

## Kavinace® OS

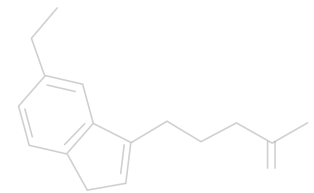
Target the cellular consequences of sleeplessness with a formulation proven to improve both sleep onset and efficiency after the first serving<sup>1,2\*</sup>

### Patient Profile

- Requires quick relief from sleeplessness
- Needs help falling asleep
- Unhealthy lifestyle habits
- Known or suspected immune activity

## Key Ingredients

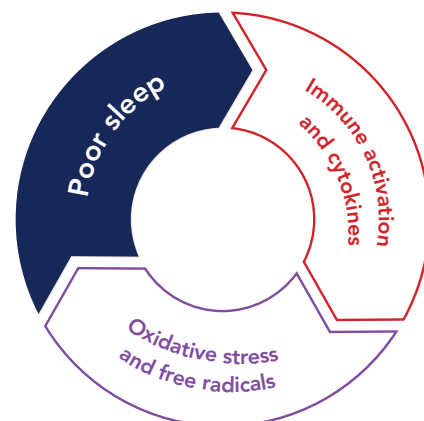
<p>L-theanine</p>	<ul style="list-style-type: none"> <li>■ Amino acid that acts as a glutamate receptor antagonist, shown to <b>reduce stress</b><sup>**</sup></li> <li>■ Modulates the immune response by regulating the secretion of <b>INF-γ, IL-2 and IL-10</b>, cytokines shown to increase after strenuous activity and <b>display sleep regulating properties</b><sup>20-23</sup></li> </ul>
<p>Astaxanthin</p>	<ul style="list-style-type: none"> <li>■ Carotenoid that can have inhibitory effects on macrophage activity, IL-1 and IL-6 expression, and Nf-kB phosphorylation, impacting chronic and acute immune responses<sup>17,18*</sup></li> <li>■ Shown to <b>improve sleep onset</b> when taken with zinc<sup>6*</sup></li> </ul>
<p>Magnesium (as magnesium bisglycinate chelate)</p>	<ul style="list-style-type: none"> <li>■ Cofactor for multiple mechanisms in the body including the <b>production of serotonin</b> and acting as a <b>GABA agonist</b><sup>7*</sup></li> </ul>
<p>Zinc (as zinc bisglycinate chelate)</p>	<ul style="list-style-type: none"> <li>■ Essential micronutrient cofactor for the antioxidant enzyme <b>superoxide dismutase (SOD)</b><sup>**</sup></li> <li>■ Zinc deficiency is directly associated with increased biomarkers of oxidative stress and inflammatory cytokines<sup>9</sup></li> </ul>
<p>Melatonin</p>	<ul style="list-style-type: none"> <li>■ <b>Antioxidant</b> and hormone important for the <b>regulation of the sleep-wake cycle</b><sup>10*</sup></li> <li>■ Melatonin restores the <b>circadian rhythm dependent activity of mast cells</b>, important for the allergic and inflammatory cascade and immune response to pathogens<sup>19</sup></li> </ul>



## The Science

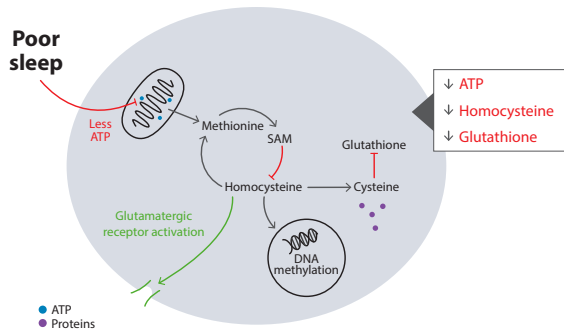
- The nervous system works with the immune system to regulate the sleep-wake cycle and the immune response<sup>11,12</sup>
- During daytime activity, the immune system generates free radicals and depletes antioxidants<sup>13</sup>
- During bedtime hours, accumulated free radicals can stimulate the immune system with the adaptive immune system at its most active<sup>12</sup>
- Poor sleep has been shown to increase oxidative stress markers, perpetuating the Immune-Sleep Cycle<sup>13-15</sup>

## The Immune-Sleep Cycle



\*These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure or prevent any disease.

