical paraspinal muscles may manifest as increased procedure-related pain.\textsuperscript{374} Use of larger grounding pads and thinner RF cannulas (22 or 20 gauge) with shorter active tips (5 mm) for high-risk procedures may mitigate the risk of cutaneous burns.\textsuperscript{386}

**Recommendations**

A pre-procedure check of all RFA equipment to ensure that it is properly functioning, and positioning grounding pads in an optimal location and orientation is recommended. Applying a large, properly positioned grounding pad on a dry, clean-shaven skin that is devoid of scars or tattoos may minimize the risk of tissue burns; grade C recommendation, low-to-moderate level of certainty.

**Impact of RFA on spinal instrumentation in proximity of the procedure**

Anterior approaches for fusing the cervical spine are more frequently used than in the lumbar spine. However, it is not uncommon for patients who have had cervical spine surgery with instrumentation to undergo facet joint RF denervation at levels adjacent to the operated segments. In a retrospective study evaluating cervical medial branch RFA outcomes, Cohen et al found that 59% of 29 patients with prior cervical spine surgery experienced a positive outcome, which was no different than the success rate in surgically-naïve patients.\textsuperscript{29} No serious complications were noted in these patients. In another retrospective study on patients with hardware in the posterior cervical elements, Ellwood and colleagues\textsuperscript{316} reported no complications following 56 ablations performed in 36 patients, including 11 who underwent cervical RFA. However, concerns have been raised that the use of RFA in patients with existing posterior spinal instrumentation can cause thermal injury to surrounding structures through heating of the hardware.\textsuperscript{387} Yet, as noted and recommended in the multi-society consensus guidelines for lumbar facet joint procedures,\textsuperscript{31} there is no reported evidence of superficial or deep burns, denervation of the lateral branches or ventral rami, or coagulation of blood vessels with lumbar facet joint RFA procedures in patients with posterior spinal instrumentation.

**Recommendations**

Multiplanar fluoroscopic image-guided techniques should be used to ensure that the RF cannula is not in contact with the pedicle screw in patients with posterior cervical instrumentation to avoid thermal injury to surrounding tissues. Depending on the type of surgery performed, consideration should also be given to the probability of accurately targeting cervical medial branches in the presence of spinal hardware placed using a posterior approach; grade C recommendation, low level of certainty.

**QUESTION 19: IN WHICH PATIENTS SHOULD REPEAT RFA BE CONSIDERED AND WHAT IS THE LIKELIHOOD FOR SUCCESS?**

**DO REPEAT DIAGNOSTIC BLOCKS NEED TO BE REPEATED AND, IF SO, AT WHAT INTERVAL?**

**Rationale for repeating RFA and defining a positive outcome**

Pain relief after cervical medial branch RFA has shown durability but not permanence, with the median time to pain recurrence ranging from 6 to 14 months (table 21). In clinical practice, RFA is commonly repeated when pain returns. Pain physicians
Successful outcome:

endpoint. The 3-guidelines recommendation of ≥2-ized controlled lumbar facet RFA trial, ACTTION guidelines on .

