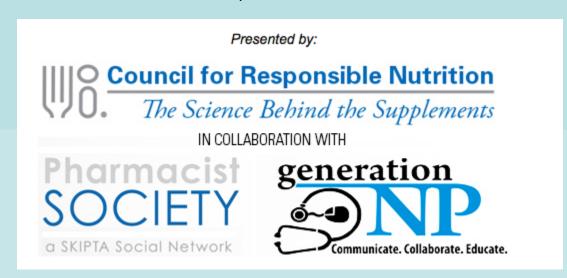


The Role of Dietary Supplements in Maintaining Cognitive Health and Mental Hygiene

James Greenblatt, M.D.

Chief Medical Officer, Walden Behavioral Care





Integrative Medicine



"What we have now is a 'sick care' system that is reactive to problems. The integrative approach flips the system on its head and puts the patients at the center, addressing not just symptoms, but the real causes of illness. It is care that is preventive, predictive and personalized."



Predictive Personalized Preventative Participatory Ralph Snyderman, MD Chancellor Emeritus Duke University School of Medicine

Lee Hood, MD, PhD P4MI Chairman and ISB President



The Problem



- Depression is one of the most serious and costly health problems in the world today
- Major Depression accounts for the 2nd longest number of days lost to disability in the U.S.
- Approximately 15% of adults will experience severe depressed mood during their lifetime
 - Approx. 15% of these adults will commit suicide
- Suicide is the 3rd leading cause of death in youth aged 15 to 24
- An Average of 25 Suicides per day occur for US Veterans (that we are aware of).
- One death from suicide every 40 seconds. It is predicted that by 2020, the rate of death from suicide will increase to one every
 20 seconds.



Diagnosis and Treatment



General Medical Treatment:

Symptoms



Measure Physiology



"Anti"-physiology treatment



Measure physiology and symptoms

Psychiatric Treatment:

Symptoms



"Anti"-Symptom treatment given



Measure symptoms

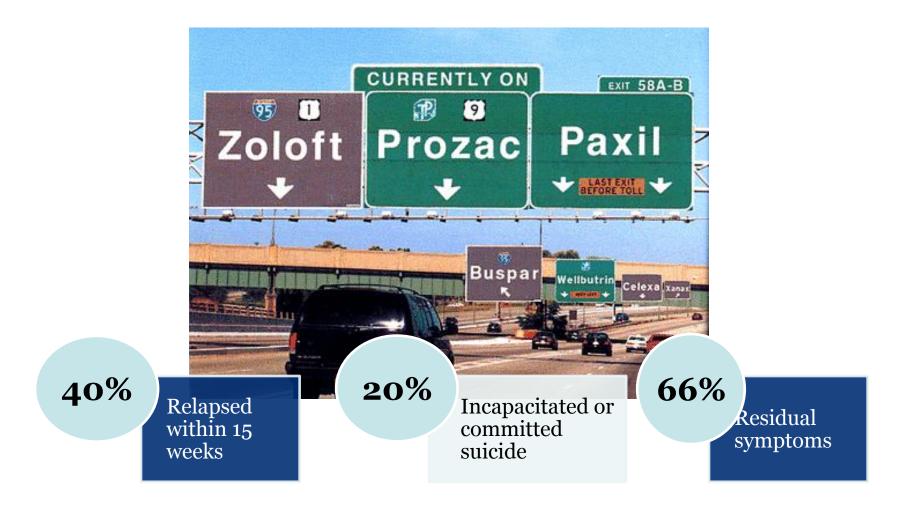
Psychiatric Testing





Polypharmacy Highway





THE ZEEBRA Approach



T – Take Care of Yourself – Stress, Sleep, Sugar

H – Hormones – Thyroid, Sex Hormones

E – Exclude – Celiac, Allergies

Z – Zinc and Other Minerals – Mg, Li

E – Essential Fatty Acids – Cholesterol

E – Exercise and Energy

B – B Vitamins & Other Vitamins – B12, Vitamin D

R – Restore – GI tract, Probiotics

A – Amino Acids and Protein – Digestive and Absorption



Cultural Chaos





The Brain



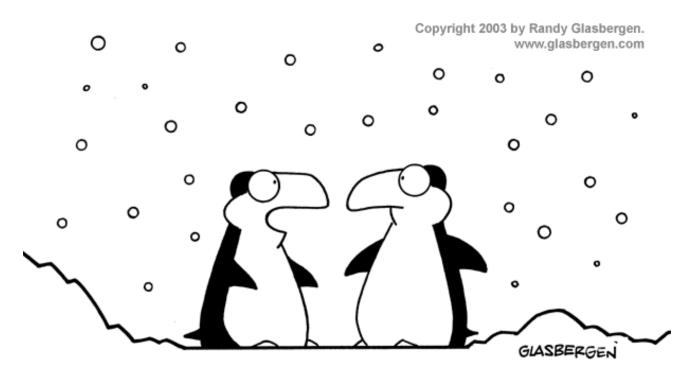


60% of the dry weight of the brain is fat

Approximately 30% of fatty acid pool cannot be made de novo and must be obtained through diet





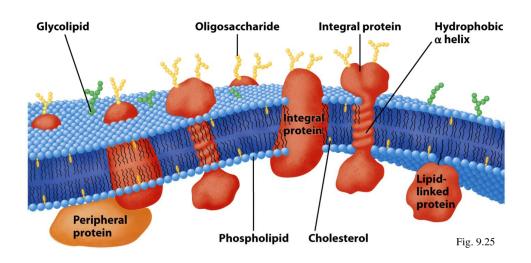


"Low fat diets don't work. I eat fish every day and my butt still drags on the ground!"



Omega-3 Fatty Acids





Inflammation research studies suggest that levels of omega-3 fatty acids may directly influence the magnitude of the inflammatory response to stress and depression.

Neurotransmitters

Every aspect of neurotransmission involves adequate functioning of omega 3s.



Omega-3s and Suicide in the Military



- Study of 800 U.S. servicemen and women who committed suicide between 2002 and 2008 (compared to 800 who didn't)
- Personnel with medical records showing low blood levels of DHA were 62% more likely to have been suicide victims than those with the highest levels
- Study found that U.S. service personnel generally have low levels of DHA in their blood



Omega-3 Fatty Acids Augmentation of Antidepressants



- 42 patients (40.5 y/o) with dietary intake of Omega 3
 3 gms/day
- DBPC 1.8 gms EPA .4gms DHA Omega 3 supplements or placebo BID x 8 weeks
- Celexa 20-40mg
- Higher proportion of patients achieved full remission in Omega 3 group versus the placebo group
 - 44% versus 18%

Gertsik L et al. J Clin Psychopharmacol 2012;32(1):61-4.



Prevention is Possible



- Omega-3s may have the ability to delay or prevent psychosis
- 81 adolescents or young adults with subthreshold psychosis
- 1.2 g omega-3 fatty acids or placebo daily for 12 weeks
- After 40 weeks:
 - 5% (2 out of 41 individuals) in omega-3 group developed psychosis
 - 28% (11 of 40 individuals) in placebo group developed psychosis

Amminger, et al., Archives of General Psychiatry, 2010, 67(2): 146-154.



Inflammation as a Predictive Biomaker as Response





Original Article

Molecular Psychiatry advance online publication 24 March 2015; doi: 10.1038/mp.2015.22

Inflammation as a predictive biomarker for response to omega-3 fatty acids in major depressive disorder: a proof-of-concept study

M H Rapaport¹, A A Nierenberg², P J Schettler¹, B Kinkead¹, A Cardoos², R Walker² and D Mischoulon²

¹Department of Psychiatry and Behavioral Sciences, Emory University School of Medicine, Atlanta, GA, USA
²Depression Clinical and Research Program, Massachusetts General Hospital, Harvard Medical School, Boston, MA, USA

Correspondence: Dr MH Rapaport, Department of Psychiatry and Behavioral Sciences, Emory University School of Medicine, Suite 4000 WMB, 101 Woodruff Circle, Atlanta, GA 30322, USA. E-mail: mrapapo@emory.edu

Received 19 September 2014; Revised 21 January 2015; Accepted 23 January 2015 Advance online publication 24 March 2015



Importance of Cholesterol



- The brain is a "cholesterol rich" organ
- Cholesterol is involved in the synthesis of all steroid hormones
- Activates serotonin and oxytocin receptors
- Bile salt formation for fat digestion and absorption of fat soluble vitamins
- Synthesis of vitamin D



Prevention is Possible



Low Cholesterol is associated with...

