ETHNOBOTANICAL, PHYTOCHEMICAL, PHARMACOLOGICAL AND HOMOEOPATHIC REVIEW OF ARNICA MONTANA LINN

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ABSTRACT
Arnica Montana is a popular traditional remedy used in folk medicine and Homoeopathy to alleviate pain, inflammation and swelling of muscles and joints associated with rheumatoid arthritis and other inflammatory conditions. It is a high-altitude perennial plant that belongs to Asteraceae (Compositae) family and is native to the mountain slopes in Europe, northern Asia, Siberia, Canada and America. This review was conducted to summarize the available scientific information obtained from literature, medical databases, laboratory studies and human clinical studies on Arnica Montana Linn. Arnica Montana Linn, contains essential oil, fatty acids, thymol, pseudoguaianolide, sesquiterpene lactones (helenalin, 11α, 13-dihydrohelenalin and their fatty acid esters) and flavonoid glycosides (spinacetin, hispidulin, patuletin andisorhamnetin). The anti arthritic efficiency of Arnica Montana is attributed to a synergism of phenolic and flavonoid compounds. The analgesic and anti-inflammatory property of Arnica Montana has been attributed at Helenalin. Helenalin also has anti-tumour activity against a variety of chemically induced tumours. It has been pharmacologically proven for arthritis, anti-inflammatory, analgesic, anti-oxidant, wound healing and post operative healing properties. The homeopathic preparations of Arnica Montana have been pharmacologically proven beneficial for arthritis, anti-inflammatory action, anti-haemorrhagic action, wound healing, post operative healing, cellulitis and furunculosis.

KEYWORDS: Arnica Montana, Ethnobotany, Homoeopathy, Pharmacology, Phytochemistry.

INTRODUCTION
Arnica Montana is a popular traditional remedy used in folk medicine and Homoeopathy to alleviate pain, inflammation and swelling of muscles and joints associated with rheumatoid arthritis and other inflammatory conditions.

Arnica Montana belongs to Asteraceae (Compositae), the largest family of flowering plants with its approximately 1,620 genera and more than 23,600 species. Asteraceae has especially diverse distribution in the tropical and subtropical regions of North America, the Andes, eastern Brazil, southern Africa, the Mediterranean region, central Asia, and southwestern China. While majority of Asteraceae species are herbaceous, an important component of the family is also constituted by shrubs and trees. The family contains several species that are important sources of medicinal plants (Arnica, Artemisia, Achillea Millifolium, Calendula, Chamomile, Echinacea, Taraxacum, Solidago), ornamental plants (Aster, Calendula, Chrysanthemum, Dahlia, Sunflower, Zinnia) and Food (Artichokes, Chicory, Lettuce). Insecticides pyrethins are made from Chrysanthemums.

Taxonomy
- Kingdom: Plantae
- Division: Tracheophyta
- Subdivision: Spermatophyta
- Class: Magnoliopsida
- Order: Asterales
- Family: Asteraceae (Compositae)
- Genus: Arnica L.
- Species: Arnica montana L.
- Botanical name: Arnica montana Linn.

Vernacular Names
- English: Celtic Nard, Leopard’s Bane, European Arnica, Mountain Arnica, Mountain Tobacco
- Danish: Guldblomme
- Finnish: Etelänarnikki
- French: Arnique
- German: Arnika, Berg-Wohlverleih
- Swedish: Etelänarnikki, Slättergubbe, Slättergubbe

Habit and Habitat
Arnica is a high-altitude perennial plant native to the mountain slopes in Europe, northern Asia, Siberia, Canada and America.
Botanical Discription
Arnica Montana L. is a perennial herb, 20–50 cm high. Aerial portion consists of a basal rosette of entire obovateleaves up to 17 cm long, five to seven veins, from the centre of which projects an erect, simple, glandular hairy stem up to 0.6 m high. Stem bears two to four pairs of cauline leaves, ovate, elliptic-oblong, lanceolate or oblanceolate, with rounded or rounded-toothed apex and clothed with numerous non-glandular and glandular hairs, up to 16 cm long and 5 cm wide. Peduncles are one to three, bearing alternate bracteoles, extending from the uppermost pair of cauline leaves; glandular–puberulent each terminating in a hemispherical or turbinate capitulum bearing orange-yellow flowers, which are tubular. Tubular floret has a regular, 5 toothed, yellow, tubular corollas about 7 to 8 mm long; 5 epipetalous stamens with syngenesious anthers; pappus and ovary as in ligulate florets. Fruits, black to brown, five-ribbed, with a bristle tuft of hairs. The drug as a whole has a faint, but rather agreeable apple like odour and bitter, acrid taste. Flowers too have a pleasant sweet and aromatic odour and bitter, acrid taste.  