Results: Success Duration of benefit

<table>
<thead>
<tr>
<th>Author, year</th>
<th>Patients</th>
<th>Joints included</th>
<th>Design</th>
<th>Successful outcome</th>
<th>Active group initial RFA failure: 2 of 5 (40%) had successful procedure Control group: 4 of 5 (80%) had success</th>
<th>Median duration for initial RFA 263 days vs 253 days for repeat RFA</th>
</tr>
</thead>
<tbody>
<tr>
<td>Lord et al 1998</td>
<td>24 pts diagnosed with cervical facet pain based on three blocks including one with saline</td>
<td>C3–C7</td>
<td>Randomized, placebo-controlled</td>
<td>Complete pain relief for ≥90 days. Duration of benefit was time for pain to return to ≥50% of baseline</td>
<td>C2–3; Initial RFA success rate: 4 of 10 (40%) Repeat RFA success rate: 2 of 2 (100%) C5–C7: Initial RFA success rate: 7 of 10 (70%) Repeat RFA success rate: 3 of 3 (100%)</td>
<td>Median duration of initial successful RFA 161 days vs 237 days for repeat RFA</td>
</tr>
<tr>
<td>Lord et al 1995</td>
<td>19 pts with chronic cervical facet pain after MVC diagnosed by dual blocks</td>
<td>C2–C7</td>
<td>Retrospective</td>
<td>Complete relief for a clinically significant duration</td>
<td>Initial RFA success (pts): 5 of 12 (42%) Repeat RFA success rate (procedures): 10 of 13 (77%)</td>
<td>Median duration of initial successful RFA 217 days for repeat RFA</td>
</tr>
<tr>
<td>Lord et al 1998</td>
<td>12 pts with cervical facet pain from C2–3 after MVC diagnosed by dual comparative or placebo-controlled blocks</td>
<td>C2–3</td>
<td>Retrospective</td>
<td>Complete relief in the region treated for ≥90 days. Duration of benefit was time for pain to return to ≥50% of baseline</td>
<td>Initial RFA success (pts): 5 of 12 (42%) Repeat RFA success rate (procedures): 10 of 13 (77%)</td>
<td>Median duration of initial successful RFA 217 days for repeat RFA</td>
</tr>
<tr>
<td>McDonald et al 1999</td>
<td>28 pts with cervical facet pain from C2–3 after MVC diagnosed by dual comparative or placebo-controlled blocks</td>
<td>C3–C7</td>
<td>Retrospective</td>
<td>Complete relief for a clinically significant duration</td>
<td>Initial RFA success rate: 18 of 28 (64%) Repeat RFA success rate: 7 of 8 (87.5%) Initial RFA failure: 10 of 28 (36%) Repeat RFA success rate after initial RFA failure: 2 of 6 (33%)</td>
<td>Median duration of initial successful RFA 421 days vs 181.5 days for first repeat RFA. Median duration of all successful repeat RFA: 218.5 days</td>
</tr>
<tr>
<td>Govind et al 2003</td>
<td>49 pts with complete pain relief with dual comparative blocks (2 pts with bilateral pain)</td>
<td>TON/C2–3</td>
<td>Retrospective</td>
<td>Complete relief of headache and restoration of function for 90 days</td>
<td>Initial RFA success rate: 43 of 49 (88%). Repeat RFA success rate: 12 of 14 (86%)</td>
<td>Mean duration of relief after initial successful RFA: 297 days. Mean duration of relief after repeat RFA: 217 days</td>
</tr>
<tr>
<td>Barnsley, 2005</td>
<td>35 pts with complete relief of pain with dual comparative blocks and no relief with placebo injection. 12 pts received repeat RFA</td>
<td>C2–C7</td>
<td>Retrospective</td>
<td>Complete relief of neck pain</td>
<td>Initial RFA success rate: 26 of 35 (74%) Repeat RFA success rate: 8 of 9 (89%) Initial RFA failure rate: 7 of 33 (21%) Repeat RFA success rate after failure: 2 of 3 (67%)</td>
<td>Mean duration of relief after initial successful RFA: 36 weeks Duration of relief after repeat successful RFA not reported</td>
</tr>
<tr>
<td>Husted et al 2008</td>
<td>22 pts with ‘good’ relief from 2 MBBs and ≥50% relief after RFA</td>
<td>C2–C7</td>
<td>Retrospective</td>
<td>≥50% pain relief</td>
<td>Initial RFA success rate: 22 of 22 (100%) Repeat RFA success (procedures): 39 of 41 (95%) 21 pts had a second RFA 11, 4, 2, 1 patients had a third, fourth, fifth, sixth, and seventh repeat procedure</td>
<td>Mean duration of initial successful RFA: 12.5 months. Mean duration for subsequent successful RFA: 11.5 months</td>
</tr>
<tr>
<td>Rambaransingh et al 2010</td>
<td>73 pts with cervical and lumbar pain who received ≥3-point pain reduction (on 0–11 scale) with RFA. 15/73 pts underwent repeat cervical RFA</td>
<td>Levels unreported</td>
<td>Observational</td>
<td>Improvement in pain and disability</td>
<td>Pain reduction and functional improvement remained significantly improved from baseline with each repeat procedure. 43% success rate (≥50% improvement) vs 64% with second RFA and 86% with third RFA</td>
<td>Mean duration of relief after initial successful RFA: 9.9 months. Mean duration of relief after second successful RFA: 10.5 months (not reported for third RFA)</td>
</tr>
<tr>
<td>Spelda, 2011</td>
<td>151 pts with ≥80% relief from dual comparative MBB blocks. 26 pts underwent repeat RFA, 47 procedures</td>
<td>Levels unreported</td>
<td>Observational</td>
<td>≥50% pain reduction for ≥24 months</td>
<td>Initial RFA success: 112 of 151 (76%) Repeat RFA success rate after initial successful RFA (procedures) 34 of 40 (85%) Repeat RFA success rate after initial failure: 4 of 7 (57%)</td>
<td>Mean duration of success after initial RFA 12 months vs 9.7 after repeat RFA</td>
</tr>
<tr>
<td>MacVicar et al 2012</td>
<td>104 pts with complete relief of pain from dual comparative blocks</td>
<td>C2–7</td>
<td>Observational</td>
<td>Successful outcome: ≥80% pain relief for at least 6 months with complete restoration of ADLs</td>
<td>Initial RFA success rate: 69 of 104 (66%) Repeat RFA success rate after initial success: 23 of 24 (96%) Repeat RFA success rate after repeat failure: 2 of 2 (100%)</td>
<td>Median duration of pain relief after initial RFA 17 months and 20 months in two practices. Median duration of pain relief after repeat RFA 15 months</td>
</tr>
</tbody>
</table>

ADLs, activities of daily living; MVC, motor vehicle collision; pts, patients; RFA, radiofrequency ablation.

should educate patients about the duration of expected relief and the potential need for repeated treatment(s). To discuss the efficacy of repeat cervical RFA, ‘success’ needs to be defined. A successful RFA must result in a meaningful reduction of pain for a clinically relevant period of time. The definitions vary per study (see table 21) but, overall, most studies require ≥50% pain relief for ≥3 months. However, more recent randomized studies evaluating lumbar medial branch RFA have used the IMMPACT guidelines recommendation of ≥2-point (or ≥30%) decrease in average pain, with 3 months designated as the primary endpoint. The 3-month time point stems from patient and physician surveys performed in preparation for a randomized controlled lumbar facet RFA trial, ACTTION guidelines on unique aspects of interventional clinical trials, the MINT and FACTS trials evaluating lumbar medial branch RFA, and FDA guidance on randomized trials evaluating pharmaceutical treatments.

