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## Asthashine capsules: an excellent choice to boost muscle resilience.

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### ABSTRACT

Astaxanthin is a strong antioxidant which by reducing oxidative stress will support muscle function. Oxidative stress has harmful effects on muscle health. Clinical studies have shown that astaxanthin increases muscle endurance, lowers lactic acid and prevents muscle atrophy in aging. The effects of astaxanthin on muscle are explained by its ability to protect membranes from oxidation and thereby enhance mitochondrial function and reduces inflammation and muscle damage. This article reviews the current available scientific literature regarding the effect of astaxanthin from the algae *Haematococcus pluvialis* in Astashine capsules in muscle resilience.

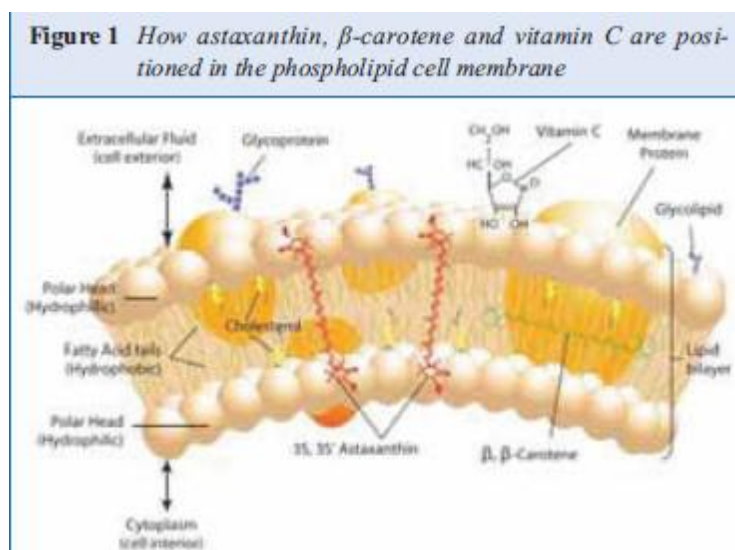
**Keywords:** Astashine capsules, Muscle resilience, Oxidative stress.

### INTRODUCTION

An imbalance between reactive free radicals and antioxidant defense leads to an oxidative stress state. Physical activity generates more free radicals due to greater stress on muscle fibers and an increased metabolism [1].

Up to 5% of the total oxygen consumed by mitochondria will end up as free radicals. Intensive training leads to higher oxygen consumption and therefore can increase the generation of free radicals up to 100 times. Although endogenous

antioxidants are directly increased by strenuous exercise, their protection against oxidative stress is not adequate during later recovery [2]. Furthermore, reduced levels of antioxidants have been suggested to be a factor in the process of muscle atrophy and a cause for why muscle mass is gradually lost with age, which weakens our body and makes us more prone to falls and injuries and to age-related illnesses exacerbated by inactivity [3].



**Molecular structure of Astaxanthine**

**Figure 1** How astaxanthin, beta-carotene and vitamin C are positioned in the phospholipid cell membrane

The unique molecular structure of astaxanthin enables it to be positioned both inside and outside the cell membrane affording better protection to the membrane as compared to I-carotene or vitamin C. These can only be positioned, respectively, inside or outside the lipid bilayer.

### Composition of Astashine capsules

Astaxanthin - 2mg

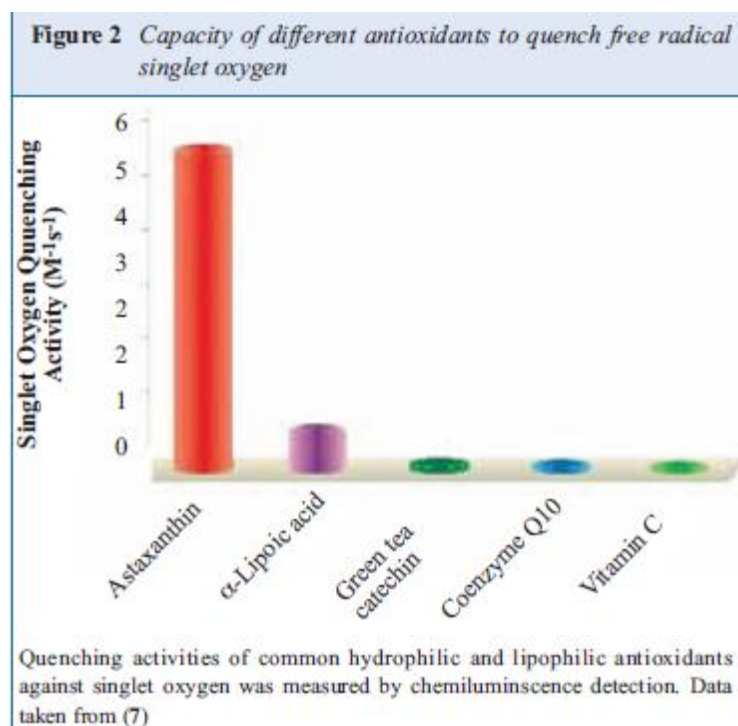
(Naturally derived from Haematococcus pulvis extract, which is microencapsulated)