Ethnobotany
Arnica Montana has been extensively used as a remedy to heal wounds, reduce inflammation and soothe muscle aches by Europeans and Native North American Indians for centuries. Arnica is also used as an external application in arthritis, bruises, sprains and swellings; and as a footbath for tender feet. It is one of the natural remedies most often used for rheumatologic conditions. It is also used as a topical counterirritant for treatment of pain and inflammation resulting from minor injuries, bruises, ecchymoses, haematomas and petechiae.

Phytochemical Properties
The major constituents of Arnica Montana Linn. include the essential oil (0.5%), fatty acids (content not specified), thymol (content not specified), pseudoguianolide, sesquiterpene lactones (0.2–0.8%) and flavonoid glycosides (0.2–0.6%). The primary sesquiterpene lactones are helenalin, 11α, 13-dihydrohelenalin and their fatty acid esters. Flavonoids include glycosides and/or glucuronides of spinacetin, hispidulin, patuletin and isorhamnetin, among others.

Experimental Pharmacology
Arthritis
A study conducted in Centre for Interdisciplinary Research in Basic Sciences, Jamia Millia Islamia, New Delhi and Oncology Lab, AIIMS, New Delhi that investigated Arnica Montana flower methanol extract against inflammation and oxidative stress in a collagen-induced arthritis (CIA) rat model concluded that oral administration of Arnica Montana flower methanol extract reduced clinical signs and improve the histological and radiological status of the OA in hind limb joints in rats. Treated rats had lower expression levels of nitric oxide, tumor necrosis factor-α, interleukins (IL-1β, IL-6 and IL-12) and titer of anti-type II collagen antibody compared with untreated CIA rats. Furthermore, by inhibiting these mediators Arnica Montana flower methanol extract also contributed towards the reversal of disturbed antioxidant levels and per-oxidative damage. The anti arthritic efficiency of Arnica Montana is attributed to a synergism of phenolic and flavonoid compounds, the dominant active principles, detected in a methanol extract, which was found efficient on a collagen-induced arthritis (CIA) rat model.  

Analgesic and anti-inflammatory activity
The analgesic and anti-inflammatory property of Arnica Montana has been attributed at Helenalin. Intraperitoneal administration of 2.5–5.0 mg/kg bw of helenalin significantly (P < 0.001) inhibited carrageenan-induced hind paw oedema in rats by 77% after 72 hours. Intraperitoneal administration of 20.0 mg/kg bw of helenalin strongly inhibited acetate acid-induced writhing by 93% in mice but did not have analgesic effects in mice in the hot-plate test. Intraperitoneal administration of 2.5 mg/kg bw of helenalin to rats inhibited arthritis induced by Mycobacterium butyricum by 87%. In vitro, helenalin, 5.0 μmol/l, significantly (P < 0.01) suppressed the activity of prostaglandin synthetase in mouse and rat homogenates, and human polymorphonuclear neutrophils, indicating an anti-inflammatory effect.

Antioxidant activity
Arnica Montana ethanolic extracts were evaluated for their chemical composition, antioxidant activity and protective effect against hydrogen peroxide-induced oxidative stress in a mouse fibroblast-like NCTC cell line at Department of Cellular and Molecular Biology, NIRDs, Romania. This study reported that Arnica Montana extracts are rich in flavonoids and phenolic acids, showed a good antioxidant activity and cytoprotective effect against oxidative damage in fibroblast-like cells. These results provide scientific support for the traditional use of Arnica Montana in treatment of skin wounds, bruises and contusions.

Anti-tumour activity
Helenalin (a primary sesquiterpene lactone in Arnica) is cytotoxic to a wide variety of cancer cell lines in vitro, with a median effective dose (ED50) range of 0.03–1.0 μg/ml (24–27). Intraperitoneal administration of 1.5–33.3 mg/kg by weight of helenalin to mice and rats had anti-tumour activity against a variety of chemically induced tumours.