Likelihood of success with repeat cervical medial branch RFA

The results from multiple studies are remarkably consistent, with an average of 84% of patients obtaining a successful repeat RFA after an initial successful RFA. In these individuals, the duration of benefit ranges between 7 and 20 months. This average differs slightly from the systematic review and meta-analysis by Smuck et al which reported a non-weighted
88% average success rate (range 67–95%) in previous studies when the first RFA was successful, but a success rate of only 38% (range 0–67%) when the initial RFA was unsuccessful. This discrepancy results from inappropriate meta-analysis application which included studies by Lord et al, Lord et al, and McDonald et al in which some duplicate patients were included. The results of denervation of the C2–3 joint appear to be comparable to outcomes with lower cervical levels in that 77–84% of patients respond with ≥50% pain relief after repeat RFA for 7.2–7.9 months (table 2).

Number of times RFA can be repeated at the same level(s)
Considering that many patients experience neck pain and undergo cervical medial branch RFA after MVCs, questions about the likelihood of success for repeat procedures and how many times procedures should be repeated have major relevance for litigation attorneys and insurance companies. No studies have directly examined the number of times that RFA can be repeated at the same level. Speldewinde et al followed 26 patients who underwent repeat treatment up to five times. Although the authors did not break down the success rate for each procedure, they reported an average duration of 9.7 months among subsequent RFA successes. McDonald et al described four patients who underwent multiple repeat procedures, with the benefit from each procedure lasting for at least 90 days. Husted et al followed a group of 22 patients who had up to seven repeat RFAs, with each repeat RFA having a 90–100% success rate and an average duration of benefit of 11.5 months. Rambaransingh et al reported the results of repeat cervical medial branch RFA in 14 patients (nine of whom underwent a third RFA), noting a significant decrease in pain intensity, pain frequency, and disability in pre- and post-treatment measurements. These improvements were similar in magnitude for each RFA, suggesting no or minimal loss of effectiveness with repeat procedures.

Decision to repeat RFA determined by initial RFA response
In general, the decision to repeat a procedure should depend on the magnitude of pain, results of the previous procedure(s), and the concordance of location and quality. In patients who undergo repeat cervical medial branch RFA after an initially successful procedure, McDonald et al noted the initial median duration of pain relief to be 14 months, which compared favorably with a 7.3-month median duration of relief for repeated successful procedures. Although this represents a meaningful attenuation in the duration of benefit for subsequent RFAs, several studies have reported more modest declines. Speldewinde et al reported the mean duration of benefit for subsequent RFAs to be 9.7 months, which was less than the 12-month duration of the initial procedure. MacVicar et al reported that the median duration of benefit after initial successful RFAs was 17 and 20 months in two private practice groups. Successful repeat cervical medial branch RFA procedures resulted in a median duration of benefit of 15 months in both groups. Govind et al noted a small decrement in effect, with the initial TON RFA lasting a mean of 9.9 months and the repeat RFA lasting an average of 7.2 months. In a study by Husted et al the authors reported a mean duration of initial benefit of 12.5 months, which was only slightly longer than the 11.5-month mean duration of benefit after subsequent procedures.

Other authors have reported longer benefits after repeat procedures compared with initial ones. Rambaransingh et al reported a mean duration of 9.9 months pain relief after initially successful lumbar and cervical medial branch RFA versus 10.5 months of relief after a repeat procedure. In the Lord et al study evaluating TON RFA, 12 patients had a median duration of benefit of 5.4 months after the initial procedure versus 7.9 months after repeat denervation. The variability in the duration of response to repeat cervical facet RFA in comparison to the initial denervation is likely due to several factors including technical details (needle placement, lesion size, duration of recurrence of pain prior to repeat RFA), progression of degenerative changes since the last treatment (eg, worsening facet arthropathy), and potential differences in the calculations of duration of benefit (ie, inclusion or exclusion of treatment failures).

Rationale for repeating prognostic blocks before repeat RFA
Whereas all studies specified that a positive response to prognostic MBB was a prerequisite for RFA, no studies were identified in which the prognostic MBB was repeated prior to subsequent RFA. Lord et al noted that, if an RFA was unsuccessful, repeat prognostic blocks could be used to test the technical success of the RFA if performed at the same levels. Likely, repeat MBBS at the levels of a previously successful RFA have not been studied in part due to the established high success rate of repeat RFA. However, if the patient’s pain has changed in location or quality and/or the examination is no longer similar to the pre-RFA baseline, a repeat prognostic MBB can be considered. If an initial RFA is unsuccessful and is not attributable to technical failure, repeat MBB at a different location may be warranted.

Initial and repeat procedure interval
Cohen et al identified no significant difference in the baseline duration of cervical pain between patients with a successful outcome and those who failed RFA treatment. However, in multicenter studies by the same group of authors, a shorter duration of pain correlated with better outcomes in studies examining predictors of response for lumbar medial branch RFA and sacroiliac joint lateral branch RF denervation. No studies were identified which assessed the effect of pain duration on the success of repeat cervical medial branch RFA. Prolonged denervation of the paraspinous musculature after lumbar medial branch RFA has been shown to last over 12 months in some patients. The high success rate of repeat cervical medial branch RFA
after a successful initial RFA treatment suggests that subsequent procedures may be performed shortly after recurrence of pain. However, RFA should never be performed pre-emptively since prolonged denervation of muscles may lead to irreversible interstitial fibrosis and the replacement of myocytes by adipocytes.

