### ARTHROBEN™

Medical Food for the dietary management of osteoarthritis and musculoskeletal inflammation related to athletic or other physical activity

### **INGREDIENTS**

Arthroben <sup>™</sup>	Amount per serving
Serving size	11 g
Flavocoxid	250 mg
Fortigel™	5 g
Verisol™	2.5 g

**ARTHROBEN**™ is available in a delicious tasting apple flavored powder

### **RECOMMENDED USE**

Mix 11 grams (approx. 1 1/2 tablespoons) in 8 ounces of water per day, or as directed by a physician

### REFERENCES:

1. Levy R, et al. Flavocoxid is as effective as naproxen for managing the signs and symptoms of osteoarthritis of the knee in humans: a short-term randomized, double-blind pilot study. Nutr Res. 2009 May; 29(5):298-304. 2. Burnett BP, et al. Safety Evaluation of a Combination, Defined Extract of Scutellaria baicalensis and Acacia catechu. J Food Biochem. 2007;31:797-825. 3. Burnett BP, et al. A medicinal extract of Scutellaria baicalensis and Acacia catechu acts as a dual inhibitor of cyclooxygenase and 5-lipoxygenase to reduce inflammation. J Med Food. 2007 Sep;10(3):442-51. 4. Adam M, What effects do gelatine preparations have? Therapy of osteoarthritis (in German), Therapiewoche 1991, 41, 2456-2461. 5. Clark, Sebastianelli et al., 24-Week study on the use of collagen hydrolysate as a dietary supplement in athletes with activity-related joint pain, Current Medical Research and Opinion, Vol. 24, No. 5, 2008, 1485 - 1496.6. Edward J. Frech and Mae F. Go, "Treatment and chemoprevention of NSAID-associated gastrointestinal complications", Therapeutics and Clinical Risk Management, 2009, pp. 65-73 7. Singh G. Recent considerations in nonsteroidal anti-inflammatory drug gastropathy. Am. J. Med. 1998 105(1B):31S-38S 8. Vonkeman HE and van de Laar MAFJ. Nonsteroidal anti-inflammatory drugs: adverse effects and their prevention. Semin Arthritis Rheum. 2010 39(4):294–312. 9. Back M, et al. Cyclooxygenase2 inhibitors and cardiovascular risk in a nationwide cohort study after the withdrawal of rofecoxib. Eur Heart J. 2011;21. [Epub ahead of print] 10. Moodley I. Review of the cardiovascular safety of COXIBs compared to NSAIDS. Cardiovasc J Afr. 2008 MarApr 19(2):1027. 11. Belknap SM. NSAIDs were associated with increased risk for mortality, regardless of time since first MI. Ann Intern Med. 2013 Jan 15;158(2): JC10. 12. Roubille C, et al. Cardiovascular adverse effects of anti-inflammatory drugs. Antiinflamm Antiallergy Agents Med Chem. 2012 Dec 31. 13. Arch Intern Med. 2000;160(6):777-784. doi:10.1001/archinte.160.6.777. 14. Singh Gurkirpal, MD, "Recent Considerations in Nonsteroidal Anti-Inflammatory Drug Gastropathy", The American Journal of Medicine, July 27, 1998, p. 31S 15. Wolfe M. MD, Lichtenstein D. MD, and Singh Gurkirpal, MD, "Gastrointestinal Toxicity of Nonsteroidal Anti-inflammatory Drugs", The New England Journal of Medicine, June 17, 1999, Vol. 340, No. 24, pp. 1888-1889. 16. Larson AM, et al. Acetaminophen-induced acute liver failure: results of a United States multicenter, prospective study. Hepatology. 2005 42(6):1364-72. 17. Bessems JG and Vermeulen NP. Paracetamol (acetaminophen)-induced toxicity: molecular and biochemical mechanisms, analogues and protective approaches. Crit Rev Toxicol. 2001 31(1):55-138. 18. Patrício JP, et al. Relative Cardiovascular and Gastrointestinal Safety of Non-selective Non-steroidal Anti-inflammatory Drugs Versus Cyclo-oxygenase-2 Inhibitors: Implications for Clinical Practice. Clin Drug Investig. 2013 Jan 22



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# ARTHROBEN™

MEDICAL FOOD

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### **NSAIDs**

Selective or unbalanced inhibition of an individual COX or LOX pathway is believed to result in most of the common side effects associated with chronic NSAID use.