Wound healing
The molecular aspects of wound healing properties of Arnica Montana were evaluated in a laboratory study at University of Verona and University of Bologna, Italy to elucidate This study evaluated the whole plant extract, in a wide range of dilutions, in THP-1 human cells,
differentiated into mature macrophages and into an alternative IL-4-activated phenotype involved in tissue remodelling and healing. Arnica montana affected the expression of several genes on macrophages differentiated towards the wound healing phenotype. CXC chemokine ligand 1 (CXCL1), coding for an chief chemokine, exhibited the most consistent increase of expression, while also CXC chemokine ligand 2 (CXCL2), Interleukin8 (IL8) and bone morphogenetic protein (BMP2) were slightly up-regulated, suggesting a positive influence of A. montana on neutrophil recruitment and on angiogenesis. This exploratory study concluded that Arnica montana is a promoter of healing, since some of the genes it modifies are key regulators of tissue remodelling, inflammation and chemotaxis.\[19\]

**Clinical Pharmacology**

**Arthritis**

An open multicenter trial of Arnica Montana gel in OA knee was conducted in the Department of Rheumatology, Valens Clinic for Rheumatism, Switzerland to investigate the safety and efficacy of an Arnica Montana fresh plant gel, applied twice daily, in 26 men and 53 women with mild to moderate osteoarthritids (OA) of the knee. After 3 and 6 weeks, significant decreases in median total scores on the Western Ontario and McMaster Universities Osteoarthritis Index (WOMAC) were evident in the intention-to-treat and per-protocol populations (both P < .0001). Scores on the pain, stiffness, and function subscales also showed significant reduction. The overall local adverse-event rate of 7.6% included only one allergic reaction. The patient reported tolerability of the gel as “good” or “fairly good was 87% and and 76% of the patients said they would use it again. Topical application of Arnica montana gel for 6 weeks was a safe, well-tolerated, and effective treatment of mild to moderate OA of the knee.\[19\]

A double-blind study on 204 patients comparing Arnica Montana with Ibuprofen in topical applications for hand OA found no difference in terms of efficacy and side effects were less frequent for Arnica.\[20\]

The equipotency of Arnica with NSAID in the local treatment of hand OA was acknowledged by a Cochrane review.\[21\]

**Muscle pain and muscle damage**

Twenty well-trained males matched by maximal oxygen uptake (VO2 Max) completed a double-blind, randomised placebo-controlled trial to test the efficacy of topical Arnica in reducing pain, indicators of inflammation and muscle damage, and in turn improve performance in well-trained males experiencing delayed onset muscle soreness (DOMS). Topical Arnica was applied to the skin superficial to the quadriceps and gastrocnemius muscles immediately after a downhill running protocol designed to induce DOMS and reapplied every 4 waking hours for the duration of the study. Performance measures (peak torque, countermovement and squat jump), pain assessments (visual analogue scale (VAS) and muscle tenderness) and blood analysis (interleukin-1 beta, interleukin-6, tumour necrosis factor-alpha, C-reactive protein, myoglobin and creatine kinase) were assessed at seven time points over five days (pre-, post-, 4, 24, 48, 72 and 96 hours after the downhill run). Participants in the topical Arnica group reported less pain as assessed through muscle tenderness and VAS 72 hours post-exercise. The application of topical Arnica did not affect any performance assessments or markers of muscle damage or inflammation.\[22\]

**Post-operative healing**

A review of clinical trials done to evaluate prevention and/or treatment of post-procedure ecchymosis or edema with oral arnica and topical arnica reported improvement post-procedure with arnica (4/13) in randomised control trials. It however concluded that there is insufficient data to support use of arnica for post procedure and recommend additional research to determine its efficacy and safety.\[23\]

A review of Effectiveness and Safety of Arnica montana in post-surgical setting, pain and inflammation concluded that it is more effective than placebo and may represent a valid alternative to non-steroidal anti-inflammatory drugs, at least when treating some specific conditions.\[24\]

A systematic review with meta-analysis retrieved for 11 trials with a total of 627 patients was done in the College of Medicine, Korea in the year 2017. This study concluded that the occurrence of periorbital edema and ecchymosis that usually occurs after surgical operation of Rhinoplasty was statistically decreased in the arnica administration groups versus the control group during the first 7 days postoperatively.\[25\]