Recommendations
The committee recommends that if an initial RFA is successful (clinically meaningful pain relief for ≥3 months), it should be offered to patients who experience return of their cervical pain, assuming that the pain is similar in character and location to the initial pain; grade B, moderate level of certainty. Given the mean duration of benefit and drop-off in success rates noted in some studies with repeat RFA, we recommend repeating the procedure no more than two times a year; grade B, moderate level of certainty. Although there are a paucity of studies examining cervical medial branch RFA success rates after repeating prognostic blocks, the high success rates reported after empirically repeating RFA indicate they are not necessary in most people; grade C, low level of certainty.

QUESTION 20: SHOULD THERE BE DIFFERENT STANDARDS IN SELECTING PATIENTS FOR RFA IN CLINICAL TRIALS AND CLINICAL PRACTICE?
Clinical trials are the reference standard to determine the efficacy and effectiveness of novel therapeutics to treat pain, including invasive therapies such as cervical facet treatments. The current push for more evidence-based medicine has caused invasive therapies for pain to come under intense scrutiny due to a lack of evidence for efficacy. Invasive therapy trials face numerous challenges including difficulties with sham control groups, blinding, cost, and an inability to enroll subjects. This has resulted in invasive therapies bypassing the early phase clinical trial model for efficacy and jumping to ‘real world’ patient populations that reflect effectiveness. Whereas early phase clinical trials typically employ strict selection criteria to reduce variables that may affect outcomes, later phase selection criteria are loosened (as more is learned about the treatment) to evaluate outcomes in ‘real world’ patient populations. Differences between clinical research and clinical practice for invasive therapies such as cervical facet interventions may be more accentuated compared with those between medication prescribing and industry-sponsored drug trials. Understanding these differences is necessary to determine both the efficacy and clinical effectiveness of cervical facet therapies.

Compared with lumbar procedures, cervical procedures are associated with greater technical difficulties and risks. Balancing the risk/benefits of cervical facet blocks and RFA in clinical trials may result in more stringent selection criteria compared with clinical practice, which takes into consideration physician preference and individual factors (personalized medicine). This guideline critically evaluates the literature which was used to develop guidelines to inform clinical practice, which was subsequently modified to come up with recommendations for clinical trials. Whereas most recommendations show no differences between the two, for others there are distinctions. Table 22 summarizes the differences in recommendations between clinical trials and clinical practice. Details for the basis of these recommendations can be found in the previous sections which provide a thorough review of the literature for each question.

Patient selection for diagnostic blocks
Similar to all chronic pain treatments, patient selection for diagnostic cervical MBB plays a critical role in determining the likelihood of a positive outcome. There is consensus that failure of at least 3 months of conservative therapy is a reasonable threshold that should be implemented in both clinical trials and practice, although practice guidelines could allow for flexibility in extenuating circumstances. There is agreement that the natural course of acute pain (including neck) is favorable, allowing conservative therapies in the first 3 months to facilitate recovery. However, prospective studies on the course of neck pain indicate that up to 40% of patients who do not respond to conservative treatment still suffer neck pain at 12 months.

There are no physical examination signs that reliably predict response to facet joint blocks, though paraspinal tenderness and pain with certain movements (which are limited by the facet joints) may be weakly associated with cervical facetogenic pain. However, some studies have delineated referred pain patterns of individual facet joints and may improve the identification of putatively painful joints. Therefore using pain referral maps and/or standardized tests such as algometry for paraspinal tenderness should be considered for clinical trials. Bilateral neck pain is often more multifactorial than unilateral pain and should be excluded from clinical trials unless the study is specifically designed to address this clinical problem or is properly powered for subgroup analysis. As with all clinical trials, mental health exclusion criteria to identify people with existing anxiety, depression, or post-traumatic stress should be more stringent in efficacy trials than in clinical practice due to the negative influence that these comorbid conditions have on pain outcomes.

MBB cut-offs and patient-reported outcomes
Percent pain reduction after a diagnostic MBB is an important indicator of facet pain. Although there is no clear evidence on what threshold should be used, consensus supports ≥50% pain reduction, and this applies to both clinical trials and practice, possibly with a higher cut-off considered for whiplash injuries and younger age groups enrolled in studies. For clinical trials, since the magnitude of pain relief is likely to be greater for MBB than RFA, using higher cut-offs may theoretically lead to improved outcomes, but the clinical evidence does not support this consideration. Despite interest in using non-pain measures (such as function) as selection criteria, there is no evidence to support any modality as a sole criterion and using them is not feasible in clinical practice to evaluate short-term MBB results. However, in clinical trials, these measures may be useful to evaluate in exploratory analyses using a composite score that includes pain relief. When selecting outcome measures for clinical trials evaluating RFA and in accordance with IMMPACT guidelines, non-pain outcome domains should be measured. Since these outcomes were chosen because they are important to patients and for physicians to evaluate success, simple non-pain outcome measures such as function, satisfaction, and healthcare utilization (including analgesic consumption) may be assessed in clinical practice.

Injection technique
Injection technique is important to increase the chances of precision delivery to the targeted structure(s). There are only two recommended differences concerning technique. First, AO/AA injections require fluoroscopic guidance with an injection of contrast. DSA has been shown to detect vascular uptake that would otherwise be missed with fluoroscopy alone. However, DSA is not widely available and increases radiation exposure.
The consequences of inadvertent vascular injection for upper joint injections include false-negative results and seizures.403

Number of MBBs
The use of dual blocks may increase the RFA success rate. However, this will invariably result in false-negative blocks and deny some patients the benefit of an RFA. There are also patient inconveniences and increased costs and risk exposure to consider. Nonetheless, for clinical trials that aim to show efficacy, the benefits of a higher percentage of true positives (enriched population) may outweigh the cons and should be considered in trial design.