### Oxidative stress and its effect on muscles

Oxidative stress damages proteins, lipids, DNA and alters the function of muscle cells [1]. Increased oxidative stress triggers inflammation by activating pro-inflammatory cytokines which leads to muscle pain, stiffness and injuries. In addition, oxidative stress damages mitochondrial membranes, which reduces their capacity to

generate energy. Mitochondria provide as much as 95% of the body's energy, primarily by burning carbohydrates and fats. As a consequence of reduced mitochondrial function, the muscles will be supplied with less energy causing muscle fatigue; this is a major factor in muscle atrophy in aging [1,3]. Moreover, oxidation of the red blood cell (RBC) membranes, followed by poorer motility and

A reduced blood flow, might lower their ability to transport oxygen out to muscles. It has been demonstrated that physical activity increases oxidation of the RBC membranes [4,5]. Oxidation of RBCs and impaired mitochondrial function might finally result in lowered aerobic capacity, which increases lactic acid and exhaustion. In addition, increased oxidative stress might alter muscle contraction and damage enzymes in the aerobic and anaerobic pathways resulting in declined muscle power and fatigue [1].



**Figure 2** Capacity of different antioxidants to quench free radical singlet oxygen

Quenching activities of common hydrophilic and lipophilic antioxidants against singlet oxygen was measured by chemiluminescence detection.

#### ***Astaxanthin and its benefits on muscle function and endurance***

Several clinical studies have reported that astaxanthin from the alga *Haematococcus pluvialis* has benefits on muscle function and endurance. Astaxanthin is related chemically to other lipid antioxidants of the carotenoid family such as lutein, zeaxanthin, beta-carotene and lycopene. It is found typically in seafood such as salmon and crabs.

Astaxanthin is fat soluble and, as can be seen in *Figure 1*, its unique molecular structure enables it to stretch through the cell membrane. [6]. Nishida *et al* [7] found that astaxanthin had the greatest capacity to quench singlet oxygen as compared with several other antioxidants (*Fig 2*). In other *in vitro* systems, astaxanthin has shown antioxidant capacity up to 500 times higher than vitamin E and 10 times higher than  $\beta$ -carotene [8]. Unlike many

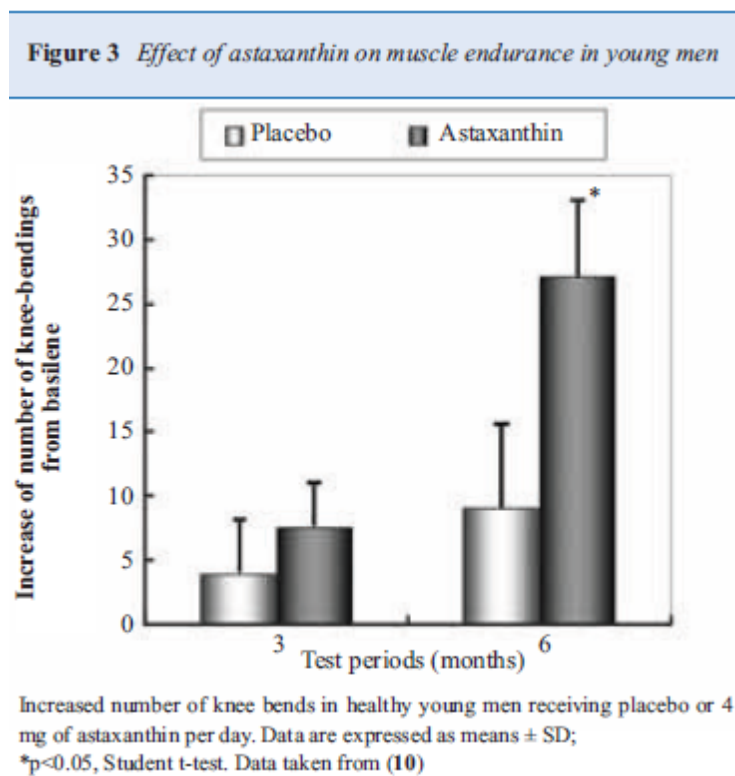
other antioxidants, astaxanthin has been classified as a pure antioxidant and does not show any pro-oxidative effects [9].

