

# Rhodiola: Rhodiola rosea, Artic root, Golden root

### **Common Indications:**

- Adaptogen/Adrenal stress
- Immunoprotection
- Fatigue
- Improving Cognition
- Depression
- Antioxidant

#### **General Comments:**

Rhodiola has a long history of use throughout Russia and China as an adaptogen and as an element for sports performance enhancement. It has shown itself to be an aid in supporting resilience, stress management, and enhancing mental and physical work capacity. Its actions have been used to fight stress induced depression, support nervous and immune system function, and even use in prevention of high altitude sickness.<sup>1,2,3,4,5,6,7,8,9</sup>

#### Benefits & Mechanism of Action:

#### Adaptogen/Anxiety

The activity of neurotransmitters such as serotonin, dopamine, and norepinephrine have been reportedly influence by use of Rhodiola. There is an adaptogenic effect that appears cardio protective and stabilizes central nervous system activities, possibly thru its impact on heat shock proteins.<sup>10,11,12,13,14,15,16</sup> It is believed the changes in monoamine levels are due to inhibition of the activity of enzymes responsible for monoamine degradation and facilitation of neurotransmitter transport within the brain.

A clinical study found that an extract of rhodiola significantly improved generalized anxiety disorder symptoms, with a reduction in HARS scores similar to that found in clinical trials.<sup>17</sup> When administered to physicians, supplementation of rhodiola favorably influenced fatigue and mental performance during the first two weeks on night duty.<sup>18</sup>

In addition to these central effects, rhodiola has been reported to prevent both catecholamine release and subsequent cyclic AMP elevation in the myocardium, and the depletion of adrenal catecholamines induced by acute stress.<sup>19,20.21</sup> Rhodiola's adaptogenic activity might also be secondary to induction of opioid peptide biosynthesis

and through the activation of both central and peripheral opioid receptors.<sup>22</sup>

### Immunoprotection

Rhodiola may also help improve immune function through regulation of IL-2 in Th1 cells and IL-4, IL-6, IL-10 in Th2 cells.<sup>23,24</sup>

### Fatigue

A 2009 phase III clinical trial found that a standardized rhodiola extract had anti-fatigue activity that increases mental performance, particularly the ability to concentrate, and decreases cortisol response to awakening stress in burnout patients with fatigue syndrome.<sup>25</sup>

### Improving Cognition

Rhodiola extract initiates the release of monoamine neurotransmitters. This activates the cerebral cortex and limbic system. This activation improves cognitive function, memory, attention, and learning. An extract of rhodiola in combination with vitamins/minerals was administered to 120 adults aged 50-89 years and reported to improve cognition in 81% of patients.<sup>26,27</sup>

### Depression

A review of the clinical studies supporting the effectiveness of rhodiola supplements was conducted in 2011. Eleven randomized, placebo-controlled human studies were investigated. It was found that rhodiola supplements may have beneficial effects on physical performance, mental performance, and mental health conditions like depression.<sup>28,29</sup> The authors concluded that more research seems warranted.

A 2012 systematic review supported the antidepressant effects of Rhodiola rosea.<sup>30</sup> A 2015 phase II randomized placebo controlled trial found that rhodiola may be a potential treatment alternative for patients with mild to moderate depression who are intolerant to the adverse effects of conventional antidepressants.<sup>31</sup>

Students receiving a standardized extract of rhodiola demonstrated significant improvements in physical fitness, psychomotor function, mental performance, and general well being. Subjects receiving the rhodiola extract also reported statistically significant reductions in mental fatigue, improved sleep patterns, a reduced need for sleep, greater mood stability, and a greater motivation to study.<sup>32</sup>

#### Antioxidant

In human and animal studies, rhodiola extract reduced oxidative stress and increased endogenous antioxidant production. It also protects human erythrocytes from glutathione depletion and glyceraldehyde-3-phosphate dehydrogenase inactivation.33,34,35,36

### Dose: 100-300mg, 1-3 times daily

**Standardization:** Rhodiola should be standardized to contain at least 3% rosavins and 1% salidroside.

Cautions & Side Effects: Rhodiola has been reported to be safe in recommended doses.

Symptoms that may indicate acute toxicity include:

- Insomnia
- Irritability
- Fatigue
- Headache

Rhodiola rosea is a mild stimulant, with no addictive potential, and should be taken in the morning to avoid sleep problems.