- Increased cancer rates
- Increased violent behavior and aggression
- Increased anxiety
- Increased depression and suicide
- Increased death rate
- Increased incidence of stroke
- More difficult recovery from drug addiction
- Increased rate of school suspensions



Integrative Psychiatry



- 42 patients (40.5 y/o) with dietary intake of Omega 3 <3 gms/day
- DBPC 1.8 gms EPA .4gms DHA Omega 3 supplements or placebo for 8 weeks
- Celexa 20-40mg
- Higher proportion of patients achieved full remission in Omega 3 group versus the placebo group
 - 44% versus 18%



Not a Quick Fix





It takes at least 10 weeks for cerebral membranes' highly unsaturated fatty acid levels to recover following chronic deficiency.

Bourre, et al. Prostaglandins Leukot Essent Fatty Acids 1993.



The Human Brain: The Franklin Institute





"By modifying natural fats, we have altered the basic building blocks of the human brain – weakening cerebral architecture. And, like unstable buildings that come apart in an earthquake or storm, poorly structured human brains are failing to cope with the mounting stress of modern life."

Non-Gastrointestinal Manifestations of Celiac Disease



Most common presentation of age: older child to adult

- Nutrient malabsorption
- Iron deficiency
- Short stature
- Hair Loss
- Dermatitis Herpetiformis
- Irritability and behavioral changes
- Liver enzyme elevation

- Delayed puberty (no menstrual cycle)
- Infertility
- Osteoporosis/ osteopenia
- Malnutrition
- Fatigue
- Depression



Deficiencies in Celiac Disease



When a disorder such as Celiac Disease (CD) affects the intestine, the absorption of almost all elements is impaired.

•Essential fatty acids

•Iron

•Fat soluble vitamins (A,D,E,K)

•Magnesium

•Folic Acid

•B Vitamins

•Zinc

One study involving 12 patients presenting with CD and depression who had been on a gluten-free diet for a year without improvement of the depression, showed that malabsorption not only was an issue, but was also inevitable even if the diet was 100% gluten-free.1

Hallert, C. (1982). *Biol Psychiatry* 17(9): 959-61.



Zinc Supplements for Depression



- Serum zinc levels are inversely correlated to depression scale1
- Women who took the multivitamin and zinc showed a significant reduction in anger-hostility and depression-dejection versus women who did not2
- Patients who received 25mg of Zinc daily had significantly reduced HDRS and BDI scores compared to placebo group3



¹Amani R et al. *Biol Trace Elem Res.* 2010; 137(2):150-8.

²Sawada T et al. *Eur J Clin Nutr* 2010;64(3):331-3.

³Nowak et Al. *Pol J Pharmacol* 2003; 55:1143-1147.

Zinc Supplementation for Depression



- 12 week study with 60 patients with unipolar depression
- 25 mg Zinc daily or placebo
- Zinc supplementation significantly reduced depression scores in antidepressant treatment resistant patients

Siwek, et al., *J Affect Disord*. 2009 Nov;118(1-3):187-95.



Zinc in Depression: A Meta-Analysis



- Seventeen studies, measuring peripheral blood zinc concentrations in 1643 depressed and 804 control subjects, were included
 - Zinc concentrations were lower in depressed subjects than control subjects
- Depression is associated with a lower concentration of zinc in peripheral blood

Swardfager W et al. Biol Psychiatry 2013 Jun. [E-pub ahead of print.]



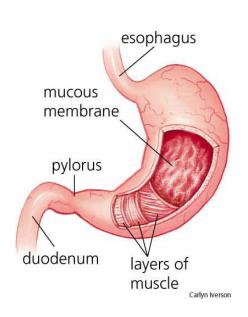
Zinc and Digestive Enzymes



• Zinc increases the activity of digestive enzymes



- Zinc deficiency influences ability to form hydrochloric acid (HCl)
- Zinc deficiency causes insufficient gastric acid production





Zinc Dependent Enzymes



Enzyme	Digests
Trypsin	Protein
Chymotrypsin	Protein
Elastase	Protein
Carboxypeptidase	Protein
Lipase	Fat
Amylase	Polysaccharides
Maltase	Maltose
Sucrase	Sucrose
Lactase	Lactose
Pepsin	Protein