Other antioxidants may show pro-oxidative effects under certain conditions, which increase oxidative stress and cause damage on cells

#### **CLINICAL STUDY REPORTS ON ASTAXANTHIN IN ASTASHINE CAPSULES**

##### ***Effect of astashine capsules on muscle endurance***

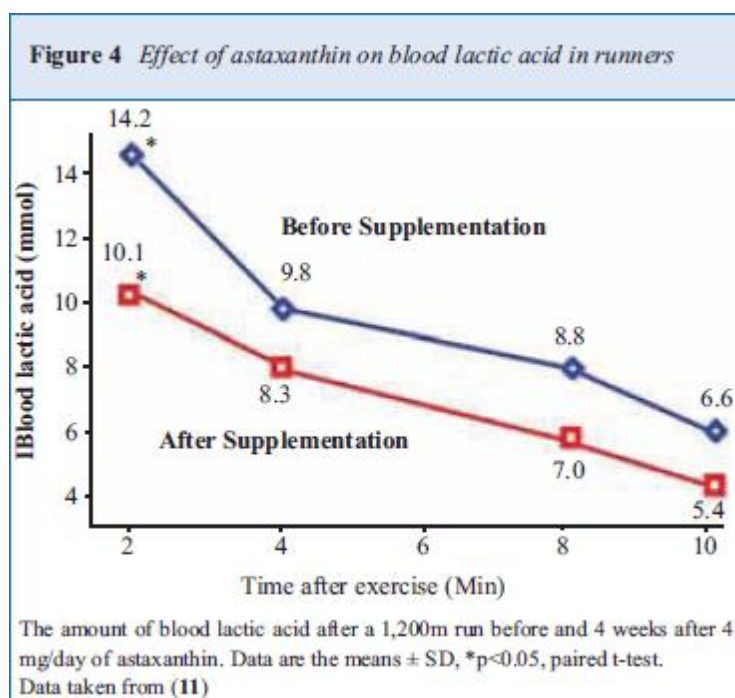
A randomized, double-blind study has shown that astaxanthin increases muscle endurance [10]. Forty-two healthy men were supplemented with 4 mg astaxanthin/day for 6 months. Standardized exercise tests demonstrated that the average number of knee bends performed increased only in the astaxanthin treated group at three months, and by six months significant improvements were observed [*Fig 3*].



**Figure 3** Effect of astaxanthin on muscle endurance in young men

Increased number of knee bends in healthy young men receiving placebo or 4 mg of astaxanthin per day. Data are expressed as means  $\pm$  SD; \* $p < 0.05$ , Student t-test. Data taken from [10]

In addition, Sawaki *et al* [11] demonstrated that a daily dose of 6 mg astaxanthin/ day for 4 weeks resulted in lower levels of lactic acid during a 1,200 meter sprint (**Fig 4**).