It may induce temporary vivid dreams and mild nausea.<sup>37</sup>

### Medication interactions

Medications with increased effects while taking rhodiola include:

- Adriamycin reduced toxicity
- Cyclophosphamide
- Antidepressants
- P-glycoprotein substrates
- Cytochrome 3A4 substrates
- Losartan

Patients with the following disease states or conditions should not use rhodiola:

• Bipolar depression

### **References:**

General Comments

- 1. Rhodiola rosea. Monograph. Altern Med Rev. 2002;7(5):421-3.
- 2. Battistelli M, De Sanctis R, De Bellis R, Cucchiarini L, Dacha M, Gobbi P. Rhodiola rosea as antioxidant in red blood cells: ultrastructural and hemolytic behaviour. Eur J Histochem. 2005;49(3):243-54.
- 3. De Bock K, Eijnde BO, Ramaekers M, Hespel P. Acute Rhodiola rosea intake can improve endurance exercise performance. Int J Sport Nutr Exerc Metab. 2004;14(3):298-307.
- 4. Goel HC, Bala M, Prasad J, Singh S, Agrawala PK, Swahney RC. Radioprotection by Rhodiola imbricata in mice against whole-body lethal irradiation. J Med Food. 2006;9(2):154-60.
- 5. Iaremii IN, Grigor'eva NF. [Hepatoprotective properties of liquid extract of Rhodiola

rosea]. Eksp Klin Farmakol. 2002;65(6):57-9.

- 6. Kanupriya, Prasad D, Sai Ram M, et al., Cytoprotective and antioxidant activity of Rhodiola imbricata against tert-butyl hydroperoxide induced oxidative injury in U-937 human macrophages. Mol Cell Biochem. 2005;275(1-2):1-6.
- 7. Kim SH, Hyun SH, Choung SY. Antioxidative effects of Cinnamomi cassiae and Rhodiola rosea extracts in liver of diabetic mice. Biofactors. 2006;26(3):209-19.
- Kucinskaite A, Briedis V, Savickas A. [Experimental analysis of therapeutic properties of Rhodiola rosea L. and its possible application in medicine]. Medicina (Kaunas). 2004;40(7):614-9.
- Kwon YI, Jang HD, Shetty K. Evaluation of Rhodiola crenulata and Rhodiola rosea for management of type II diabetes and hypertension. Asia Pac J Clin Nutr. 2006;15(3):425-32.

# Adaptogen/Adrenal stress

- 10. Ming DS, Hillhouse BJ, Guns ES, et al., Bioactive compounds from Rhodiola rosea (Crassulaceae). Phytother Res. 2005;19(9):740-3.
- 11. Kelly GS. Rhodiola rosea: a possible plant adaptogen. Altern Med Rev. 2001;6(3):293-302.
- Abidov M, Grachev S, Seifulla RD, Ziegenfuss TN. Extract of Rhodiola rosea radix reduces the level of C- reactive protein and creatinine kinase in the blood. Bull Exp Biol Med. 2004;138(1):63-4.
- Perfumi M, Mattioli L. Adaptogenic and central nervous system effects of single doses of 3% rosavin and 1% salidroside Rhodiola rosea L. extract in mice. Phytother Res. 2006 Oct 27; [Epub ahead of print].
- 14. Pogorelyi VE, Makarova LM. [Rhodiola rosea extract for prophylaxis of ischemic cerebral circulation disorder]. Eksp Klin Farmakol. 2002;65(4):19-22.
- 15. Tolonen A, Pakonen M, Hohtola A, Jalonen J. Phenylpropanoid glycosides from Rhodiola rosea. Chem Pharm Bull (Tokyo). 2003;51(4):467-70.
- 16. Walker TB, Robergs RA Does Rhodiola rosea possess ergogenic properties? Int J Sport Nutr Exerc Metab. 2006;16(3):305-15.
- 17. Bystritsky A, Kerwin L, Feusner JD. A Pilot Study of Rhodiola rosea (Rhodax((R))) for Generalized Anxiety Disorder (GAD). J Altern Complement Med. 2008;14(2):175-80.
- Darbinyan V, Kteyan A, Panossian A, Gabrielian E, Wikman G, Wagner H. Rhodiola rosea in stress induced fatigue--a double blind cross-over study of a standardized extract SHR-5 with a repeated low-dose regimen on the mental performance of healthy physicians during night duty. Phytomedicine. 2000;7(5):365-71.
- 19. Maslova LV, Kondrat'ev BI, Maslov LN, Lishmanov IB. The cardioprotective and antiadrenergic activity of an extract of Rhodiola rosea in stress. Eksp Klin Farmakol. 1994;57:61-63.
- 20. Mishra KP, Padwad YS, Jain M, Karan D, Ganju L, Sawhney RC. Aqueous extract of Rhodiola imbricata rhizome stimulates proinflammatory mediators via phosphorylated IkappaB and transcription factor nuclear factor-kappaB. Immunopharmacol Immunotoxicol. 2006;28(2):201-12.
- 21. Ohsugi M, Fan W, Hase K, et al., Active-oxygen scavenging activity of traditional

nourishing-tonic herbal medicines and active constituents of Rhodiola sacra. J Ethnopharmacol. 1999;67(1):111-9.