**Homoeopathic Experimental Pharmacology**

**Anti-inflammatory**

The anti-inflammatory effect of Arnica montana 6CH was evaluated using acute and chronic inflammation models. In the acute, model, carrageenin-induced rat paw oedema, the group treated with Arnica montana 6CH showed 30% inhibition compared to control (P < 0.05). In the chronic model, Nystatin-induced oedema, the group treated 3 days previously with Arnica montana 6cH had reduced inflammation 6 h after the inflammatory agent was applied (P < 0.05). In a model based on histamine-induced increase of vascular permeability, pretreatment with Arnica montana 6cH blocked the action of histamine in increasing vascular permeability.\[26\]

**Cerebral Ischemia**

Homeopathic drugs were studied in the middle cerebral ischemia occlusion (MCAO) model of rat at Jamia Hamdard, New Delhi. The content of antioxidants, thiobarbituric acid reactive substances (TBARS) was elevated significantly whereas the level of reduced
glutathione (GSH) was depleted significantly in the MCAO of rats as compared to the sham group. The rats pretreated with C200 potency once daily for 5 days orally (1 drop or 21 μl) and post treated after 24 hr of MCAO with C30 potency 3 times a day for 5 days orally (1 drop or 21 μl) of Croton, Phosphorus, Arnica and Crocus. The pre and post treated potencies of C200 and C30 respectively has protected the activities of these enzymes significantly when compared with the animals of MCAO group. The study has shown that the Homeopathic drugs have protected most of the studied parameters significantly but further studies are required to comments on the mechanism and reproducibility of Homeopathic drugs.[27]

Wound healing
An animal model study concluded that Homeopathic medicines- Arnica and Staphysagria have a significant role in wound healing and cicatrization process.[38]

Homoeopathic Clinical Pharmacology
Arthritis
A multi-centric observational study carried out by Central Council for Research was conducted between the years 1984-2005 to identify a group of most effective homeopathic medicines in osteoarthritis. A total of 1049 out of 1323 osteoarthritis patients enrolled were followed up regularly. Homoeopathic simillimum was prescribed after detailed case taking. Arnica (n=10) was among the most effective medicines. Others included were Arsenicum album (n=10), Bryonia alba (n=86), Calcarea carbonica (n=92), Calcarea fluoricum (n=15), Causticum (n=8), Graphites (n=15), Lycopodium clavatum (n=168), Medorrhinum (n=21), Natrum mur. (n=11), Pulsatilla (n=26), Rhus toxicodendron (n=279) and Sulphur (n=83). [26]

Anti-haemorrhagic activity
The preliminary results of a randomised, double-blind, placebo-controlled study concluded that stem tincture of Arnica montana was found to possess anti-haemorrhagic activity in women of age group 20–35 years, which reduced postpartum blood loss.[30]

Cellulitis
A case reported of a 55-year-old female patient with pain in the right hand and numbness in the fourth finger after cellulitis in the palmar area which was treated with antibiotics, but pain symptoms remained at 7 points on a 0 to 10-point visual analog scale (VAS) for pain despite intake of oral ibuprofen. Ten Arnica patches (3X dilution according to the Homeopathic Pharmacopoeia of the United States) were dispensed to the patient and applied at night. A substantial decrease in pain symptoms (VAS = 1) and a marked decrease in numbness was reported after 3 days. This case report concluded that Arnica patch on the hand provided a marked reduction of pain and recovery of functionality of the hand and may thus represent an alternative NSAIDS.[31]

Furunculosis
A prospective, multicentre observational study was conducted on furunculosis with nine predefined trial medicines. 539 patients were enrolled of which 397 patients completed regular follow up. Predefined trial medicines were Arnica montana, Arsenicum album, Antimonialum crudum, Berberis, Calcarea carbonica, Calcarea phosphoric, Echinacea, Hepar sulphuris, Sulphur and these were prescribed as per individualization on holistic principles of homoeopathy. A furunculosis symptom score (FSS) was designed to assess the severity of illness. There was significant difference in the mean total FSS between baseline and end of the treatment favouring homeopathic care. The results were found to be statistically significant (p < 0.05) with all nine predefined trial medicines.[32]

Post-operative healing
To assess whether Arnica administration affects recovery from hand surgery a double-blind, randomized comparison of oral and topical Arnica administration versus placebo was done for 37 patients undergoing bilateral endoscopic carpal-tunnel release between June 1998 and January 2000 in the Department of Plastic Surgery of Queen Victoria Hospital in West Sussex, England. Homeopathic Arnica tablets and herbal Arnica ointment compared to placebos. No difference in grip strength or wrist circumference was found between the 2 groups. However, there was a significant reduction in pain experienced after 2 weeks in the Arnica-treated group (P<.03).[33]