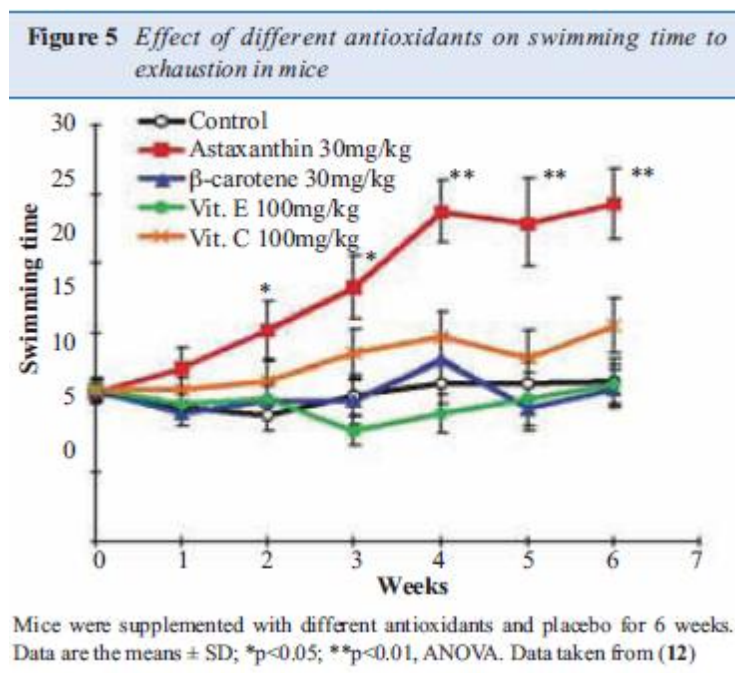


**Figure 4** Effect of astaxanthin on blood lactic acid in runners

The amount of blood lactic acid after a 1,200m run before and 4 weeks after 4 mg/day of astaxanthin. Data are the means  $\pm$  SD, \* $p$ <0.05, paired t-test. Data taken from (11)

The formation of lactic acid is a result of insufficient oxygen to muscles and leads to fatigue; a lower level of lactic acid will, therefore, improve endurance.

The effect of astaxanthin on muscle endurance is further supported by studies on mice. Ikeuchi *et al* [12] found that mice supplemented with astaxanthin for 5 weeks could swim for a significantly longer time to exhaustion as compared to placebo and other antioxidants (**Fig 5**).



**Figure 5** Effect of different antioxidants on swimming time to exhaustion in mice

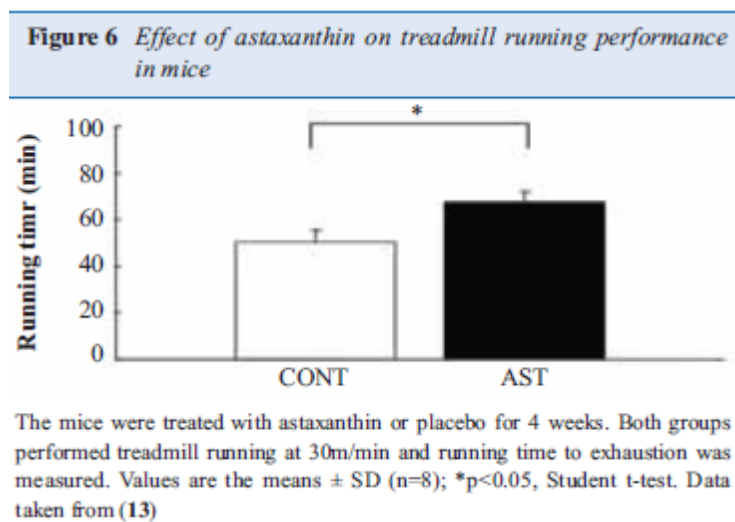
Mice were supplemented with different antioxidants and placebo for 6 weeks.

Data are the means  $\pm$  SD; \* $p$ <0.05; \*\* $p$ <0.01, ANOVA. Data taken from [12]

In the astaxanthin group, blood lactate concentration was significantly lower than the

control group while at the same time muscle and liver glycogen were higher.

