Case Report Of Cryoneurolysis For The Treatment Of Refractory Intercostobrachial Neuralgia With Postherpetic Neuralgia

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Abstract: Postherpetic neuralgia is a common and potentially debilitating neuropathic pain condition. Current pharmacologic therapy can be inadequate and intolerable for patients. We present a case of a gentleman with refractory postherpetic neuralgia in the intercostobrachial nerve distribution that was successfully treated with cryoneurolysis/cryoanalgesia therapy.

Keywords: cryoneurolysis, cryoablation, postherpetic neuralgia, intercostobrachial neuralgia, neuropathic pain

Introduction
Postherpetic neuralgia (PHN) is one of the most common painful chronic neuropathic pain conditions and it occurs following an acute herpes zoster infection. Herpes zoster, and PHN generally presents along a unilateral thoracic dermatome, with 20% involving multiple adjacent dermatomes. The neuropathic pain associated with postherpetic neuralgia is persistent, severe and substantially interferes both with quality of life and recreational and daily activities. It even can induce the change from independent living to requiring dependent care. Unfortunately, there is not a single treatment that is completely effective or disease modifying for PHN. Considering that less than half of patients report greater than 50% pain relief with first-line medical therapy, existing medical treatment of postherpetic neuralgia is often refractory to multiple interventions.

We present a case of a 71-year-old gentleman with postherpetic neuralgia initially in the left upper thoracic dermatomes, but with most severe pain along the distribution of the left intercostobrachial nerve (ICBN), which was refractory to initial treatment attempts. The patient ultimately responded to diagnostic local block followed by cryoneurolysis targeted along the ICBN anatomically. Despite multiple failed prior treatment attempts, the patient demonstrated sustained analgesia subsequent to the cryoneurolysis of the ICBN. To the authors’ knowledge, this is the first reported case of refractory PHN that presented as ICBN neuralgia, which was successfully treated with cryoneurolysis/cryoanalgesia therapy.

Case Report
Full informed written consent for the procedure and publication of this case report was obtained from the patient. No further institutional approvals were required. A 71-year-old gentleman with a past medical history of coronary artery disease, hyperlipidemia,
anxiety and depression presented to the pain clinic with the chief complaint of persistent left back, flank and excruciating left arm pain. The pain started, approximately, 5 months prior to initial presentation, when he was diagnosed with acute herpes zoster in the same distribution (upper thoracic, approximately T2-5 dermatomes on left). The pain was severely limited to the patient’s quality of life and ability to participate in activities of daily living (ADL).

The patient further described his pain as constant, severe, burning, pins and needles sensation. Average pain severity was reported as >8/10 (on NRS scale 0–10) and significantly interfered with ADLs by impeding exercise, chores, self-care, socializing and sexual relations. The patient’s history and the physical exam revealed his pain and allodynia were mainly located in the left upper arm along the anatomic ICBN distribution.

Initially, the patient’s herpes zoster infection was treated with a course of valacyclovir, with the resolution of his rash. However, the pain persisted subsequently in a similar distribution. Initial treatments included (sequentially) gabapentin, pregabalin, lidocaine patch, amitriptyline and oxycodone. Despite multiple pharmacologic agents, he had limited-to-no relief and found the side effects including drowsiness and cognitive changes to be prohibitive. Epidural steroid injections were also attempted, but yielded minimal and no sustaining pain relief. The patient had attempted trigger point injections, intercostal nerve blocks (ICB), as well as stellate ganglion blocks. He had significant relief from the ICB; however, the pain seemed mainly focused and isolated in severity along the ICBN and there was no sustained analgesia when the local anesthetic had worn off. Given the reproducibility of the majority of his pain along the specific region of the ICBN, the decision was made to specifically target the ICBN for neural blockade and neurolysis. Following two successful diagnostic blocks (>50% pain reduction for the duration of local anesthetic) of the ICBN with local anesthesia only, it was agreed upon with the patient to pursue cryoneurolysis for greater sustained pain relief.