Twenty-nine patients undergoing face-lift surgery at a tertiary care center were treated perioperatively with either homeopathic Arnica Montana or placebo in a double-blind fashion. Postoperative photographs were analyzed using a novel computer model for color changes, and subjective assessments of postoperative ecchymosis were obtained.Patients taking perioperative homeopathic A. montana exhibited less ecchymosis, and that difference was statistically significant (P<.05) on 2 of the 4 postoperative data points evaluated.[34]

A prospective, placebo-controlled, double-blind study in which patients were randomly assigned to the administration of homeopathic Arnica Montana or placebo concurrent with unilateral upper eyelid blepharoplasty followed by contralateral treatment at least 1 month later reported no statistically significant difference in area of ecchymosis or rank order of ecchymosis severity for days 3 and 7 after treatment.[35]

A prospective randomized double-blind study was conducted to compare the effect of oral perioperative Arnica Montana homoeopathic preparation (AMHP) with placebo among patients scheduled for rhinoplasty surgery with nasal bone osteotomy by a single surgeon. Ecchymosis was measured in digital "three-quarter"-view photographs at 3 postoperative time points. Each bruise was outlined with Adobe Photoshop and change in

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intensity was calculated. Compared with 13 subjects receiving placebo, 9 taking AMHP had a statistically significant difference in ecchymosis on day 7 (P = 0.097). Color change showed a statistically significant difference on day 9/10 (P = 0.074). The study concluded that Arnica montana accelerated postoperative healing, with quicker resolution of the extent and the intensity of ecchymosis.\textsuperscript{36}

**Homoeopathic Uses**

In Homoeopathy, the whole plant of Arnica Montana is used for medicinal purposes. The Mother Tincture is prepared according to old method, class III of mother tincture preparation and has drug strength 1/10 (Arnica Montana in coarse powder 100 g, purified water 400 ml and strong alcohol 635 ml to make one thousand millilitres of the Mother Tincture). 2Xpotency contains one part tincture, three parts purified water and six parts strong alcohol. 3X and higher potencies are prepared with dispensing alcohol.\textsuperscript{6}

The standard Homoeopathic Materia Medicae describe that Arnica Montana is suited to cases when any injury, however remote, seems to have caused the present trouble. It is useful after traumatic injuries with blunt objects, a muscular tonic for complaints from overuse of any organ, muscular sprains and compound fractures. It is useful in stupor from concussion with involuntary faeces and urine. Useful in conjunctival or retinal haemorrhage form injuries or cough. Venous stasis; patient complains of ecchymosed skin as from bruises. There is tendency to haemorrhage, low-grade fever, tissue degeneration, septic conditions and abscesses that do not mature. Arnica controls haemorrhage, prevents suppuration and aids absorption. Arnica is useful in traumatism of grief, remorse or sudden financial loss. The patient is nervous, cannot bear pain; whole body over-sensitive. In gout and rheumatism patient has fear of being touched or struck by persons coming near him. Cannot walk erect on account of a bruised sort of feeling in the pelvic region. Sore, lame, bruised feeling all through the body, as if beaten. Uneasiness in the painful parts; has to change position often, every place seems too hard. There is tendency to small, painful boils, one after another that are extremely sore. Constipation with loaded rectum, faeces will not come away; ribbon like stools from enlarged prostrate or retroverted uterus. Soreness of parts and/ or retention or incontinence of urine after labour; prevents post-partum haemorrhage and puerperal complications. The patient is worse from least touch; motion; rest; wine; damp cold; and better by lying down, or with head low.\textsuperscript{37,38,39}

**CONCLUSION**

Arnica Montana Linn. is an important medicinal plant in used in Homoeopathy. In recent years, Arnica Montana has received much attention and its medicinal virtues have been evaluated on modern scientific lines such as phytochemical analysis, pharmacological screening and clinical trials. Its anti arthritic efficiency is attributed to a synergism of phenolic and flavonoid compounds. The analgesic and anti-inflammatory property is attributed at Helenalin. Helenalin also has anti-tumour activity against a variety of chemically induced tumours. The important pharmacological properties of this medicinal plant include arthritis, anti-inflammatory, analgesic, anti-oxidant, wound healing and post operative healing. The homoeopathic preparations of Arnica Montana have been pharmacologically proven beneficial for arthritis, anti-inflammatory action, anti-haemorrhagic action, wound healing, post operative healing, cellulitis and furunculosis.

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