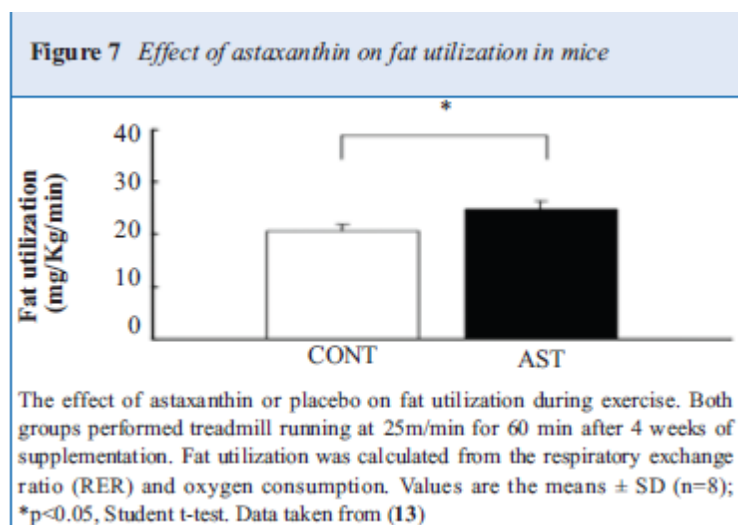
Another study by Aoi *et al* demonstrated similar results [13]. Running time to exhaustion for mice fed astaxanthin was significantly longer compared to control group as shown in *Figure 6*.



**Figure 6** Effect of astaxanthin on treadmill running performance in mice

The mice were treated with astaxanthin or placebo for 4 weeks. Both groups performed treadmill running at 30m/min and running time to exhaustion was measured. Values are the means  $\pm$  SD (n=8); \*p<0.05, Student t-test. Data taken from [13]

Plasma lactate was significantly increased by exercise, an effect that was prevented by the addition of astaxanthin to the diet. Additionally, astaxanthin increased muscle glycogen and increased fat utilization (**Fig 7**).



**Figure 7** Effect of astaxanthin on fat utilization in mice

The effect of astaxanthin or placebo on fat utilization during exercise. Both groups performed treadmill running at 25m/min for 60 min after 4 weeks of supplementation. Fat utilization was calculated from the respiratory exchange ratio (RER) and oxygen consumption. Values are the means  $\pm$  SD (n=8); \*p<0.05, Student t-test. Data taken from [13]

A better fat utilization during exercise contributes to a reduced level of lactic acid. It has been shown that astaxanthin enhanced fat burning by protecting the mitochondrial membrane bound enzyme, carnitine palmitoyltransferase

I (CPT I), which plays an important role in the entry of fatty acids into the mitochondria [13]. A randomized, double blind study on humans has



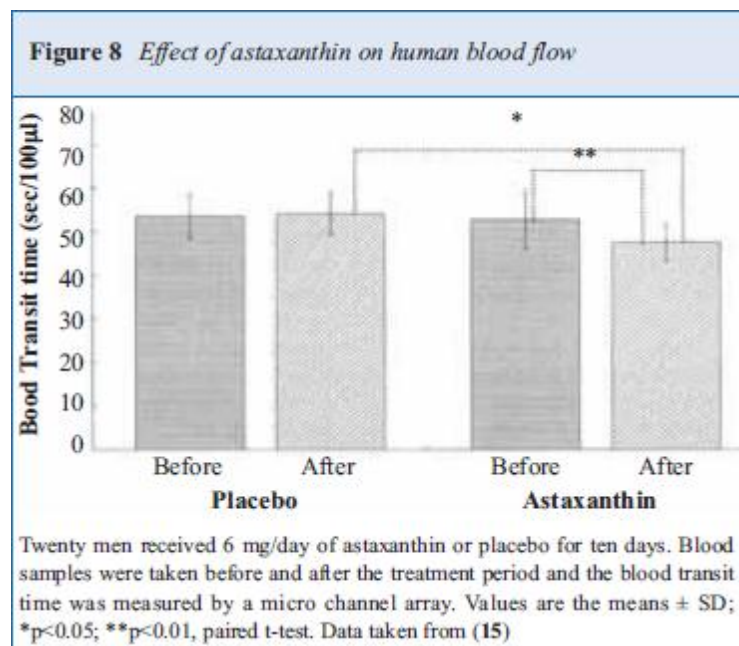
confirmed that astaxanthin increases fat utilization during exercise [14]. In that study, 32 subjects were supplemented with 2 x 6 mg astaxanthin/day, or placebo, for 6 weeks. The subjects were instructed to perform continuous exercise for a 40-min period 3 times per week during this 6-week period. After the 6 weeks, it was found that the astaxanthin group had a significantly reduced percentage of body fat while there were no difference in the placebo group. These results indicated that astaxanthin increases muscle endurance and reduces lactic acid during intensive training by promoting the use of fat and sparing glycogen.

