Topical Nonsteroidal Anti-Inflammatory Drugs for Chronic Musculoskeletal Pain in Adults

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Objective
To assess the efficacy and safety of different topical NSAIDs compared with oral NSAIDs and with a placebo in the treatment of chronic musculoskeletal pain.

Type of Review
An interventional review of the evidence for the treatment of any chronic pain with a topical NSAID and one of a series of systematic reviews on the use and efficacy of topical analgesics.

Relevance to Nursing
Chronic pain can be debilitating and adversely affect quality of life. Pain management and support for patients living with chronic pain is an essential part of holistic nursing care. Topical NSAIDs are increasingly used for pain relief, with their use as a primary treatment option recommended by the National Collaborating Centre for Chronic Conditions (2008). Osteoarthritis (OA) is the most common type of joint disease and a primary cause of pain and physical disability in older adults. Nurses need to be able to identify medications that are effective in the relief of chronic pain. A wide variety of topical NSAIDs are becoming available, and evidence-based practice must guide choices on efficacy. Therefore, a systematic review was warranted.

Characteristics of the Evidence
This review included 34 randomized double-blind controlled trials involving 7,688 participants. Participants were aged 16 years or older with chronic musculoskeletal pain (chronic not defined), most with a diagnosis of primary OA of the knee or hand confirmed by independent radiologic examination prior to trial commencement. Participants were excluded for pregnancy or lactation, known sensitivity to NSAIDs, coexistent skin disease at site of application, secondary OA, or systemic inflammatory disease.

To be eligible for inclusion, participants had to have been treated with a topical NSAID or comparator for at least two weeks with at least 10 participants per treatment arm. Topical NSAIDs had to be applied at least once per day. Twenty-three of the 34 studies compared topical NSAIDs with a placebo, three studies compared topical NSAIDs with a placebo and oral NSAIDs, three with only an oral NSAID, and two compared topical NSAIDs with a different topical NSAID. One study compared a topical NSAID with a placebo and a non-NSAID topical treatment, and two compared a topical NSAID with a non-NSAID topical treatment. A variety of different topical NSAIDs were used within the studies, including diclofenac, ketoprofen, piroxicam, felbinac, flurbiprofen, piroprofen, nimesulide, flufenamate, indomethacin, and ibuprofen applied as solutions, gels, or patches; 17 studies used diclofenac. Treatment application was defined as application of a set quantity of gel or solution or a patch. Actual dose of the medication was not normally calculated but was defined in terms of number of treatments per day and a specified quantity of agent. Administered oral NSAIDs were all in tablet form. Difficulties in calculation of the topical application dose meant that comparisons between studies were not possible. Outcome analysis of the study data used calculations of relative risk (RR), numbers needed to treat (NNT), or numbers needed to harm.

Methodologic quality of included studies was assessed using a five-point scale that considered randomization, blinding, and study withdrawal and dropouts. A risk of bias tool was used to report on allocation of concealment, sequence generation, blinding, and additional risks such as study size and missing data.

Outcomes of interest were clinical success (defined as a 50% reduction in pain or an equivalent measure), adverse events (local or systemic), and number of withdrawals (whether through lack of efficacy or adverse event). Only patient-reported outcomes were used, although measurement tools for documenting pain were varied.

Summary of Key Evidence
Results were presented according to clinical success, any topical NSAID versus placebo (subdivided according to study duration: 2–3, 4–6, or 8–12 weeks), topical NSAID versus active comparator, and adverse events. The following results were obtained.
- For clinical success comparing topical NSAIDs with placebo, data were insufficient to compare any individual topical NSAID other than diclofenac.
- The NNT for successful treatment with diclofenac in studies of 2–3 weeks' duration (n = 4) was 5 (95% confidence...