

Evexia



HEMP OIL



CASPEROME®



melofeed®



Evexia is a complementary feed for dogs and cats

✓ CONTAINS HEMP OIL

✓ WITH NATURAL HEMP DERIVATIVES (TERPENES, CANNABINIDS, ETC.)

✓ TRY IT FOR 30 DAYS!

Candioli
PHARMA

Evexia

CONTAINS HEMP OIL
PRODUCT FROM PROCESSING OF HEMP PLANT
FREE OF THC (tetrahydrocannabinol, responsible for the psychotropic effects of cannabis)

THE ENTOURAGE EFFECT OF CANNABIS SATIVA

For some neurologic conditions, such as chronic pain, phytocomplex from Cannabis sativa, i.e. terpenes, flavanoids and phytocannabinoids, are thought to be more effective than a single one molecule, thanks to what is defined as «entourage effect». This is a synergistic effect of the different substances contained in the whole plant^{1,2,3}.

The entourage effect is the term used to describe enhancement of efficacy, with related improvement in overall therapeutic effectiveness, derived from combining phytocannabinoids and other plant-derived molecules^{1,2,3,15}.

Phytocannabinoids and terpenes from Cannabis sativa interact with the endocannabinoid system, miming the effects of endogenous cannabinoids^{1,3,11,16,17}, thus playing a role in homeostatic endogenous modulation of nociception¹¹.

Evolution and progression of osteoarthritis are due to the combination of the following factors^{9,22,23}:

- degeneration
- inflammation
- oxidative damage

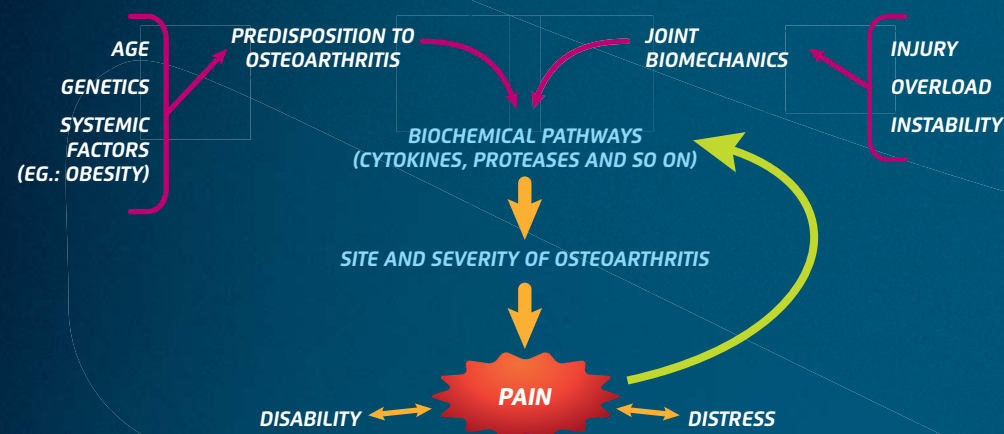


Fig. 1 Current model of osteoarthritis (OA) (From Dieppe PA, Lohmander LS. Lancet. 2005; 365:965-73) mod from Beale BS 2011

CHRONIC ORTHOPEDIC PAIN IS MULTIFACTORIAL

Chronic orthopedic pain²³ is, at least in part, the consequence of **nociceptive stimulation** of^{8,9,10,14,23}:

- joint capsule
- synovial membrane
- ligaments and subchondral bone

activated by inflammatory cytokines (PG, LT, IL, TNF- α) and free radicals (NO)^{5,8,9,20,21}

Sensitization of pain pathways during OA involves changes in joint nociceptors and nociceptive variations at the **spinal cord**, **encephalon**, and **thalamocortical levels**¹⁹.