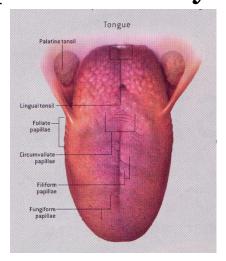
Decreased Taste and Smell

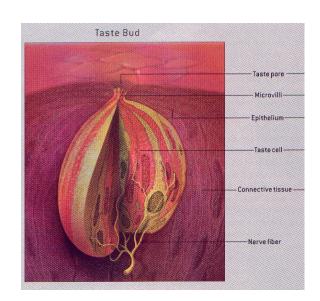


Zinc deficiency alters taste and smell receptors

Taste is mediated through a salivary zinc

dependent enzyme

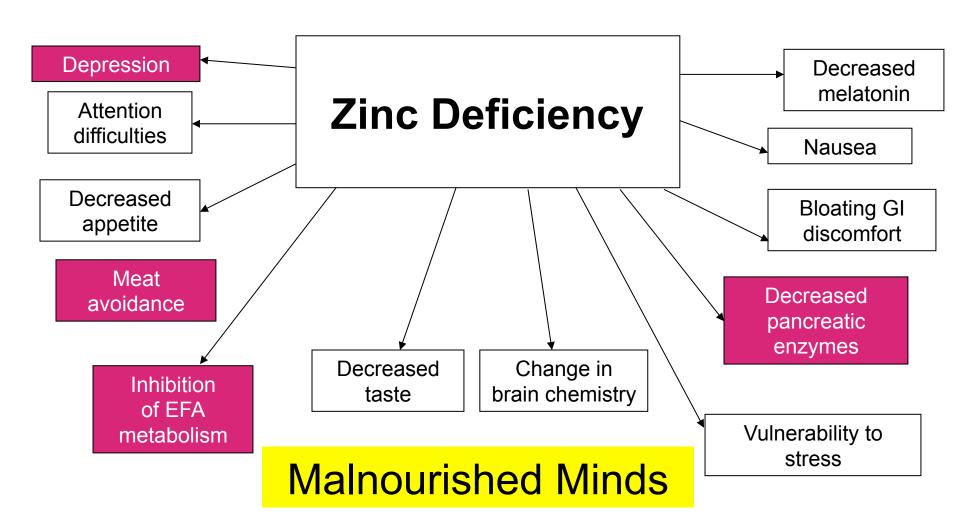




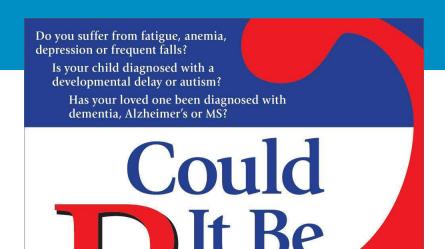


Zinc Deficiency









Please Note: Although the reference range for vitamin B12 is 200-1100 pg/mL, it has been reported that between 5 and 10% of patients with values between 200 and 400 pg/mL may experience neuropsychiatric and hematologic abnormalities due to occult B12 deficiency; less than 1% of patients with values above 400 pg/mL will have symptoms.



The underground classic that has saved lives



SALLY M. PACHOLOK, R.N., B.S.N. JEFFREY J. STUART, D.O.





Symptoms of B₁₂ Deficiency



MENTAL

- Irritability
- Apathy
- Personality Changes
- Depression
- Memory Loss
- Dementia
- Hallucinations
- Violent Behavior

PHYSICAL

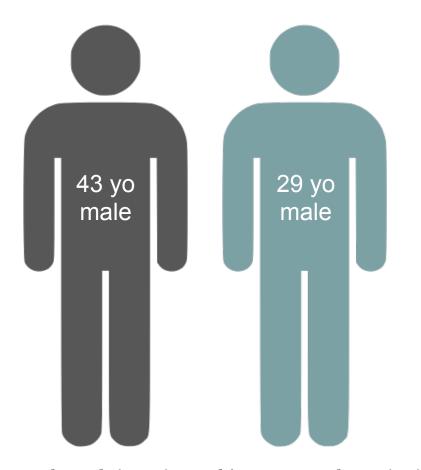
- Clumsiness
- Weakness
- Pernicious Anemia
- Chronic Fatigue
- Tremors
- G.I. Problems
- Diminished sense of touch and pain

BIOCHEMICAL INDIVIDUALITY



B12 for Treatment Resistant Depression





- Both males failed to respond to 3+ trials of antidepressants
- Vegetarians
- Low vitamin B12 levels
- Treatment with 1000 mg/ day of Vitamin B12
- Both noticed improvements within 3-4 weeks

Kale et al. (2009). Psychiatry Research. 175(1-2):47-53.



