CLINICAL INQUIRIES

Evidence Based Answers from the Family Physicians Inquiries Network

ONLINE EXCLUSIVE

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FAST TRACK

Tricyclic antidepressants, anticonvulsants, and narcotics all reduce postherpetic pain.

What measures relieve postherpetic neuralgia?

Evidence-based answer

Tricyclic antidepressants, gabapentin, and pregabalin effectively reduce pain (strength of recommendation [SOR]: **A**, at least 2 good-quality randomized controlled trials [RCTs] and/or meta-analyses). Opioids have

demonstrated pain relief in 3 RCTs (SOR: **A**, consistent RCTs). Capsaicin and the lidocaine 5% patch relieve pain and decrease allodynia (SOR: **B**, recommendations from meta-analyses and lower-quality RCTs).

Evidence summary

Postherpetic neuralgia (PHN) is defined as pain lasting 1 to 3 months after resolution of acute herpes zoster (shingles) rash. It occurs in approximately 10% to 15% of patients and can cause significant morbidity.

Tricyclic antidepressants provide effective pain relief

Five systematic reviews have concluded that tricyclic antidepressants (TCAs) are effective treatments for PHN.¹⁻⁵ Amitriptyline, the best studied TCA, provides at least moderate pain relief in two-thirds of patients with a pooled number needed to treat (NNT) for TCAs of 2.64 (95% confidence interval [CI], 2.1-3.54)⁵ (TABLE).

Selective serotonin reuptake inhibitors—including fluoxetine, paroxetine, citalopram, and sertraline—have been studied in a variety of neuropathic pain syndromes, but not for treating PHN.¹

Anticonvulsants help, too

Five systematic reviews found gabapentin to be effective, with a range of NNT from

2.8 to 5.3 for as much as 50% pain reduction based on the visual analog score (VAS).²⁻⁶ Pregabalin is also effective, with an NNT of 4.93 (95% CI, 3.34-6.07) for up to 50% pain reduction.^{7,8} Limited data are available concerning the effectiveness of valproate.⁵

A look at the role of narcotics

Four systematic reviews found that controlled-release oxycodone reduced pain by 50%, based on the VAS.²⁻⁵ Another systematic review reported only limited evidence of effectiveness.⁶ In pooled results from systematic reviews, opioids decreased pain by 50% on the VAS (NNT=2.67; 95% CI, 2.10-3.77).⁶

An RCT of 76 patients demonstrated that morphine, with methadone as back-up, both reduced the intensity of pain and relieved pain more than placebo.⁹

Tramadol, a selective opioid agonist, showed moderate effectiveness in a small RCT (N=125), with an NNT of 4.76 (95% CI, 2.61-26.97).^{3,5,6} The mean pain intensity, degree of pain relief, and amount of rescue medication required

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CLASS	DRUG	DOSE	NNT	SIDE EFFECTS
Tricyclic antidepressants⁵	Amitriptyline	Up to 150 mg/d (mean 120 mg/d)	2.64	Sedation, dry mouth, blurred vision, constipation, urinary retention
	Nortriptyline	Up to 150 mg/d (mean 89 mg/d)		
	Desipramine	Up to 150 mg/d (mean 65-73 mg/d)		
Anticonvulsants ^{3,5}	Gabapentin	1800-3600 mg/d	2.8-5.3	Somnolence, dizziness, edema, dry mouth
	Pregabalin	150-600 mg/d	4.93	
Opioids ⁵	Oxycodone	Variable	2.67	Constipation, nausea, vomiting, sedation, dizziness, dependence
	Long-acting morphine/methadone	15-225 mg/d (morphine) (mean 91 mg/d for morphine, 15 mg/d for methadone)	2.67	
	Tramadol	100-400 mg/d (mean 275 mg/d)	4.76	Dependence
Topicals⁵	Capsaicin 0.075% cream	Applied 3-4 times per day	3.26	Burning skin
	Lidocaine 5% extended release patch	Max 3 patches per day	2.0	Mild skin reaction

were all better in the tramadol group than the placebo group.

Evidence for topical therapy is limited

The anesthetic lidocaine patch 5% has shown efficacy in PHN with allodynia based on 3 RCTs of lower quality (short duration, recruitment of patients who had improved on lidocaine previously, no report of baseline levels of pain); the NNT was 2 (95% CI, 1.4-3.3). A systematic review of these 3 RCTs concluded that evidence is insufficient to recommend the lidocaine patch as treatment for PHN. 10

Capsaicin, a topical counterirritant, reduced pain in fewer than 20% of patients in 2 RCTs reported in systematic reviews, with an NNT of 3.26 (95% CI, 2.26-5.85).²⁻⁶ Blinding was limited in these studies because of the stinging associated with treatment.

Recommendations

A 2004 practice parameter of the American Academy of Neurology recommends TCAs (amitriptyline, nortriptyline, desipramine, and maprotiline), gabapentin, pregabalin, opioids, topical lidocaine, and capsaicin to treat PHN (level of evidence: A), but notes that amitriptyline has significant cardiac effects in the elderly compared with nortriptyline and desipramine.³

In 2006, the European Federation of Neurological Societies determined that TCAs, gabapentin, pregabalin, and opioids had established efficacy (level of evidence: A), but recommended opioids as second-line therapy because of potential adverse events with long-term use.⁴ The federation's guidelines designate capsaicin, tramadol, topical lidocaine, and valproate as drugs with lower efficacy or limited strength of evidence (level of evi-

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dence: **B**). Nevertheless, they recommend considering topical lidocaine for elderly patients with allodynia and small areas of pain.⁴

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FAST TRACK

Both topical lidocaine and capsaicin have less evidence of efficacy as treatment for PHN.