Informed procedural consent was obtained from the patient. In the supine position, the patient's arm was abducted and externally rotated. The area was prepped with a chlorhexidine solution and a 25-gauge needle was used to deposit local anesthetic solution just distal to the axillary fossa, from the deltoid muscle toward the triceps muscle along the anatomic localization of the ICBN (Figure 1). Ten milliliters of a solution containing 5 mL 0.25% bupivacaine, 4 mL of 1% lidocaine and 10 mg of triamcinolone was injected. After nerve stimulation was obtained, cryoneurolysis was performed using a handheld cryoneurolysis device (Iovera). 3 A 3 probe cryoablation tip probe (with three 27 g needles, approximately 7.6 mm in length) was used along the aforementioned treatment line. The tip of the cryoneurolysis probe was placed along the distribution of the ICBN. The device was used on the dotted line shown in Figure 1, moving circumferentially to cover the entire cross-section of the ICBN. The superficial nature of the field block allowed for the use of the cryoablation needles used in this case as described in Figure 2 to cover the length and depth of the intercostobrachial nerve. Nerve stimulation was not used to identify the exact localization as this nerve is generally located via a field block along anatomical landmarks only.

Multiple cycles of cryotherapy lasting 60 s in the treatment line as described above were administered with three 27-g needles at a depth of approximately 5.1 mm. The tip of the three-prong needle (Figure 2) achieved a cooling temperature of maximum –88 degree Celsius. The patient tolerated the procedure well, without any complications reported. At one-month postprocedure, the patient reported greater than 50% pain relief (NRS <4 on the NRS scale) in the ICBN distribution with significant improvement in self-reported quality of life and activity. The patient did not require any further interventions and reported that the quality of life indicators mentioned earlier were no longer significantly debilitating his lifestyle. He did not need any further follow-up due to improved analgesia.

**Discussion**

Herpes zoster is a viral disease caused by reactivation of latent varicella zoster virus (VZV), characterized by a
vesicular rash distributing in a dermatomal pattern. The incidence of this disease in the United States is estimated to be 3.4 per 1000 persons, increasing significantly after age 50 to 11 per 1000 persons by age 90. Approximately 10–20% of these cases in older individuals will develop PHN. PHN is a neuropathic pain condition defined by severe pain often with allostynia in a dermatomal pattern persisting for months after VZV occurrence and resolution. The pain often precludes functioning, preventing sleep, recreation and can be associated with depression. While the reactivation in acute herpes zoster virus is thought to propagate from the dorsal root ganglion, it is thought that chronic pain with PHN is related to an ongoing inflammatory response, which may also lead to injury of the peripheral nerve in addition to the DRG and higher order somatosensory processing. The authors of this case report hypothesize that this may be the pathophysiologic basis for successful treatment of PHN with cryoablation of the intercostobrachial nerve in the patient presented. This theory has been supported by multiple reported effective peripheral nerve treatments for painful PHN. Currently, the usual treatment is multimodal, often including a combination of different pharmacologic classes of medications in addition to topical agents. These include gabapentinoids, tricyclic antidepressants, and opioids. Unfortunately, treatment of PHN is often inadequate with less than half of patients reporting greater than 50% pain relief with first-line medical therapy. Furthermore, the adverse effects of the mentioned standard treatments are often additive which limit dose escalation and tolerability, especially in older patients.

