Hyposerotonergic<sup>™</sup> state or condition occurs when serotonin concentrations are not enough, low, inadequate, depleted, deficient, or suboptimal on a modified normal diet.<sup>™</sup> Hypodopaminergic<sup>™</sup> state or condition occurs when dopamine concentrations are not enough, low, inadequate, depleted, deficient, or suboptimal on a modified normal diet.™

# E MEDICAL FOOD

**Hypoglutathionemia<sup>™</sup> state or condition** occurs when glutathione concentrations are not enough, low, inadequate, depleted, deficient, or suboptimal on a modified normal diet.<sup>™</sup>

Giving only serotonin precursors can deplete dopamine and glutathione.™ Giving only dopamine precursors can deplete serotonin and glutathione.™ Giving only glutathione or glutathione precursors can deplete serotonin and dopamine.™

Hinz Medical Foods<sup>™</sup> / NeuroResearch Centers, Inc.<sup>™</sup> 1150 88<sup>th</sup> Ave W Duluth, MN 55808 +1-218-626-2220 www.HinzMedicalFoods.com

For the management of hyposerotonergic conditions or states that may accompany

## **Migraine Headaches**

A hyposerotonergic condition or state often accompanies migraine headaches (see the right column).

After diagnosing a migraine, formulate a differential diagnosis to rule out accompanying issues, including a hyposerotonergic condition or state.

Identify the presence of a hyposerotonergic condition or state with an empirical trial of the hyposerotonergic condition or state protocol (see below).

Management of the hyposerotonergic condition or state which may accompany the migraine headache requires establishing serotonin concentrations higher than are possible with modification of the normal diet. "Generally, it was stated that migraine patients have low platelet 5-HT, a sign of hyposerotonergic status of migraine." Panconesi A. Serotonin and migraine: a reconsideration of the central theory J Headache Pain (2008) 9:267-276

"The low platelet serotonin in migraine almost certainly reflects reduced concentrations of the amine in other parts of the body" Anthony, M. et al. Journal of Neurology, Neurosurgery, and Psychiatry 1989;52:182-184

"In addition, patients with migraine have low plasma serotonin between migraine attacks"

Lin, Y. et al. Association of Suicide Risk with Headache Frequency Among Migraine Patients with and without Aura Frontiers in Neurology March 2010 Vol 10 Article 228

## "In the current study, we identified low levels of serum serotonin in migraine patients"

Ren, C. et al. Biochemical and Biophysical Research Communications Volume 496, Issue 2, 5 February 2018, Pages 267-273

# "It has been suggested that migraine might be characterized as a symptom of chronic low serotonin levels or, alternately, a dysregulation of the serotonin system"

Pergolizzi, J. et al. Red wine triggers may lead to betterunderstanding of migraine headache: a narrative review J. Wine Research 2019, VOL. 30, NO. 1, 15–30

#### "Accordingly, although plasma levels of 5-HT (serotonin) do not necessarily reflect brain 5-HT levels, migraine has been considered a syndrome of chronically low brain 5-HT levels."

Deen, M. et al. High bradin serotonin levels in migraine between attacks: A 5-HT4 receptor binding PET study NeuroImage Clnical 18 (2018) 97-102



#### Hyposerotonergic Condition Protocol™

		AM	NOON	4 pm
Day-0	Level 1	3 R&R		3 R&R
Day-7	Level 2	3 R&R	3 R&R	2 R&R Sans
Day-14	Level 3	3 R&R	3 R&R	4 R&R Sans
Day-21 - If symptoms are still present after seven days on level 3 submit a				
specimen for serotonin and dopamine assay to DBS Labs, 1-877-476-7229				

**Figure 1**: Dosing levels 1-3 of the hyposerotonergic condition protocol do not require lab testing. Do not increase to level 4 through level 9 without first obtaining a serotonin and dopamine assay. Only increase to the next level if symptoms are present after seven days.



Hinz Medical Foods™ For the management of Drug-induced hyposerotonergic conditions



Platelets contain 99% of whole blood serotonin<sup>1</sup>

Medical science literature documents the magnitude of SSRI-induced serotonin depletion:

- SSRIs may deplete<sup>™</sup> 80% of platelet serotonin in two weeks<sup>2</sup>
- SSRIs may deplete 90% of platelet serotonin in three weeks<sup>3</sup>
- Eventually, lab assay of platelet serotonin may become undetecable<sup>4</sup> For a more in-depth depletion bibliography, go to https://hinzmedicalfoods.com/bibliography-1/

Drugs that work with serotonin do not work if there are not enough (depleted) serotonin.™

When SSRIs deplete serotonin, the only way to increase the synthesis of the total number of serotonin molecules in the system is by administering the required nutrients as found in R&R<sup>™</sup>.

### A POTENTIAL SOLUTION - R&R<sup>™</sup>

R&R<sup>™</sup> can achieve serotonin concentrations higher than a modified normal diet alone while addressing the ability of serotonin precursors to deplete dopamine (catecholamines) and glutathione (thiols).