Lithium: Mineral, Medicine, Miracle







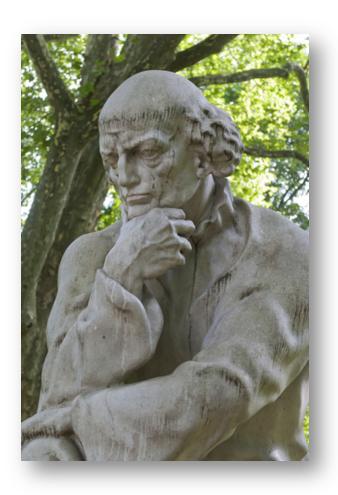
Dosage



• "The dosage makes it either a poison or a remedy."

- Paracelsus

Swiss-German physician, physician, botanist, alchemist, astrologer (1493-1541)





Nutritional Lithium and Suicide



- Examined 27 Texas counties from 1978-1987
- Examined lithium levels in tap water in the 18 municipalities in Japan in relation to the suicide standardized mortality ratio
- Austrian study with nationwide sample of 6460 lithium measurements

Overall suicide rate and the suicide mortality ratio were inversely associated with lithium levels

Ohgami H, et al. (2009). *Br J Psychiatry*, 194(5), 464-5. Schrauzer GN, Shrestha KP. (1990). *Biol Trace Elem Res*, 25(2),105-13. Kapusta, N.D., et al. (2011). *The British Journal of Psychiatry*, 198, 346-350.



Lithium Cases



- Irritability
- Road rage; waiting room rage
- Family history of mood and substance abuse
- Domestic violence

PATRICIA

43 year old therapist

Diagnosed with depression and alcohol abuse at age 18

Strong family
history of
alcoholism
Sober for 10 years



Lithium for Prevention of Alzheimer's



- Comparison of 66 elderly patients with Bipolar Disorder who were on chronic lithium therapy and 48 similar patients without recent lithium therapy
- Alzheimer's disease was diagnosed in 3 patients (5%) on lithium and in 16 patients (33%) who were not on lithium (P<0.001)
- Lithium treatment reduced the prevalence of Alzheimer's disease in patients with bipolar disorder to levels in the general elderly population

Nunes PV et al. *Br J Psychiatry*. 2007 Apr;190:359-60.



Lithium and Cognitive Decline



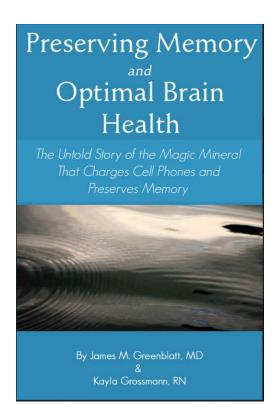
- Lithium modulates several biological cascades related to dementia/Alzheimer's
- Modulating these cascades can decrease amyloid and tau-induced neurotoxicity and cell damage

Can lithium reduce risk and/or progression from Alzheimer's?



Lithium





LowDoseLithium.org

Contents

I. Restoring Brain Health

Ch. 1 Something is Wrong

Ch. 2 Nutritional Psychiatry

Ch. 3 From Prescription Pad to Health Food Store

II. Lithium as a Mineral

Ch. 4 Lithium as a Mineral

III. Lithium as a Medicine

Ch. 5 Lithium as a Medicine: Depression

Ch. 6 Lithium as a Medicine: Substance Abuse

Ch. 7 Lithium as a Medicine: ADHD

Ch. 8 Lithium as a Medicine: Anger and Aggression

Ch. 9 Lithium as a Medicine: Dementia and Alzheimer's

IV. Lithium as a Miracle

Ch. 10 Lithium as a Miracle: Neuroprotection and Neurogenesis

Ch. 11 Playing God

V. Conclusion

Ch. 12 Peace of Mind

VI. Appendix - Resources and References



Spirituality



- Studies have concluded that people who have faith in religion:
 - cope better with stress
 - suffer from less depression
 - respond better to medical treatment

Murphy PE et al. *Journal of Clinical Psychology* 2009; 65 (9):1000-1008. Schettino JR et al. *Mental Health, Religion & Culture* 2011; 14 (8):805-818. Vasegh, S. *Journal of Cognitive Psychotherapy: An International Quarterly* 2011; 25 (3):177-188.