Figure 1. Cellular Impact of Sleep Deprivation



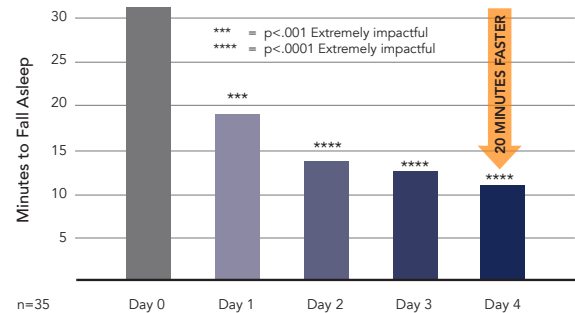
## One night of poor sleep changes cellular metabolic function (Figure 1).<sup>1</sup>

Metabolic changes worsen the Immune-Sleep Cycle, perpetuating more sleepless nights<sup>1</sup>

- Decreased antioxidant capacity increases susceptibility to negative effects of free radicals and oxidative stress<sup>1</sup>
- Oxidative stress modulates immune regulators like Nf-kB<sup>16</sup>
- Wake promoting substances (hormones and neurotransmitters) respond to immune activation and further disrupt sleep<sup>11, 15</sup>

The uniquely formulated blend of **Kavinace OS** comprehensively intervenes at every portion of the **Immune-Sleep Cycle\***

Figure 2. Kavinace OS Decreases Sleep Latency



## Researched ingredient blend, proven results\*

Data gathering participants were prescreened for poor sleep (PSQI), received one serving of Kavinace OS at bedtime for seven days, and submitted a daily sleep diary

- **Fall asleep quickly.** Reductions in sleep latency were reported after the first serving of Kavinace OS, with ongoing improvement through day four<sup>2</sup>
- **Sleep better.** Improvement in sleep efficiency, or total sleep time relative to time in bed, was reported after one serving of Kavinace OS<sup>2\*</sup>

**Kavinace OS provides quick relief from sleeplessness at the symptom and cellular level\***

NeuroScience supplements undergo rigorous, product specific third-party testing to guarantee label claims of each ingredient and the absence of heavy metals and microbes

Suggested Use: Take 2 capsules at bedtime or as directed by your healthcare provider.

Supplement Facts	
Serving Size: 2 Capsules	
Servings Per Container: 30	
Amount Per Serving	% Daily Value
Magnesium (as magnesium bisglycinate chelate)	25 mg 6%
Zinc (as zinc bisglycinate chelate)	15 mg 136%
Selenium (as selenomethionine)	200 mcg 364%
Melatonin	5 mg †
<b>Proprietary Blend</b>	214 mg †
L-theanine, Trans-resveratrol ( <i>Polygonum cuspidatum</i> ) (root), and Astaxanthin ( <i>Haematococcus pluvialis</i> ).	
† Daily Value not established.	

Other ingredients: Vegetable capsule (hypromellose, water), organic rice concentrate, microcrystalline cellulose, dicalcium phosphate, citric acid, and glycine.



Item Number	Available Sizes	Serving Size
20053	60 capsule	2 Capsules

- Trivedi M, et al. PLoS One. 2017;12(7):e0181978.
- Data on file. 2019. NeuroScience, Osceola, WI 54020.
- Kakuda T, et al. Biosci Biotechnol Biochem. 2002;66(12):2683-6.
- Kimura K, et al. Biol Psychol. 2007;74(1):39-45.
- Wu H, et al. Mar Drugs. 2015;13(9):5750-66.
- Saito H, et al. Mol Nutr Food Res. 2017;61(5):1600882.
- Schwalfenberg G, et al. Scientifica. 2017;4179326.
- Neddi S, et al. J Mol Biol. 2014;426(24):4112-4124.
- Prasad AS and Bao B. Antioxidants (Basel). 2019 Jun 6;8(6). pii:E164.
- Pandi-Perumal S, et al. Prog Neurobiol. 2008;85(3):225-53.

- Pongratz G and Straub R. Arthritis Res Ther. 2014;16:504.
- Cermakian N, et al. Chronobiol Int. 2013 Aug;30(7):870-88.
- Gulec M, et al. Prog Neuropsychopharmacol Biol Psychiatry. 2012 Jun 1;37(2):247-51.
- Bryant P, et al. Nat Rev Immunol. 2004;4:457-67.
- Luster FS, et al. Sleep. 2012;35(6):727-34.
- Lugin et al. 2014; Hussain et al. 2016; Liu et al. 2017.
- Lee S, et al. Astaxan Inhib Nitric. 2003; 16(1):97- 105.
- Kishimoto Y, et al. Eur J Nutr (2010) 49: 119.
- Carpentieri et al./Pharmacological Research 65 (2012).

- Juszkiewicz et al. 2019. Journal of the International Society of Sports Nutrition 16:7.
21. Kushikata T, Fang J, Krueger JM. 1999. J Interferon Cytokine Res 19: 1025-1030.
22. Kubota T, Brown RA, Fang J, Krueger JM. Am J Physiol Regul Integr Comp Physiol 281: R1004-R1012, 2001.
23. Kubota T, Majde JA, Brown RA, Krueger JM. 2001. J Neuroimmunol 119:192-198.

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