Therefore, pain in OA has a different origin^{9,16}:

- inflammatory
- nociceptive
- neuropathic

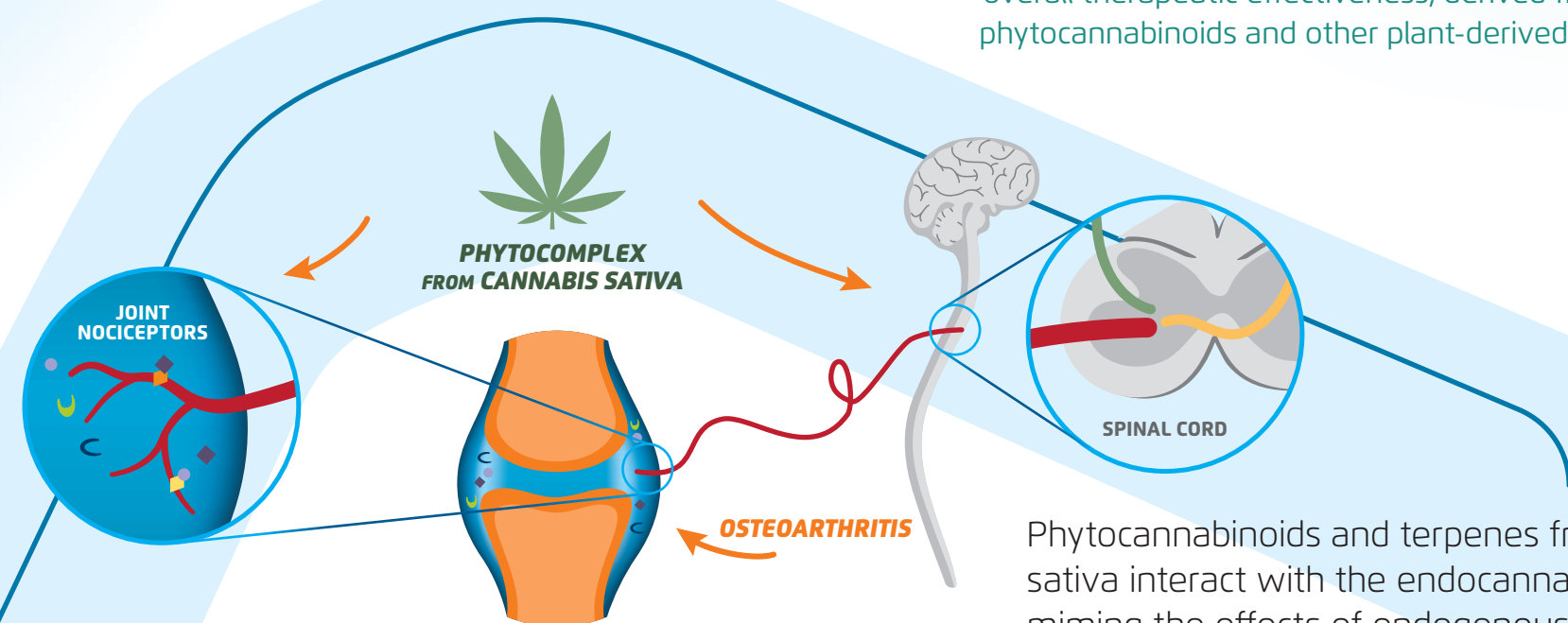


Fig. 2 Elements of the endocannabinoid system in the joints. mod da O'Brien, 2018

CONTAINS CASPEROME

The control of joints inflammation and oxidative damage is important in the natural modulation of pain caused by OA^{5,8,10,20,21}



