

Alternative Agents for Inflammation and Pain

Devils claw

Devils claw is an African plant that was used traditionally for digestive diseases and appetite stimulation as well as for arthritis. Clinical trials have supported its efficacy in the treatment of acute and chronic pain. Preclinical studies suggest that in addition to anti-inflammatory and analgesic effects, the herb has antioxidant and chondroprotective activity. In high quality human clinical trials, devils claw has been as effective as NSAIDs in alleviating pain due to hip and knee arthritis and low back pain. Comparative trials suggested that devils claw is also as safe or safer than NSAID therapy, and no clinical interactions with drugs used in the treatment of arthritic conditions have been reported.

Corydalis

Corydalis has analgesic and possible anti-inflammatory effects, and has been used in China for pain relief for many centuries. Corydalis strengthened the analgesia produced by electroacupuncture (Hu, 1994). The most actively studied constituent, has been effective in reducing nerve pain, painful menstruation, abdominal pain and headache. The mechanism of action appears to involve opioid, GABA-ergic, dopaminergic and cannabinoid receptors. Interestingly, corydalis has been shown to improve healing and clinical symptoms associated with stomach and intestinal ulcers in people.

Boswellia

A 2008 Cochrane review found 7 controlled trials examining the efficacy of boswellia as an anti-inflammatory agent in such varied human conditions as asthma, rheumatoid arthritis, Crohn's disease, colitis, and osteoarthritis. All trials showed clinical benefit. A prospective, open, multicenter clinical trial in dogs with osteoarthritis utilized a standardized boswellia resin extract. Overall, 71% of the dogs that were assessed exhibited "good" or "very good" results at 2 and 6 weeks of treatment. Investigators claimed that after 6 weeks, anywhere from 40% to 70% of the dogs were symptom-free.

Bromelain

Bromelain is a complex of proteolytic enzymes, glycoproteins, and proteinase inhibitors extracted from pineapple. It has been shown (in a formula with another proteolytic enzyme) to have anti-inflammatory activity clinically equivalent to that of the NSAID, diclofenac. Enzyme therapy is thought to have multiple activities that benefit patients with chronic inflammatory disease: they may inhibit release of inflammatory mediators, modulate adhesion molecules, and activate fibrinolysis. Some of its pharmacologic properties may be mediated through nonprotein factors, as well. Although most clinical trials concentrate on mitigation of arthritis pain, bromelain has been used for treatment of other inflammatory diseases as well.

References

Bromelain

- Orsini RA; Plastic Surgery Educational Foundation Technology Assessment Committee. Bromelain. *Plast Reconstr Surg.* 2006 Dec;118(7):1640-4.
- Klein G, Kullich W, Schnitker J, Schwann H. Efficacy and tolerance of an oral enzyme combination in painful osteoarthritis of the hip. A double-blind, randomised study comparing oral enzymes with non-steroidal anti-inflammatory drugs. *Clin Exp Rheumatol.* 2006 Jan-Feb;24(1):25-30.
- Akhtar NM, Naseer R, Farooqi AZ, Aziz W, Nazir M. Oral enzyme combination versus diclofenac in the treatment of osteoarthritis of the knee--a double-blind prospective randomized study. *Clin Rheumatol.* 2004 Oct;23(5):410-5.
- Kerkhoffs GM, Struijs PA, de Wit C, Rahlfs VW, Zwipp H, van Dijk CN. A double blind, randomised, parallel group study on the efficacy and safety of treating acute lateral ankle sprain with oral hydrolytic enzymes. *Br J Sports Med.* 2004 Aug;38(4):431-5.
- Wallace JM. Nutritional and botanical modulation of the inflammatory cascade--eicosanoids, cyclooxygenases, and lipoxygenases--as an adjunct in cancer therapy. *Integr Cancer Ther.* 2002 Mar;1(1):7-37; discussion 37.
- Maurer HR. Bromelain: biochemistry, pharmacology and medical use. *Cell Mol Life Sci.* 2001 Aug;58(9):1234-45.
- Walker AF, Bundy R, Hicks SM, Middleton RW. Bromelain reduces mild acute knee pain and improves well-being in a dose-dependent fashion in an open study of otherwise healthy adults. *Phytomedicine.* 2002 Dec;9(8):681-6.
- Leipner J, Iten F, Saller R. Therapy with proteolytic enzymes in rheumatic disorders. *BioDrugs.* 2001;15(12):779-89.