### RFA technique and considerations
RFA cannula placement is critical to a successful outcome. There is widespread agreement that both anteroposterior and posteroanterior; RFA, radiofrequency ablation; TON, third occipital nerve; US, ultrasound.

#### Table 22 Guidelines for clinical trials versus clinical practice

<table>
<thead>
<tr>
<th>Table 22</th>
<th>Guidelines for clinical trials versus clinical practice</th>
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</thead>
<tbody>
<tr>
<td>Factor</td>
<td>Clinical trial</td>
</tr>
<tr>
<td><strong>Patient selection</strong></td>
<td></td>
</tr>
<tr>
<td>History and physical examination of cervical facet joints</td>
<td>Inclusion criteria: ► Paraspinal muscle tenderness, ideally under fluoroscopy ► Pain with movement (eg, extension, rotation, lateral flexion) ► Pain consistent with referral maps Exclusion criteria: ► Radicular symptoms ► Bilateral neck pain unless study adequately powered ► Active psychological comorbidities</td>
</tr>
<tr>
<td>Failure of conservative treatment</td>
<td>No recommendation</td>
</tr>
<tr>
<td>Radiological findings for facet joint pain</td>
<td>No recommendation</td>
</tr>
<tr>
<td>Patient reported outcomes</td>
<td>Follow IMMPACT and other relevant guidelines, more detailed than clinical practice</td>
</tr>
<tr>
<td>Pain relief cut-off for positive MBB</td>
<td>≥50% (consider higher cut-off for efficacy trials or subgroup analysis)</td>
</tr>
<tr>
<td>Functional measures</td>
<td>Sole criterion not recommended for assessing MBB results (composite with pain relief should be considered for RFA effectiveness)</td>
</tr>
<tr>
<td>Repeat RFA</td>
<td>≥30% for at least 3 months per IMMPACT and lumbar facet guidelines281</td>
</tr>
<tr>
<td>Repeat diagnostic MBB for repeat RFA</td>
<td>No – previous RFA should be an exclusion criterion for studies not evaluating repeat RFA</td>
</tr>
<tr>
<td>Injection technique</td>
<td></td>
</tr>
<tr>
<td>AO and AA imaging</td>
<td>Pre-injection CT or MRI, fluoroscopy and real-time contrast injection with strong consideration of DSA or CT</td>
</tr>
<tr>
<td>Approach</td>
<td>Posterior/posterior oblique</td>
</tr>
<tr>
<td>Volume and prognostic test</td>
<td></td>
</tr>
<tr>
<td>Medial branch block</td>
<td>≤ 0.3 mL</td>
</tr>
<tr>
<td>Diagnostic block (IA vs MBB)</td>
<td>MBB</td>
</tr>
<tr>
<td>Diagnostic block (single vs dual)</td>
<td>Single block (consider dual blocks only for efficacy studies)</td>
</tr>
<tr>
<td>Imaging</td>
<td>Fluoroscopy or US</td>
</tr>
<tr>
<td>Approach</td>
<td>Lateral (TON, C3–C7)</td>
</tr>
<tr>
<td>Caudal nerve block</td>
<td>Posterior oblique (C8)</td>
</tr>
<tr>
<td>Volume and steroid use for AA/AO injections</td>
<td>Non-particulate ≤1 mL</td>
</tr>
<tr>
<td>Sedation</td>
<td>Avoid</td>
</tr>
<tr>
<td>RFA technique</td>
<td></td>
</tr>
<tr>
<td>Stimulation</td>
<td>Motor for all levels</td>
</tr>
<tr>
<td>Needle orientation</td>
<td>Parallel (preferable) or near-parallel</td>
</tr>
<tr>
<td>Lesion size</td>
<td>No recommendation due to unknown risk/benefit</td>
</tr>
<tr>
<td>Cannulae confirmation</td>
<td>PA/lateral and possibly contralateral oblique</td>
</tr>
<tr>
<td>Implanted devices</td>
<td>Exclude patients</td>
</tr>
<tr>
<td>Grounding pad placement</td>
<td>Dry, hairless skin devoid of tattoos</td>
</tr>
<tr>
<td>Spinal hardware</td>
<td>Exclude patients unless specifically addressing outcomes in this population</td>
</tr>
<tr>
<td>Repeat RFA</td>
<td>No more than twice/year</td>
</tr>
<tr>
<td>Post-lesion steroids</td>
<td>No steroids for efficacy study unless administered to all patients</td>
</tr>
<tr>
<td>Anticoagulation</td>
<td>Exclude subjects who cannot stop anticoagulants</td>
</tr>
</tbody>
</table>

AA, atlanto–axial; AO, atlanto–occipital; CT, computed tomography; DSA, digital subtraction angiography; IA, intra-articular; MBB, medial branch block; MRI, magnetic resonance imaging; PA, posteroanterior; RFA, radiofrequency ablation; TON, third occipital nerve; US, ultrasound.
lateral views are necessary to verify needle location. A contra-lateral oblique approach provides a third view that may increase precision and confirm that the cannula tip is not in the neural foramen. Its use may be particularly valuable when performing lower cervical procedures in obese patients. However, additional views increase procedure time and radiation exposure and may not be feasible for all cases in clinical practice.