To order for the clinic, pharmacy, or to authorize online ordering by the patient contact NeuroResearch Centers, Inc. +1-218-626-2220 1150 88th Ave W, Duluth, MN | Brenda@NeuroAssist.com

R&R is a medical food administered enterally under the supervision of a healthcare professional for the specific dietary management of hyposerotonergic conditions or states.

<sup>1</sup>McCloskey, D, et al. Selective Serotonin Reuptoke Inhibitors (SSRIs): Measurement of Effect on Platelet Function Transl Res. 2008 March; 151(3): 168–172 <sup>2</sup>Jordan, S. et al. Serotonergic agents increase the incidence of Gl bleeds in patients with continuous/flow left ventricular assist devices J head and Jung tplt 05 Jan 2016, 35(6):823-824 <sup>3</sup>Wagner, A. et al. Effects of fluwerine the fatelet Al-imigramine binding. 5-HT uptake and 5-HT content in major depressive disorder, J of Affective Disorders, 20 (1990) 101-113 <sup>4</sup>Mauer-Spurej E. et al. The influence of selective serotonin reuptake inhibitors on human platelet serotonin Thromb Haemost 2004;91:119-28

Recommended daily starting dose:

FOR THE MANAGEMENT OF HYPOSEROTONERGIC CONDITIONS AND STATEST

two tablets, three times a day To be used under the supervision of a licensed caregiver Hyposerotonergic<sup>™</sup> conditions occur when serotonin concentrations are not enough, low, inadequate, depleted, deficient, or suboptimal on a modified normal diet. Hypodopaminergic<sup>™</sup> conditions occur when dopamine concentrations are not enough, low, inadequate, depleted, deficient, or suboptimal on a modified normal diet.



**Hypoglutathionemia™ conditions** occur when glutathione concentrations are not enough, low, inadequate, depleted, deficient, or suboptimal on a modified normal diet.

Giving only serotonin precursors can deplete dopamine and glutathione.™ Giving only dopamine precursors can deplete serotonin and glutathione.™ Giving only glutathione or glutathione precursors can deplete serotonin and dopamine.™

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# The carbidopa-induced vitamin B6 hypovitaminosis condition

**Fact #1: Carbidopa depletes vitamin B6.** The U.S. Recommended Daily Allowance (USRDA) of vitamin B6 is about 2 mg per day. Carbidopa forms an irreversible bond with vitamin B6 in a 1:1 ratio (one mg of carbidopa will permanently remove one mg of B6). The maximum recommended dosing of carbidopa is 200 mg per day.<sup>1</sup> Administering 200 mg per day may remove one hundred times the B6 USRDA. A 2020 lab study reported 79.2% of patients taking carbidopa for more than three years had a vitamin B6 hypovitaminosis condition (vitamin B6 deficiency). The lab results on almost half of these B6 deficiencies (47.3%) documented no detectable systemic vitamin B6.<sup>2</sup>

1 Lodosyn prescribing information: https://www.accessdata.fda.gov/drugsatfda\_docs/label/2014/017830s014s016s017s018s019s030lbl.pdf

2 Rojo-Sebastián, A. et al. Vitamin B6 Deficiency in Patients With Parkinson Disease Treated With Levodopa/Carbidopa clinical Neuropharmacology Vol 43, Number 5, Sep/Oct 2020FDA

Fact #2: Vitamin B6 deficiency increases the death rate (mortality). A ten-year prospective study (N = 134,480) concluded, "...dietary vitamin B6 consumption was inversely associated with risk of all-cause and CVD (cardiovascular disease) mortality." Zhoa, L. et al. Prospective cohort studies of dietary vitamin B6 intake and risk of cause-specific mortality, Volume 38, Issue 3, June 2019, Pages 1180-1187

#### Fact #3: The Parkinson's disease death rate has increased by 561% since carbidopa sales started (1975).

Sales of L-dopa (without carbidopa) began in 1960. Between 1960 and 1976, the Parkinson's disease death rate decreased from 2.6 to 1.633 per 100k. Carbidopa sales started in 1975. Between 1976 and 2019, the Parkinson's death rate increased from 1.633 to 10.8 per 100k, a 561% increase. Giving B6 to compensate for carbidopa-induced B6 hypovitaminosis condition while taking carbidopa is not an option. Fully compensating for B6 deactivates all active carbidopa in the system. Center for Disease Control National Vital Statistics Accessed from: https:// www.cdc.gov/nchs/products/index.htm



TO ORDER FOR THE CLINIC, PHARMACY, OR TO INITIATE PATIENT ONLINE ORDERING CONTACT NEURORESEARCH CENTERS +1-218-626-2220 1150 88TH AVE W, DULUTH, MN | BRENDA@NEUROASSIST.COM Hinz Medical Foods indicated for management of the hypodopaminergic, hyposerotonergic, or hypoglutathionemia conditions and states.