### *Effect of astashine capsules on aerobic power*

Muscle cells are dependent on oxygen that is used by mitochondria to generate ATP and energy from carbohydrates and fats. If the muscle does not

have a sufficient amount of oxygen, it will resort to anaerobic metabolism with increased lactic acid production and muscle fatigue.

Two studies showed that astaxanthin was able to improve oxygen transport by reducing the oxidation of RBCs thus enhancing blood flow. In the first study, a double blind, randomized study [15], twenty male subjects received 6 mg of astaxanthin or placebo for ten days. Blood samples were taken before and after the treatment period and the blood transit time were measured by channeling the blood through a micro channel array. The results showed that the group treated with astaxanthin had a transit time 10% significantly lower than the placebo group (*Fig 8*). this effect was due to the ability of astaxanthin to protect blood cells from oxidation and thus improve blood rheology.



**Figure 8** *Effect of astaxanthin on human blood flow*

Twenty men received 6 mg/day of astaxanthin or placebo for ten days. Blood samples were taken before and after the treatment period and the blood transit time was measured by a micro channel array. Values are the means  $\pm$  SD; \* $p < 0.05$ ; \*\* $p < 0.01$ , paired t-test. Data taken from [15]

In a recent randomized, double blind study on humans [16], it was demonstrated that astaxanthin is taken up by the RBCs and reduces the oxidation of the RBC membrane.

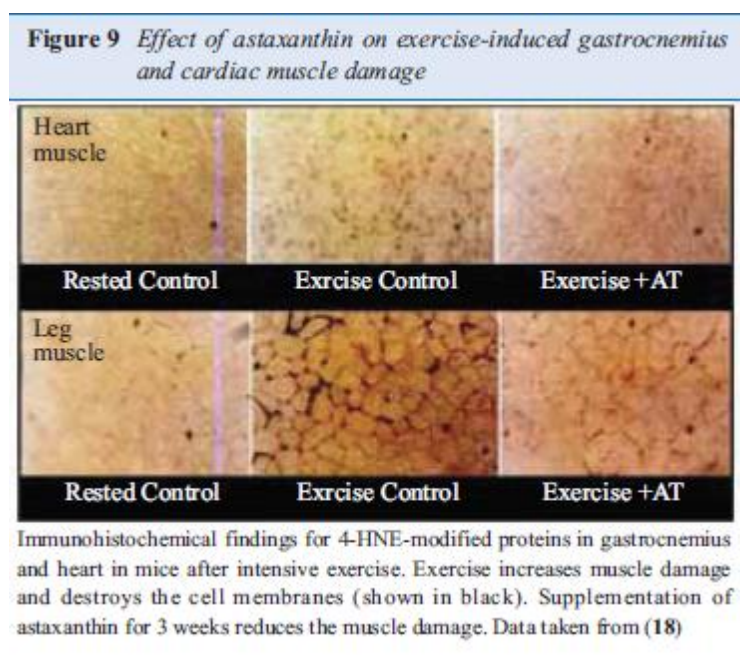
In this study, a total of thirty healthy subjects received 0 mg (placebo), 6 mg or 12 mg of astaxanthin per day for 12 weeks. The content of astaxanthin was increased both in plasma and in the RBCs among the subjects that were supplemented. Oxidation of the RBC membranes was significantly reduced by both doses of astaxanthin compared to placebo. These results indicate that astaxanthin can enhance the transportation of oxygen by reducing

the oxidation of RBCs and thus improve blood flow.

In other studies, Wolf *et al*, [17] demonstrated that astaxanthin stimulates mitochondrial respiration by maintaining a higher membrane potential in mitochondria, thus increasing, oxygen consumption. The effect is enhanced mitochondrial function followed by a better energy supply to muscles. These results indicate that astaxanthin boosts aerobic power by increasing oxygen transport to muscles and enhancing the capacity of mitochondria.

### *Effect of astashine capsules on muscle oxidation and inflammation*

Astaxanthin not only protects mitochondria and RBCs from oxidation, but it also protects heart and skeletal muscle cells from damage. In a study on mice [18], it was found that exercise increased oxidation and inflammation in heart and leg muscle while supplementation with astaxanthin contributed to significantly prevent the damage to both muscles (**Fig 9**).



