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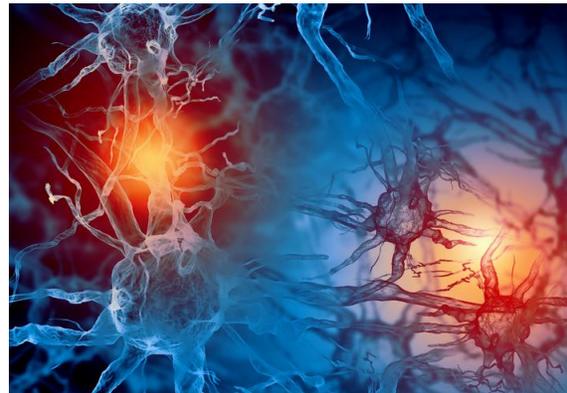


Thank you for entrusting in the compounding services at Madison Medical Compounding Pharmacy to help meet the unique medication needs of your patients. We are excited to share our monthly newsletter with you and look forward to working with you. Please don't hesitate to let us know how we can assist you and your practice.

LeAnn Chambers, Pharm.D. and Matthew Chambers, Pharm.D.

Neuropathic Pain

Neuropathic pain is a common condition that is often resistant to “standard” treatment. Patients are usually dissatisfied with the results of their treatment. There is increasing interest in the optimization and personalization of pharmacotherapy based upon the underlying mechanism(s) of neuropathic pain and the patient's symptoms. The management of chronic neuropathic pain is challenging and is best achieved with the use of a multidisciplinary team.



[Continuum \(Minneapolis, Minn.\). 2017 Apr;23\(2\). Selected Topics in Outpatient Neurology\):512-532](#)

Compounded Topical Analgesics for the Management of Chronic Neuropathic Pain

In recent years, the increase in opioid use has prompted medical professionals to find new and improved solutions to tackle chronic, refractory pain syndromes. Topical analgesics are emerging as a valued solution.

Neuropathic pain (NP) is defined by the International Association of the Study of Pain as “pain arising as a direct consequence of a lesion or disease affecting the somatosensory system”. NP can present alone or in any combination with other types of pain and affects 6–

8% of the population. NP can originate from focal lesions in the peripheral nervous system such as postherpetic neuralgia (PHN) or post-traumatic neuralgia, or from general lesions such as painful diabetic neuropathy, HIV-neuropathy, etc. Additionally, lesions to the CNS and other complex disorders (spinal cord injuries, stroke, complex-regional pain syndrome, etc.) can cause NP. NP can be described as burning, tingling, electric-like or a shooting sensation, with a combination of sensory loss and the paradoxical presentation of hypersensitivity in the painful area. The mechanism of NP includes peripheral sensitization, central sensitization in the dorsal horn of the spinal cord as well as changes in cortical and subcortical regions.

Topical versus Transdermal

Transdermal delivery of medication is accomplished through percutaneous absorption, and systemic therapeutic drug levels can be achieved, comparable to those of oral medication. Often, transdermal medications can be administered distal from the painful area. Transdermal drug delivery has advantages because it avoids first-pass hepatic metabolism which occurs with oral drugs, and the need for frequent painful injections associated with parenteral therapy.

Topical medication targets soft tissues and peripheral nerves underlying the site of application and exerts its action at the site of application by penetrating the skin through passive diffusion. Therefore, topical medication does not produce systemic side effects or drug-drug interactions. One of the main targets for topical analgesics is keratinocytes. This network of skin cells includes several receptors, neurotransmitters, and neuropeptides, which play a significant role in the development of NP.

Topical analgesics may be safer and are easier to use as compared to systemic drugs. Due to the complex nature of NP conditions, topical analgesic therapy should be employed as part of a multidrug approach. Reduction in the renal and hepatic metabolism and various comorbidities managed with multidrug therapy supports the use of topical versus systemic medication for safety concerns. However, there are some limitations to the use of topical agents in the management of NP. Topical analgesics can be used on a limited skin area due to increased risk of toxicity, and for that reason, they cannot be used in conditions with skin integrity disruption or large affected areas.

Prescribing and Practice Notes:

- In order for a topical analgesic to pass through the stratum corneum of the epidermis, it has to possess both hydrophilic and hydrophobic elements.
- The choice of base determines the extent of absorption of a topically applied medication.
- Ketamine acts as an inhibitor on voltage-gated Na⁺ and K⁺ ion channels.
- Topically applied ketamine exerts its peripheral antinociceptive effect by the activation of neuronal nitric oxide synthase.
- Clonidine is lipophilic, which aids in easy skin penetration and it has a half-life of approximately 8 hours, thus requiring three-times daily topical application.
- Phenytoin 10% cream, a nonselective voltage-gated Na channel stabilizer, GABA-a receptor agonist, showed promising results in allodynia reduction.
- Topical gabapentin 10% cream reduced allodynia and hyperalgesia in chronic sciatic constriction nerve injury in rats.
- GABA-b receptors are located in cutaneous layers on nerve endings and keratinocytes.

[Pain Manag. 2017 Nov;7\(6\):537-558.](#)

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