Nutrition and Behavior



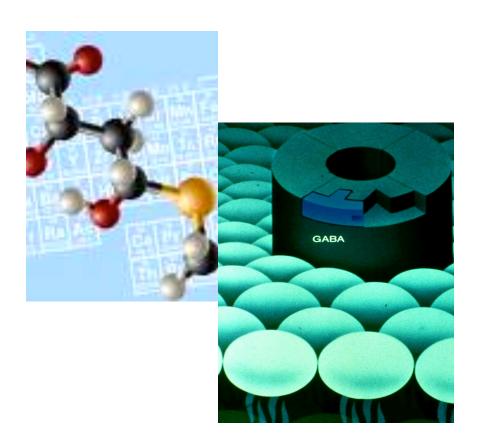
- Double-blind, placebo-controlled trial of nutritional supplements on 231 young adult prisoners, comparing disciplinary offences before and during supplementation for appx. 4 months
- Those receiving supplements committed an average of **26.3% fewer offences** (n=172)
- Those taking supplements for a minimum of 2 weeks experienced an average of **35.1% reduction** in offenses whereas placebos remained within standard error

Gesch, C.B., et al. British Journal of Psychiatry 2002;181:22-28.



Nutrition and Health

Understanding the role of Nutrition and Health is not Alternative Medicine





Integrative Therapies



- Folate/B12
- Thyroid
- Vitamin D
- Zinc/Magnesium
- Copper
- Celiac Disease
- Homocysteine
- Infections
- Cholesterol
- DHEA

- Food Allergies
- Vitamin Deficiencies
- Mineral Deficiencies
 - LITHIUM
- Amino Acids
- Heavy Metals
- Fatty Acids
- Toxins



An Integrative Approach to Mental Health



- The genetic and biochemical foundations that may predispose to symptoms
- 2. The dietary habits, pre-symptom history and present
- 3. The impact of the illness on social and psychological systems
- 4. The physical and social environment in which the symptoms occur
- 5. Understanding the patients experience and beliefs of his/her illness



Times Are Changing



Lancet Psychiatry January 26, 2015

Nutritional medicine as mainstream in psychiatry

Jerome Sarris, Alan C Logan, Tasnime N Akbaraly, G Paul Amminger, Vicent Balanzá-Martínez, Marlene P Freeman, Joseph Hibbeln, Yutaka Matsuoka, David Mischoulon, Tetsuya Mizoue, Akiko Nanri, Daisuke Nishi, Drew Ramsey, Julia J Rucklidge, Almudena Sanchez-Villegas, Andrew Scholey, Kuan-Pin Su, Felice N Jacka, on behalf of The International Society for Nutritional Psychiatry Research

Clinical Psychological Science 2015

Theoretical/Methodological/Review Article

The Emerging Field of Nutritional Mental Health: Inflammation, the Microbiome, Oxidative Stress, and Mitochondrial Function

ASSOCIATION FOR PSYCHOLOGICAL SCIENCE

Clinical Psychological Science 1-17 © The Author(s) 2015 Reprints and permissions: sagepub.com/journalsPermissions.nav DOI: 10.1177/2167702614555413 cpx.sagepub.com

\$SAGE

Bonnie J. Kaplan¹, Julia J. Rucklidge², Amy Romijn², and Kevin McLeod³



Integrative Psychiatry





Mental Illness may be seen as a reflection of multiple errors in physiology.

If we find the Causes, we may find the Cures.

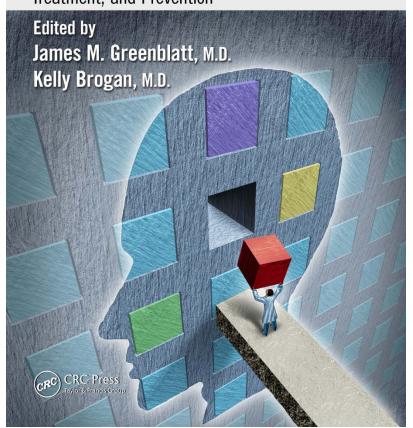
Prevention is Possible.





INTEGRATIVE THERAPIES FOR DEPRESSION

Redefining Models for Assessment, Treatment, and Prevention



DEPRESSION Solution

A *Personalized* 9-Step Method for Beating the *Physical* Causes of Your Depression



James Greenblatt, MD



Thank you!



James Greenblatt, M.D. Chief Medical Officer

http://www.waldenbehavioralcare.com/

http://jamesgreenblattmd.com/

http://www.lowdoselithium.org/