### **NSAID Side effects:**

- GI complaints, i.e., nausea, heartburn, dyspepsia, abdominal pain
- -May occur in up to 40% patients with chronic NSAID use<sup>6</sup>
- Bleeding ulcer <sup>7,8</sup>
- Kidney damage that may persist even after drug withdrawal in some cases<sup>9,10</sup>
- Cardiovascular events (MI, stroke, etc.)<sup>11,12</sup>
- Congestive heart failure
- "NSAIDs were responsible for approximately 19% of hospital admissions with CHF" 13
- —"The burden of illness resulting from NSAID-related CHF may exceed that resulting from gastrointestinal tract damage." <sup>13</sup>
- Hospitalization and death:
- —Approximately 107,000 patients are hospitalized annually for NSAID-related GI complication<sup>13</sup>
- —At least 16,500 NSAID-related deaths occur each year among arthritis patients alone. 14,15 (Statistics do not include deaths ascribed to the use of over-the-counter NSAIDS)
- -Many physicians and most patients are unaware of the magnitude of the problem. 15

### **ACETAMINOPHEN SIDE EFFECTS:**

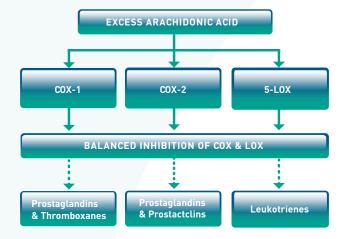
- The leading cause of acute liver failure 16
- Kidney toxicity<sup>17</sup>

"Patients with high cardiovascular or gastrointestinal risk should avoid using NSAIDs." 17

## ARTHROBEN™ produces relatively balanced inhibition of COX-1, COX-2 and 5-LOX pathways

ARTHROBEN™ balances COX-1 & COX-2 metabolism by inhibiting both enzymes relatively equally, in addition to inhibiting 5-LOX, which helps to minimize systemic side effects while effectively managing the metabolic imbalances of OA.

The unique combination of actives in ARTHROBEN™, inhibit inflammation and oxidation, increase functional mobility, and stimulate connective tissue repair.



### ARTHROBEN™

ARTHROBEN's 4 pronged approach to the dietary management of osteoarthritis:

- 1. Reduces inflammation—balanced COX and LOX inhibition
- 2. Potent antioxidant protection to reduce joint degeneration
- 3. Reduces stiffness; increases joint mobility and function
- 4. Stimulates connective tissue repair

### **KEY ACTIVES IN ARTHROBEN™**

### Flavocoxid

Flavocoxid is a proprietary blend of the flavonoids baicalin and catechin that was shown as effective as naproxin in managing knee osteoarthritis, and produced improvements in 87% of patients.<sup>1</sup>

### **Dual Action of Flavocoxid:**

- 1. Balanced inhibition of COX-1, COX-2 and 5-LOX pathways. This balanced down-regulation, though weaker than traditional NSAIDs and selective COX-2 inhibitor drugs, is not associated with the side effects commonly seen with selective COX inhibitors.
- 2. By acting as a strong antioxidant to limit free radicals and oxidative damage to cartilage and other connective tissue of the joints.

### Fortigel® & Verisol®

Fortigel® & Verisol® are standardized mixtures of collagen peptides derived from a patented process of hydrolysis of porcine or bovine type I collagen. These ingredients boost anabolic processes in connective tissues and provide building blocks for all collagen in the body.

### **Fortigel**®

- 16 human clinical trials (approximately 2800) subjects demonstrating a positive effect on joint health
- Reduced need of analgesics<sup>4</sup>
- Increased joint mobility<sup>5</sup>
- Improved radiographic markers of cartilage health

### **Verisol®**

Verisol® (like Fortigel®) stimulates anabolic processes in connective tissue

- Tested in two recent human clinical studies (pre-publication) for its effects on type I collagen in skin
- Improved skin elasticity and collagen type I content

ARTHROBEN™: The freedom to move. The power to heal.

## CLINICALLY SHOWN TO BE AS EFFECTIVE AS NAPROXEN IN MANAGING KNEE OSTEOARTHRITIS<sup>1</sup>