EXTRACT OF
BOSWELLIA
PHYTOSOMIZED

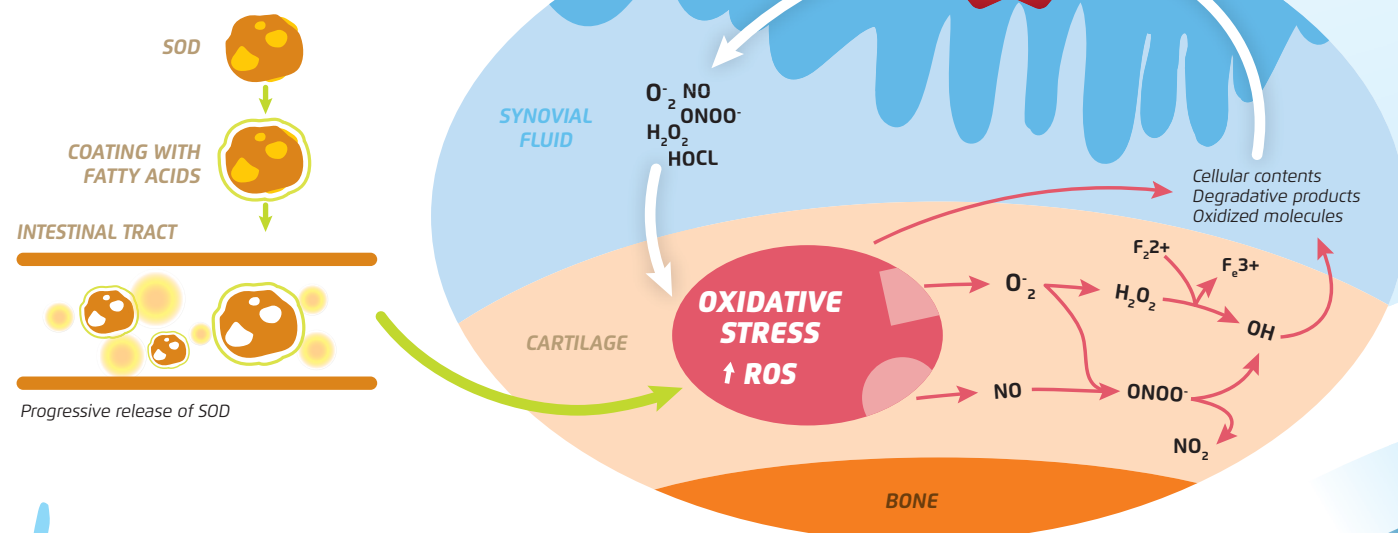
4 times more bioavailable^{7,19} thanks to the patented lecithin delivery form (phytosome) technology indena



The Boswellia extract:

- Prevents inflammatory mediators synthesis (IL-1 β , TNF- α , PGE₂)^{18,19}
- Reduces oxidative damage^{18,19}
- Enhances the activity of antioxidant defense systems (GSH, SOD, Catalase)^{18,19}

Fig. 3 The implication of ROS in cartilage degradation and related synovium inflammatory reaction. H₂O₂: hydrogen peroxide; ONOO⁻: peroxynitrite; OH⁻: hydroxyl radical; O₂⁻: superoxide anion.
from Henrotin 2003



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melofeed®

Natural protected and bioavailable SOD

The **SOD family enzymes** play a key role in controlling the level of free radicals in the cartilage^{9,21}.

In OA animal cartilage (Guinea pigs), SOD2 (Mn-SOD) levels are **significantly reduced** even at an early stage²¹, before the onset of cartilage erosions.

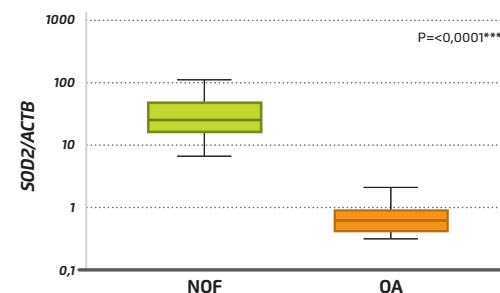
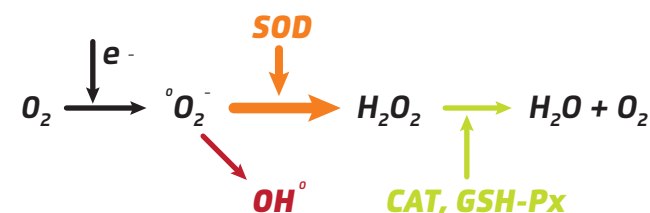


Fig. 4 Comparison of SOD gene expression between femoral neck cartilage (NOF9) and patients with osteoarthritis (OA), in relation to the ACTB gene. Values measured with RT-PCR on T2 +12 subjects: the lines inside the boxes represent the medians, the boxes show the 25th and 75th percentage, the lines outside the boxes show the maximum and minimum values.
from Scott JL 2011 mod.

Evexia



TESTED FOR ITS EFFECTIVENESS IN REDUCING PAIN IN OA DOGS¹³

✓ TRIED FOR
30 DAYS!

Research Article
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EFFECT ON PAIN AND MOBILITY OF A NEW DIET SUPPLEMENT
IN DOGS WITH OSTEOARTHRITIS: A PILOT STUDY

In this study, we have evaluated the efficacy of a new diet supplement in reducing chronic pain and improving mobility in a group of 10 dogs with Osteoarthritis (OA).