Implanted devices and spinal hardware can complicate RFA procedures as the electrical current can affect the devices and contact with the spinal hardware can lead to thermal injury. Yet, precautions can be taken to reduce the likelihood of affecting the implanted device. Having prior surgery may also increase the rate of false-positive MBB and treatment failure. Due to these considerations, it is recommended that patients with implanted devices and spinal hardware be excluded in clinical trials. The exception is for clinical trials specifically evaluating RFA in patients with spinal hardware.

The use of steroids to prevent neuritis has not been studied in the cervical spine, although the risk is at least as high as in the lumbar spine at higher levels. Given the risks of repeated steroids which may include immune suppression and bone demineralization, in clinical practice their pre-emptive use should be made after a careful risk:benefit analysis. Since a small percentage of patients may experience prolonged benefit from MBB performed with LA and steroids, we recommend standardization in clinical trials (ie, either avoiding steroids in all research subjects or using them in everyone).

Anticoagulation
The vasculature of the neck is more extensive than in the lumbar region with a higher risk of penetrating vascular structures and bleeding. Therefore, unlike in the lumbar region where continuation is routinely recommended, the decision to continue or discontinue anticoagulants should be made only after a careful risk:benefit assessment for joint blocks and RFA. Whereas a risk to benefit analysis is justified in clinical practice, it is recommended that these patients be excluded in clinical trials (table 22).

Recommendations
This committee acknowledges that employing different standards for clinical practice and clinical trials for some elements of selection and performance, particularly those that purport to show efficacy, is reasonable. These differences reflect the different goals for investigators, patients, and physicians. Specific areas in which criteria may differ include: (1) patient selection for cervical MBB and RFA (with clinical practice erring on the side of enhanced access to care); (2) more non-pain secondary outcome measures for clinical trials than in clinical practice; (3) consideration of the requirement for DSA when evaluating AO/AA injections in clinical trials; (4) requiring the contralateral oblique approach to improve precision in clinical trials; (5) excluding patients with implanted devices and spinal hardware for clinical trials evaluating cervical medial branch RFA; and (6) excluding patients on anticoagulation in clinical trials to reduce risks and dropouts. Grade B recommendation, moderate level of certainty for differences in patient selection and non-pain secondary outcome measures. Grade C, low level of certainty for requiring a contralateral oblique view and excluding patients with implanted hardware, on anticoagulants, and requiring consideration of DSA for clinical trials (table 23).

DISCUSSION
Effect of perspective
These guidelines were developed to serve as a roadmap for pain physicians who treat cervical joint pain. There are wide variations in the conclusions of experts from different specialties on the effectiveness of pain management procedures, with trials and reviews performed by those who perform procedures much more likely to yield positive results than those performed by non-interventional pain and spine physicians. For neck pain and cervical facet interventions, there are much fewer studies and reviews conducted by non-specialists compared with the literature on LBP. However, among reviews that have been published, similar discrepancies exist in that those performed by pain specialists are more likely to recommend treatments as effective. Reasons for incongruities include conscious and unconscious bias, differences in study interpretation which reflect variances in background and understanding, and perhaps a better ability for pain management physicians to select appropriate candidates and perform complex procedures.

Approval of participating organizations
All participating organizations except for one (North American Spine Society) approved or supported the document, with their representative (BJS) being the only committee member to dissent on any question (n=3). The North American Spine Society did not vote on individual questions. The American Academy of Physical Medicine and Rehabilitation approved 18 questions, abstained on two questions (laterality and differences between clinical trials and practice), and affirmed the value of the document. The American Academy of Neurology approved all questions and also affirmed the value of the document. The American Society of Anesthesiologists supported the document and approved of all questions, but since the document was not voted on by the Board of Directors or House of Delegates, the organization could not officially endorse the document. The Spine Intervention Society dissented on two questions (pain relief cut-off and number of diagnostic facet blocks) and abstained on three (utility of stimulation, differences between clinical trials and practice, and risk mitigation) but approved the document. As noted previously, the Departments of Defense and Veterans Affairs did not vote on the document, but each organization received approval from their pain medicine leadership (table 24).

Differences between the lumbar and cervical facet guidelines
Aside from the inclusion of interventions targeting the AO and AA joints, there were several differences between recommendations for lumbar facet procedures and those in the cervical spine that warrant mentioning. Because of shorter distances (ie, less depth from the skin to the anatomic target), different trajectories, and a higher incidence of critical vessels overlying the cervical facets, US may be more useful for cervical MBB than for lumbar facet blocks, although fluoroscopy is recommended for RFA in both regions. Routine advanced radiological imaging (eg, MRI or CT) was not deemed essential before lumbar facet blocks, and given that there is a higher prevalence of facetogenic pain in the neck than the back, one might have expected the same in the cervical region. However, advanced radiological imaging was conditionally recommended before upper cervical facet procedures to avoid vascular injuries in cases of anatomic variations.

Two of the most controversial questions in both the lumbar facet joint guidelines and this guideline revolved around the number of prognostic blocks and cut-offs for designating a block
as positive. Recommending two blocks in the neck but one in the lumbar spine would have been inconsistent considering that the higher published prevalence rate of facetogenic pain in the neck compared with the low back should decrease the risk of a false-positive block. In contrast, the more prominent role of the facet joints in neck pain than LBP theoretically could have led to recommending a cut-off for a positive block above 50% pain relief, but neither the studies that stratified RFA outcome by prognostic block pain relief nor the IMMPACT guidelines supported a higher threshold recommendation.19 20 23 281 285

The risks of AO, AA, and facet procedures including bleeding complications and injury to aberrant vasculature and surrounding nerves (eg, cervical nerve roots) are greater in the cervical spine than in the lumbar spine. Minimizing these risks resulted in recommendations to consider reviewing radiological or US imaging before embarking on cervical joint procedures, perform RFA procedures one side at a time to reduce the risk of cervical neuritis, and a short course of NSAIDs may reduce post-RFA pain; Grade C, low level of certainty. There is inconsistent evidence supporting the peri-procedural use of gabapentin; Grade C, low level of certainty. Discontinue anticoagulants prior to intervention as indicated; Grade C, low level of certainty. There is inconsistent evidence supporting the use of sedation for diagnostic or therapeutic IA injections; Grade C, low level of certainty.