22. Lishmanov IB, Naumova AV, Afanas'ev SA, Maslov LN. Contribution of the opioid system to realization of inotropic effects of Rhodiola rosea extracts in ischemic and reperfusion heart damage in vitro. Eksp Klin Farmakol. 1997;60:34-36.

# Immunoprotection

- 23. Li HX, Sze SC, Tong Y, Ng TB. Production of Th1- and Th2-dependent cytokines induced by the Chinese medicine herb, Rhodiola algida, on human peripheral blood monocytes. J Ethnopharmacology. 2009;123(2):257-66.
- 24. Wing SL, Askew EW, Luetkemeier MJ, Ryujin DT, Kamimori GH, Grissom CK. Lack of effect of Rhodiola or oxygenated water supplementation on hypoxemia and oxidative stress. Wilderness Environ Med. 2003;14(1):9-16.

# Fatigue

25. Olsson EM, von Schéele B, Panossian AG. A randomised, double-blind, placebocontrolled, parallel-group study of the standardised extract shr-5 of the roots of Rhodiola rosea in the treatment of subjects with stress-related fatigue. Planta Med. 2009 Feb;75(2):105-12. Epub 2008 Nov 18.

# Improving Cognition

- 26. Fintelmann V, Gruenwald J. Efficacy and tolerability of a Rhodiola rosea extract in adults with physical and cognitive deficiencies. Adv Ther. 2007 Jul-Aug;24(4):929-39.
- 27. Shevtsov VA, Zholus BI, Shervarly VI, Vol'skij VB, Korovin YP, Khristich MP, Roslyakova NA, Wikman G. A randomized trial of two different doses of a SHR-5 Rhodiola rosea extract versus placebo and control of capacity for mental work. Phytomedicine. 2003;10(2-3):95-105.

### Depression

- 28. Hung SK, Perry R, Ernst E. The effectiveness and efficacy of Rhodiola rosea L.: a systematic review of randomized clinical trials. Phytomedicine. 2011;18(4):235-44.
- 29. Abidov M, Crendal F, Grachev S, Seifulla R, Ziegenfuss T. Effect of extracts from Rhodiola rosea and Rhodiola crenulata (Crassulaceae) roots on ATP content in mitochondria of skeletal muscles. Bull Exp Biol Med. 2003;136(6):585-7.
- 30. Chan SW. Panax ginseng, Rhodiola rosea and Schisandra chinensis. *Int J Food Sci Nutr*. 2012;63 Suppl 1:75–81.
- 31. Mao JJ, Xie SX, Zee J, et al., Rhodiola rosea versus sertraline for major depressive disorder: A randomized placebo-controlled trial. Phytomedicine. 2015 Mar 15;22(3):394-9.
- 32. Spasov AA, Wikman GK, Mandrikov VB, Mironova IA, Neumoin VV. A double-blind, placebo-controlled pilot study of the stimulating and adaptogenic effect of Rhodiola

rosea SHR-5 extract on the fatigue of students caused by stress during an examination period with a repeated low-dose regimen. Phytomedicine. 2000;7(2):85-9.

# Antioxidant

- 33. Battistelli M et al. rhodiola rosea as antioxidant in red blood cells: ultrastructural and hemolytic behaviour. Eur J Histochem 49.3 (2005): 243–254.
- De Sanctis R et al. In vitro protective effect of rhodiola rosea extract against hypochlorous acid-induced oxidative damage in human erythrocytes. Biofactors 20.3 (2004): 147–159.
- 35. Kim SH et al. Antioxidative effects of Cinnamomi cassiae and rhodiola rosea extracts in liver of diabetic mice. Biofactors 26.3 (2006): 209–219.
- 36. Wing SL et al. Lack of effect of rhodiola or oxygenated water supplementation on hypoxemia and oxidative stress. Wilderness Environ Med 14.1 (2003): 9–16.

# Cautions & Side Effects

37. Van Diermen D, Marston A, Bravo J, Reist M, Carrupt PA, Hostettmann K. Monoamine oxidase inhibition by Rhodiola rosea L. roots. *J Ethnopharmacol*. 2009;122:397–401.