**Figure 9** *Effect of astaxanthin on exercise-induced gastrocnemius and cardiac muscle damage*

Immunohistochemical findings for 4-HNE-modified proteins in gastrocnemius and heart in mice after intensive exercise. Exercise increases muscle damage and destroys the cell membranes. Supplementation of astaxanthin for 3 weeks reduces the muscle damage. Data taken from [18]

The anti-inflammatory effect of astaxanthin was further established in a randomized, double-blind, placebo controlled study on 42 subjects who were supplemented with 0 mg (placebo), 2 mg or 8 mg of astaxanthin for 8 weeks [19]. The groups treated with astaxanthin showed significantly reduced DNA damage caused by oxidation and plasma C-reactive protein. C-reactive protein increases in the blood as a result of inflammation.

According to these results, astaxanthin may lower muscle pain, stiffness and fatigue by reducing inflammation and oxidation in muscles. It is found in *in vitro* studies in mice that astaxanthin acts as an anti-inflammatory by inhibiting free radicals in the cell membrane that otherwise trigger NF-KB (20), a transcription factor that activates pro-inflammatory cytokines.

### *Effect of astashine capsules on risk of muscle atrophy*

Since astaxanthin seems to be able to protect muscle cells from damage, thus prevents muscle weakness (sarcopenia) in aging. In a study on mice, long-term dietary astaxanthin intake was found to prevent muscle atrophy by reducing oxidative



stress and degenerative proteins [21]. Mice were supplemented with astaxanthin or placebo for one year. After the treatment period, the weight of the soleus muscle was significantly higher for the astaxanthin group compared to placebo. The levels of degenerative proteins, e.g. cathepsin L and ubiquitinated myofibrillar protein, were also significantly lower compared to the placebo group.

The results indicate that astaxanthin increases muscle mass by protecting muscle cells from damage. In addition, astaxanthin might prevent muscle atrophy by improving the function of mitochondria [3,17].

## SAFETY OF ASTASHINE CAPSULES

Astaxanthin has demonstrated safety in numerous human clinical trials. In one open-label clinical study on subjects with metabolic syndrome (n=17). Astaxanthin (16 mg/day, for three months) significantly raised blood bilirubin ( $p \leq 0.05$ ), potassium ( $p \leq 0.05$ ), and creatine kinase ( $p \leq 0.01$ ), although all three values remained within normal range. Also, astaxanthin significantly lowered the liver enzyme gamma-glutamyl transpeptidase (GGTP;  $p \leq 0.05$ ). Since the researchers noted this enzyme was abnormally elevated in 11 of the 17 subjects at baseline, this astaxanthin effect may have been beneficial. Animal experiments have investigated astaxanthin at levels well over 120

mg/day in human equivalents, without causing apparent harm. Hoffman-La Roche confirmed its safety with extensive tests, including acute toxicity, mutagenicity, teratogenicity, embryotoxicity, and reproductive toxicity.

## Suggested Dosage

The doses of astaxanthin used in clinical trials have ranged from 1 mg/day to 40 mg/day (with the majority in the 6-12 mg range); single-dose pharmacokinetic studies used up to 100 mg per dose. As a dietary supplement, astaxanthin should be taken along with fats, with or immediately prior to meals, to ensure its optimal absorption.

## CONCLUSION

The interest of astaxanthin is growing among researchers and new findings about muscle health and endurance continue to appear. The results indicate that astaxanthin protects muscle cells and membranes from oxidative damage; in addition, it improves muscle endurance and lowers lactic acid mainly by improving fat utilization, mitochondrial function and reducing oxidation of RBCs. Studies further demonstrate that astaxanthin in Astashine capsules decreases inflammation which might reduce muscle pain and also shows potential for preventing muscle atrophy.