<table>
<thead>
<tr>
<th>Question</th>
<th>Recommendation</th>
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<tbody>
<tr>
<td>Use of history and physical examination to identify painful AO or AA joints or to select patients for injections</td>
<td>History and physical examination cannot reliably identify painful AO or AA joints, but can guide injection decisions which could confirm the AO and AA joints as pain generators; Grade C, low level of certainty.</td>
</tr>
<tr>
<td>Use of history and physical examination to identify painful cervical facet joints and select patients for blocks</td>
<td>No pathognomonic historical or examination signs can reliably predict response to facet joint blocks in individuals with chronic neck pain; Grade C, low level of certainty</td>
</tr>
<tr>
<td>Correlation between radiological findings and prognostic block or RFA outcomes</td>
<td>The evidence is insufficient to assess the utility of radiological imaging for diagnosing cervical facet joint pain and as a prognostic indicator for the success of cervical facet blocks or RFA; Grade I. For procedural planning, radiological imaging should be considered when indicated; Grade C, low level of certainty</td>
</tr>
<tr>
<td>Conservative treatment requirement before cervical facet blocks</td>
<td>Conservative therapy should be used before prognostic blocks in those with chronic neck pain; Grade B, moderate level of certainty. The trial should be for 6 weeks, although this may vary based on a personalized approach; Grade C, low level of certainty. Grade I for concomitant use of conservative measures to accompany prognostic blocks</td>
</tr>
<tr>
<td>Image guidance for cervical facet blocks and RFA</td>
<td>Fluoroscopy or US should be used for cervical MBB; Grade A recommendation, moderate level of certainty. Fluoroscopy (vs. CT) should be used for IA injections; Grade C, low level of certainty. Fluoroscopy should be used before RFA; Grade A, high level of certainty for imaging. Grade B, moderate level of certainty for use of fluoroscopy</td>
</tr>
<tr>
<td>Optimal technique for AO and AA joint injections and risk mitigation</td>
<td>Advanced imaging should be obtained before injections; Grade C, low level of certainty. Posterior approach with real-time fluoroscopy or US in multiple views; Grade B, moderate level of certainty. Grade C, low level of certainty for steroid use; if steroids are injected, &lt;1 mL non-particulate steroids should be used</td>
</tr>
<tr>
<td>Approach for cervical MBB</td>
<td>A fluoroscopically-guided lateral approach should be considered for TON and C3–C7 MBB, but a posterior or oblique oblique approach should be used for C8 MBB. The smallest needles possible should be used; Grade I</td>
</tr>
<tr>
<td>Volumes for cervical MBB and IA injections</td>
<td>Volumes ≤0.3 mL should be used for cervical MBB and ≤1 mL for IA injections; Grade C, low level of certainty for MBB; Grade C, low level of certainty for IA injections</td>
</tr>
<tr>
<td>Therapeutic value of cervical MBB and IA injections</td>
<td>Therapeutic IA injections should not be routinely used; Grade C, low-to-moderate level of certainty. Grade D recommendation, moderate level of certainty against use of steroids during MBB</td>
</tr>
<tr>
<td>Performing bilateral cervical MBB and RFA, and limits on the number of levels treated</td>
<td>Bilateral MBB may be performed at one session, but bilateral RFA should be avoided. Treating &gt;2 spinal levels (≥3 nerves) during one session should not be done routinely; Grade C recommendation, low level of certainty</td>
</tr>
<tr>
<td>Diagnostic and prognostic utility of cervical injections</td>
<td>Cervical MBB meet most criteria for a diagnostic intervention, while IA injections meet full criteria but carry a high technical failure rate; Grade C, low-to-moderate level of certainty. MBB may be more predictive of RFA outcomes than IA injections; Grade C, low level of certainty. AO and AA local anesthetic injections may be diagnostic and predictive for steroid injections; grade C, low level of certainty</td>
</tr>
<tr>
<td>Use of sedation</td>
<td>Sedation should not routinely be given for diagnostic procedures; Grade B, moderate level of certainty</td>
</tr>
<tr>
<td>Cut-off for designating a MBB as positive and use of non-pain measures</td>
<td>≥50% pain relief should be used as the cut-off to maximize access to care; Grade C, low-to-moderate level of certainty. Non-pain measures can be used in conjunction with pain relief, but not as the sole criterion for designating a block as positive; Grade B recommendation, moderate level of certainty</td>
</tr>
<tr>
<td>Number of blocks before RFA</td>
<td>A single block should be used to select patients for RFA in the absence of extenuating circumstances; Grade B, low-to-moderate level of certainty</td>
</tr>
<tr>
<td>Orientation of electrodes for RFA</td>
<td>A near-parallel approach should be used; Grade B, low-to-moderate level of certainty. For surgeries involving the articular pillars, a modified (posterior oblique or lateral) with advanced imaging and multiple lesions may be necessary; Grade C, low level of certainty</td>
</tr>
<tr>
<td>Sensory and motor stimulation before RFA</td>
<td>Sensory stimulation should be considered when single lesions are planned and with C2–3 denervation; Grade C, low level of certainty. Motor stimulation may be beneficial for safety and efficacy; Grade B, low-to-moderate level of certainty</td>
</tr>
<tr>
<td>Evidence for larger lesions</td>
<td>There is indirect evidence that larger lesions may improve RFA results; Grade C, low-to-moderate level of certainty. Grade C, low level of certainty for larger lesions to increase duration of pain relief</td>
</tr>
<tr>
<td>Risk mitigation</td>
<td>Aspirate and use real-time fluoroscopy or DSA to prevent vascular uptake; Grade B, moderate level of certainty. Position electrode in the posterior two-thirds of C2–3 facet joint to avoid vascular damage; Grade C, low level of certainty. Discontinue anticoagulants only after a careful risk: benefit assessment; Grade I. Non-particulate steroids can be injected post-RFA when there is a high risk for post-procedure neuritis, and a short course of NSAIDs may reduce post-RFA pain; Grade C, low level of certainty. There is inconsistent evidence supporting the peri-procedural use of gabapentin; Grade I. Discussion with relevant healthcare teams and device manufacturers should be undertaken before performing RFA in a person with an implanted device and their guidance considered; Grade C, low level of certainty. RFA can be performed in individuals with hardware, but may require a modified technique; Grade C, low level of certainty.</td>
</tr>
<tr>
<td>Repeating RFA</td>
<td>RFA can be repeated in patients who obtain meaningful relief lasting ≥3 months when their baseline pain returns; Grade B, moderate level of certainty. For pain that returns in a similar quality and location as the baseline pain, repeat MBBs are not necessary; Grade C, low level of certainty</td>
</tr>
<tr>
<td>Differences between clinical trials and practice</td>
<td>Because of different objectives, differences in patient selection (more rigorous for trials) and minor differences in performance (eg. for quality assurance in studies, to increase study power) may be indicated</td>
</tr>
</tbody>
</table>