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- Vishal AA, Mishra A, Raychaudhuri SP. A double blind, randomized, placebo controlled clinical study evaluates the early efficacy of aflapin in subjects with osteoarthritis of knee. *Int J Med Sci.* 2011;8(7):615-22. Epub 2011 Oct 12.
- Abdel-Tawab M, Werz O, Schubert-Zsilavec M. *Boswellia serrata*: an overall assessment of in vitro, preclinical, pharmacokinetic and clinical data. *Clin Pharmacokinet.* 2011 Jun 1;50(6):349-69.
- Krishnaraju AV, Sundararaju D, Vamsikrishna U, Suryachandra R, Machiraju G, Sengupta K, Trimurtulu G. Safety and toxicological evaluation of Aflapin: a novel *Boswellia*-derived anti-inflammatory product. *Toxicol Mech Methods.* 2010 Nov;20(9):556-63. Epub 2010 Sep 29.
- Moussaieff A, Mechoulam R. *Boswellia* resin: from religious ceremonies to medical uses; a review of in-vitro, in-vivo and clinical trials. *J Pharm Pharmacol.* 2009 Oct;61(10):1281-93.
- Ernst E. Frankincense: systematic review. *BMJ.* 2008 Dec 17;337:a2813. doi: 10.1136/bmj.a2813.
- Basch E, Boon H, Davies-Heerema T, Foppo I, Hashmi S, Hasskarl J, Sollars D, Ulbricht C. *Boswellia*: an evidence-based systematic review by the Natural Standard Research Collaboration. *J Herb Pharmacother.* 2004;4(3):63-83.
- Reichling J, Schmokel H, Fitz J, et al. Dietary support with *Boswellia* resin in canine inflammatory joint and spinal disease. *Schweiz Arch Tierheilkd* 2004;146:71-79.

Corydalis

- Chang HM, But PPH. *Pharmacology and Applications of Chinese Materia Medica*, vol 1. Singapore: World Scientific, Inc.; 1986:521.
- Hu J, Xie J, Hu J, Zhang Y, Wang J, Chen R. Effect of some drugs on electroacupuncture analgesia and cytosolic free Ca²⁺ concentration of mice brain [Chinese]. *Zhen Ci Yan Jiu* 1994;19:55-58.
- Huang J, Fang M, Li Y, Ma Y, Cai X. Analgesic effect of *Corydalis yanhusuo* in a rat model of trigeminal neuropathic pain. *J South Med Univ* 2010;30(9):2161
- Kubo M, Matsuda H, Tokuoka K, Ma S, Shiomoto H. Anti-inflammatory activities of methanolic extract and alkaloidal components from *Corydalis tuber*. *Biol Pharm Bull* 1994b;17:262-265.
- Lin DZ, Fang YS. *Modern Study and Application of Materia Medica*. Hong Kong: China Ocean Press; 1990:323-325.
- Yuan CS, Mehendale SR, Wang CZ, Aung HH, Jiang T, Guan X, Shoyama Y. Effects of *Corydalis yanhusuo* and *Angelicae dahuricae* on cold pressor-induced pain in humans: a controlled trial. *J Clin Pharmacol.* 2004 Nov;44(11):1323-7.
- Zhu YP. *Chinese Materia Medica: Chemistry, Pharmacology, and Applications*. Australia: Harwood Academic Publishers; 1998:445-448.

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- Vlachojannis J, Roufogalis BD, Chrubasik S. Systematic review on the safety of *Harpagophytum* preparations for osteoarthritic and low back pain. *Phytother Res.* 2008 Feb;22(2):149-52.
- Chrubasik JE, Roufogalis BD, Chrubasik S. Evidence of effectiveness of herbal antiinflammatory drugs in the treatment of painful osteoarthritis and chronic low back pain. *Phytother Res.* 2007 Jul;21(7):675-83.
- Brien S, Lewith GT, McGregor G. Devil's Claw (*Harpagophytum procumbens*) as a treatment for osteoarthritis: a review of efficacy and safety. *J Altern Complement Med.* 2006 Dec;12(10):981-93.
- Brendler T, Gruenwald J, Ulbricht C, Basch E; Natural Standard Research Collaboration. Devil's Claw (*Harpagophytum procumbens* DC): an evidence-based systematic review by the Natural Standard Research Collaboration. *J Herb Pharmacother.* 2006;6(1):89-126.
- Grant L, McBean DE, Fyfe L, Warnock AM. A review of the biological and potential therapeutic actions of *Harpagophytum procumbens*. *Phytother Res.* 2007 Mar;21(3):199-209.

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