OA is a common debilitating condition affecting humans and animals. Tablets containing a preparation of natural ingredients (Cannabidiol (CBD)-rich fraction Boswellia serrata Roxb. in a Phytosome® delivery form and Cucumis melo L. extract) were administered for 30 days. Veterinary evaluations were performed and owners filled questionnaires on chronic pain (Helsinki chronic pain index-HCPI) three times during the study.

The product was well tolerated and owners reported a **good palatability** and **ease of administration**. In terms of effectiveness, results of a Generalized Linear Mixed Model (GLMM) on HCPI highlighted a **significant reduction** of pain scores at the end of the study. Based on our observations, our new dietary supplement has **beneficial effects in dogs with OA** after the treatment.

Tab. 1 The mode scores registered at T1 and T2 by owners on the ease of administration, palatability, and effectiveness and by the veterinarian on the effectiveness of the administered tablets.

	Owner		Veterinarian	
	T1	T2	T1	T2
Ease of administration	0	0		
Palatability	3	3		
Effectiveness	0	3	1	3

CONCLUSION

Our study shows the beneficial effects of a new dietary supplement on the pain management and mobility in dogs with OA

The **combination of natural ingredients** seems to work well in this formulation, being the overall effect of the treatment very satisfying. In addition to the **tolerability** and **safety** of our product, its **good palatability** and its **ease of administration** have to be considered relevant points for the success of a long lasting treatment in animals with a chronic pain like OA.

A new study is currently underway for the purpose of reinforcing these preliminary results.

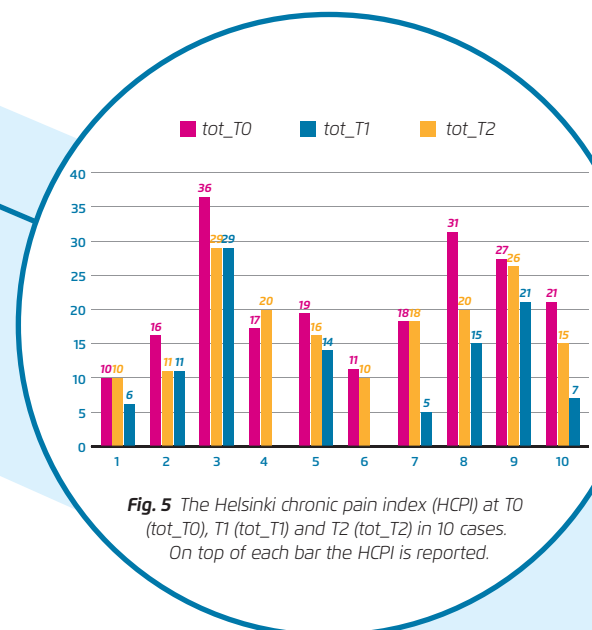


Fig. 5 The Helsinki chronic pain index (HCPI) at T0 (tot_T0), T1 (tot_T1) and T2 (tot_T2) in 10 cases. On top of each bar the HCPI is reported.

Evexia

Contains Hemp Oil



COMPOSITION

Yeasts [brewer's yeast], Hemp oil 3.0%, Magnesium stearate, Mono and diglycerides of fatty acids esterified with organic acids (glyceryl dibehenate), Products from the processing of plants (Melon) (*MeloFeed), Product from the processing of plant (Hemp)

ADDITIVES PER KG

Emulsifiers, stabilising agents, thickeners and gelling agents: Microcrystalline cellulose E460 mg 412.930 - Lecithin 1c322 mg 42.500 Organoleptic additives: Boswellia serrata Roxb. ex Colebr.: Oliban extract (*Casperome®) mg 49.375 Anti-caking agents: Colloidal silica E551b mg 21.875 Amino acids, their salts, and analogues: L-Arginine mg 12,500 Preservatives: Butylhydroxytoluene (BHT) E321 mg 100

INSTRUCTIONS FOR PROPER USE

Evexia is a complementary feed for dogs and cats. The hemp oil contained in Evexia promotes the physiological maintenance of sensory function. Give Evexia to your pet directly in the mouth or mix it with the regular feed serving for a minimum period of 20 days according to the following daily amounts:

Tablets per day			
0-7,5 kg b.w.	15-22,5 kg b.w.	30-40 kg b.w.	1 TAB 15 KG
7,5-15 kg b.w.	22,5-30 kg b.w.	40-60 kg b.w.	

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