AA, atlanto–axial; AO, atlanto–occipital; DSA, digital subtraction angiography; IA, intra-arterial; MBB, medial branch block; NSAID, non-steroidal anti-inflammatory drug; RFA, radiofrequency ablation; TON, third occipital nerve.
**Maximizing access to care**

Similar to the lumbar facet guidelines, we sought to prioritize access while reducing the total number of required procedures. Although medial branch RFA may carry slightly greater risks in the neck than the low back, the risks of properly performed cervical medial branch RFA are still much lower than the risks of alternative therapies such as chronic opioid therapy and surgery.26 408 Although low, prognostic MBB carries risks that are additive when performed multiple times. The correlative of a single positive prognostic MBB in the cervical spine being more likely to be a true positive block than in the lumbar spine is that the false-negative rate is also higher in the cervical spine, putting a greater proportion of patients at risk of not receiving a potentially therapeutic procedure if multiple blocks are required.26 275 Although the committee still advocates a personalized, shared-decision approach to treating suspected cervical facetogenic pain, we felt that performing two blocks in the cervical spine is even more difficult to justify than it is in the low back.

**Guideline limitations**

Unlike standards for which there is little room for deviation, guidelines tend to be more flexible and allow for variations based on physician judgment and unique patient characteristics, providing recommendations in areas of uncertainty. Thus, what may be an ideal treatment course for one patient may be inappropriate for another. For areas of medicine in which there is a lack of high-quality clinical trials to guide treatment and a consensus regarding best practices, the development of inclusive guidelines becomes even more important. Along similar lines, it is important to recognize that patients, payers, regulatory agencies, and even providers may have different needs and goals.

Second, we did not grade the included studies as all grading scales used to rate evidence quality consider only methodological factors while failing to consider the more important aspects of patient selection and procedural technique.35 36 409 This is relevant because not only technical performance but also patient selection (ie, not including patients with secondary gain, comorbid psychological conditions, other sources of pain, or a diffuse pain phenotype) play pivotal roles in clinical trial outcomes. This can be gleaned from the wide variability in enrollment percentages and outcome measures. Whereas technical quality scales have been developed for procedures such as ESIs, existing ones for facet blocks and RFA are dated, methodologically flawed and have not been validated.410–411

Third, by their very nature, guidelines are a byproduct of the opinions and clinical experience of the group, which in our case contained pain physicians. The clinical basis for this was that the questions we considered were mostly technical ones. Administratively, for transparency and equipoise purposes we sought out committee members only from medical societies with stakeholder interests in interventional pain management, who in turn selected representatives with a track record of peer-reviewed publications or grant funding in the area of cervicogenic headache or facet joint pain.

**CONCLUSIONS**

Clinical trials evaluating cervical facet blocks and RFA are characterized by widely disparate outcomes, and there is enormous variation in selecting patients and performing procedures. These multi-society guidelines have been developed to serve as a roadmap to improve outcomes, enhance safety, and minimize unnecessary tests and procedures. Unlike standards, which often
come from an unimpeachable authority and define standards of care, our recommendations are not meant to usurp clinician judgment or personalized medicine. As has been eloquently summarized previously, the practice of evidence-based pain medicine should consider not only the best available research but also take into account clinical experience and expertise, patient values and preferences, and practical concerns. The authors of these guidelines hope researchers, clinicians, and patients will continue to conduct and participate in high-quality research to answer some of the questions addressed in these guidelines in which evidence was not available.

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