## REFERENCES

- [1]. Finaud J, Lac G, Filaire E Oxidative stress Relationship with exercise and training *Sports Med* 36(4), 2006, 327-358
- [2]. Tian Y, Nie J, Tong TK, Baker JS, Thomas NE, Shi Q Serum oxidant and antioxidant status during early and late recovery periods following an all-out 21-km run in trained adolescent runners *Eur J Appl Physiol* 110, 2010, 971-976
- [3]. Semba RD, Blaum C, Guralnik JM, Moncrief DT, Ricks MO *et al* Carotenoid and vitamin E status are associated indicators of sarcopenia among older women living in the community *Aging Clin Exp Res* 15(6), 2003, 482-487
- [4]. Berzosa C, Gomez-Trullen EM, Piedrafita E *et al* Erythrocyte membrane fluidity and indices of plasmatic oxidative damage after acute physical exercise in humans *Eur J Appl Physiol* 111(6), 2011, 1127-1133
- [5]. Brzezczynska J, Pieniazek A, Gwozdinski L *et al* Structural alterations of erythrocyte membrane components induced by exhaustive exercise *Appl Physiol Nutr Metab* 33(6), 2008, 1223-1231
- [6]. Goto S, Kogure K, Abe K, Kimata Y, Kitahima K *et al* Efficient radical trapping at the surface and inside the phospholipid membrane is responsible for highly potent antiperoxidative activity of the carotenoid astaxanthin *Biochim Biophys Acta* 1512, 2001, 251-258
- [7]. Nishida Y, Yamashita E, Miki W Quenching activities of common hydrophilic and lipophilic antioxidants against singlet oxygen using chemiluminescence detection system *Carotenoid Science* 11, 2007, 16-20
- [8]. Miki W Biological functions and activities of animal carotenoids *Pure Appl Chem* 1(63), 1991, 141-146

- [9]. Martin HD, Ruck C, Schmidt M, Sell S, Beutner S *et al* Chemistry of carotenoid oxidation and free radical reactions *Pure Appl Chem* 71(12), 1999, 2253-2262
- [10]. Malmsten CL, Lignell Å Dietary supplementation with astaxanthin-rich algal meal improves strength endurance. - A double blind placebo controlled study in mice *Carotenoid Sci* 13, 2008, 20-22.
- [11]. Sawaki K, Yoshigi H, Aoki K, Koikawa N *et al* Sports performance benefits from taking natural astaxanthin *J Clin Ther Med* 18(9), 2002, 73-88
- [12]. Ikeuchi M, Koyama T, Takahashi J, Yazawa K Effects of astaxanthin supplementation on exercise-induced fatigue in mice *Biol Pharm Bull* 29(10), 2006, 2106-2110
- [13]. Aoi W, Naito Y, Takanami Y, Ishii T, Kawai Y *et al* Astaxanthin improves muscle lipid metabolism in exercise via inhibitory effect of CPT I modification. *Biochem Biophys Res Commun* 366(4), 2008, 892-897
- [14]. Fukamauchi M Astaxanthin: an effective anti-peroxidant in food: the effect of aerobic exercise *Food Style* 21 11(10), 2007, 22-26
- [15]. Miyawaki H, Takashi J, Tsukahara H, Takehara I Effects of astaxanthin on human blood rheology *J Clin Biochem Nutr* 43, 2008, 69-74
- [16]. Nakagawa K, Kiko T, Miyazawa T, Burdeos GC *et al* Antioxidant effect of astaxanthin on phospholipid peroxidation in human erythrocytes *Br J Nutr* 2011, 311-9
- [17]. Wolf AM, Asoh S, Hiranuma H, Ohsawa I, Lio K *et al* Astaxanthin protects mitochondria redox state and functional integrity against oxidative stress *J Nutr Biochem* 21(5), 2010, 381-389
- [18]. Aoi W, Naito Y, Sakuma K, Kuchide M, Tokuda H *et al* Astaxanthin limits exercise-induced skeletal and cardiac muscle damage in mice *Antioxid Redox Signal* 5(1), 2003, 139-144.
- [19]. Park JS, Chyun JH, Kim YK, Line LL, Chew BP *et al* Astaxanthin decreased oxidative stress and inflammation and enhanced immune response in humans *Nutr Metab* 7, 2010, 18.
- [20]. Lee SJ, Bai S-K, Lee K-S, Namkoong S, Na H-J *et al* Astaxanthin inhibits nitric oxide production and inflammatory gene expression by suppressing I $\kappa$ B kinase-dependent Nf-kB activation *Mol Cells* 16(1), 2003, 97-105.
- [21]. Shibaguchi T, Sugiura T, Furumoto T, Inoue K *et al* Effect of long term dietary astaxanthin intake on sarcopenia *Jpn J Phys Fitness Sports Med* 57, 2008, 541-552.