Cryoneurolysis/cryoablation/cryoanalgesia therapy has been well described for is a well-established treatment for chronic pain states including neuropathic pain. Recently cryoanalgseic techniques have even been applied to acute and perioperative pain states. The concept behind cryoanalgesia/neurolysis involves inducing cold temperature in the proximity of a peripheral nerve by creating ice crystals that will cause reversible axonal degeneration also known as Wallerian degeneration. The mechanism of analgesia in cryoanalgesia is multifold. There may be reduction of conduction, activation of descending inhibitory modulation, decreased excitatory neurotransmission, generalized sodium channel blockage, and likely a combination of all of the above. There are several advantages of cryoablative techniques over other neurolytic techniques including surgical, thermal and chemical ablation (eg, alcohol neurolysis). First, in cryoablation, the epineurium, perineurium and endoneurium are generally intact. This allows for nerve regeneration and reduced risk of neuroma as compared with other neurolytic modalities, which is especially common after surgical neural resection. Neuroma formation can significantly worsen pre-existing pain. Additionally, cryoablation is much less invasive compared with other techniques. Although pulsed radiofrequency (PRF) ablation has recently been described in setting of PHN, there is a larger probe 22 g for PRF vs 27 g in our case, which can be more painful for patient. Also in Makharita’s series there was PRF of the intercostal nerves and not specifically the ICBN (which will be discussed later). The adverse effects and contraindications

Figure 2 A 3-pronged cryoablution device (with three 27-g needles, approximately 7.6 mm in length) was used along the treatment line shown in Figure 1, moving circumferentially to cover the entire cross-section of the ICBN. Multiple cycles of cryotherapy lasting 60 s in the treatment line as described above were administered with three 27-g needles at a depth of approximately 5.1 mm. The tips of the three-prong needle achieved a cooling temperature of maximum −88 degree Celsius.
to cryoneurolysis include bleeding, skin discoloration and infection. As expected, the treatment area will likely become insensate. In our case, these risks were deemed to be exceedingly minimal due to the superficial nature of the cryoablation and probe used. The contraindications to cryoablation, specifically, include patients who have a history of cryoglobulinemia, Raynaud’s syndrome or cold urticaria.5 Our patient had none of these features. Recently a handheld, cryoprobe (loversa, FDA approved) has been developed and used in a variety of acute and chronic pain states.5,12,13,16 The cryoablation probe produces a focused, extremely cold temperature at its tip via the Joule Thompson effect. The basic principle is when high-pressure gas in the device (nitrous oxide) passes to a low-pressure chamber, there is the production of extremely cold temperature.5 This allows for localized cryoablation that is minimally invasive and an in-office procedure.

While cryoablation has been described for PHN as well as intercostal neuralgia and post-thoracotomy pain,11,17,18 to our knowledge this is the first reported case to specifically target the ICBN as a neuralgia, directly related to PHN. The ICBN is often affected/injured postmastectomy and there is literature regarding postmastectomy altered sensitivity with intercostobrachial neuralgia.19 The ICBN is at significant risk for injury postoperatively after procedures involving the axillary region.20 The ICBN is a cutaneous nerve that stems from the second and occasionally third intercostal nerve anatomically.21 One case series by Wisotzky et al demonstrated significant pain relief in patients with ICBN neuralgia, post breast cancer surgery with ICBN blocks.22 However, there were no patients with PHN reported, and the series only looked at 3 patients who were injected with local anesthetic and steroid; no neurolysis was performed.22

Given the severity of PHN and disabling pain to patients with limited options, we believe that cryoablation offers an alternative that is minimally invasive, with less risk compared to other forms of neurolysis. Our case report demonstrates an ICBN neuralgia related to PHN, which to our knowledge has not been reported in the literature and treated with cryoablation. While we acknowledge that additional studies are necessary and the conflicting data on efficacy of cryoablation with intercostal blocks and with post-thoracotomy pain,17,23 there have been positive results for cryoanalgesia for PHN in the dermatologic literature with cutaneous use.18 Additionally, the minimal risk to ICBN cryoablation given the superficial nature, as compared to intercostal blocks, whose risks include pneumothorax and vascular uptake/bleeding, allows for the ICBN cryoanalgesia to be a viable option that should be further explored in patients who present with features of ICBN neuralgia related to PHN.

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Disclosure

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References