## Shawn M. Talbott, PhD

CNS, LDN, FACSM, FAIS, FACN
Chief Science Officer





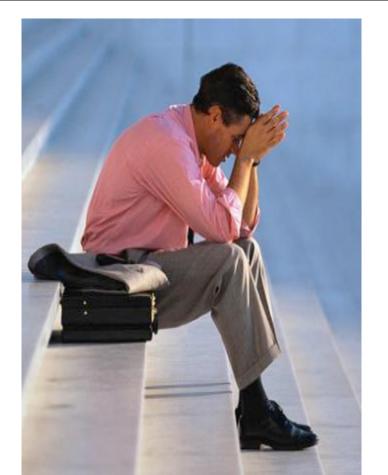
## Causes of "Imbalance" (Stress)

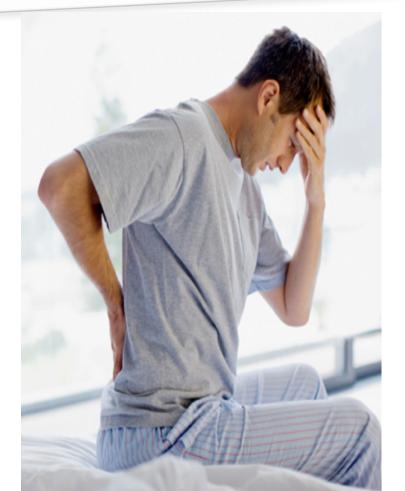
- Emotional stress (deadlines, bills, traffic...)
- Physical stress (aging, sleep deprivation, exercise...)
- Environmental stress (air/water pollution, heat, cold...)
- Non-Optimal Diet (processed foods, inadequate nutrients/phytonutrients...)

- Athletes / Dieters / Short-Sleepers / Stressed
  - Share the SAME biochemical disruptions
  - Share the SAME psychological outcomes
  - Exhibit the SAME benefits to restored biochemical balance















Prana Zone Mood Mana Swing Energy Focus Vigor Edge Flow Motivation Runner's High

## Vigor

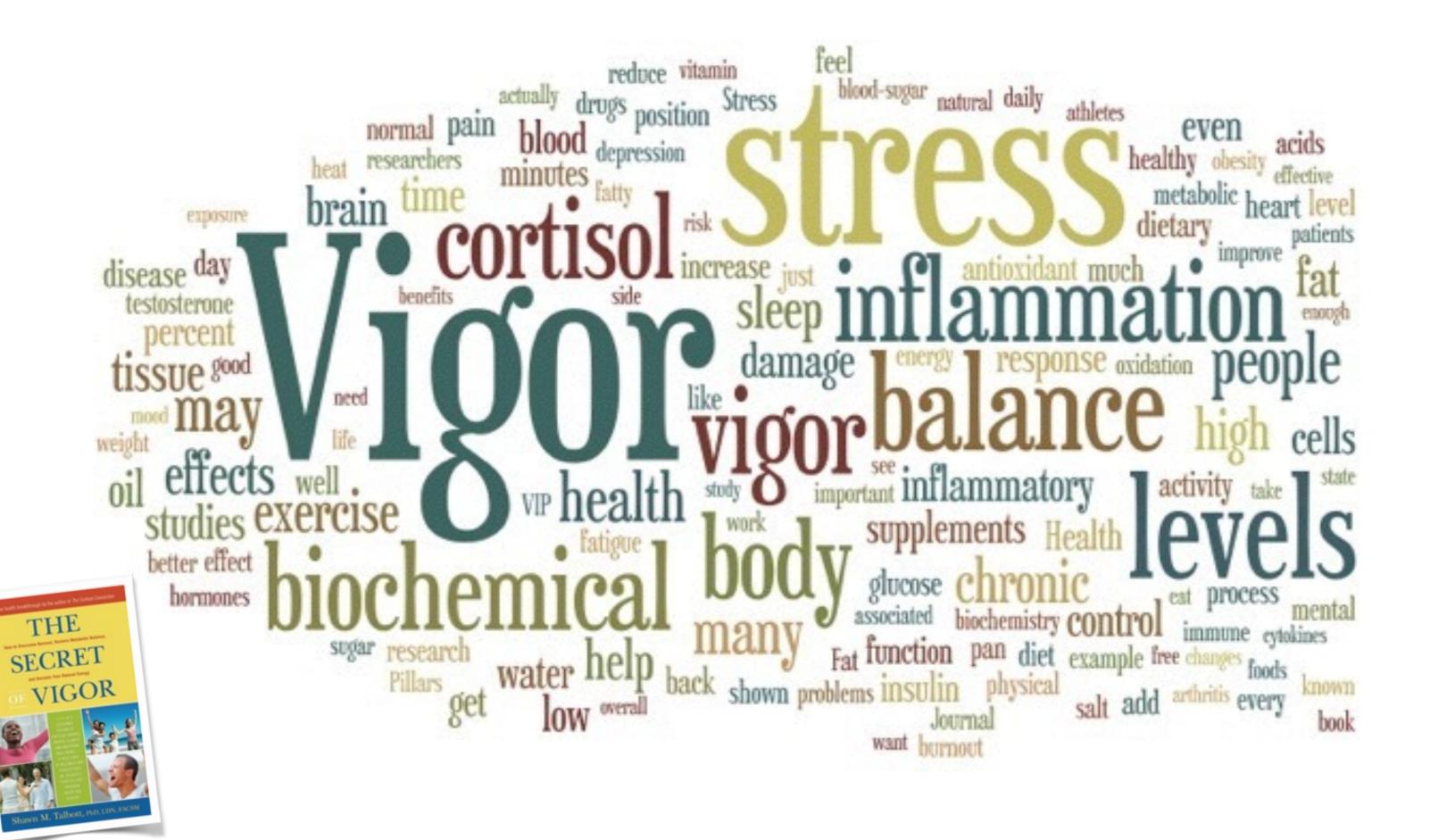
3-tiered mood state... characterized by:

Physical Energy

Mental Acuity

**Emotional Well-Being** 

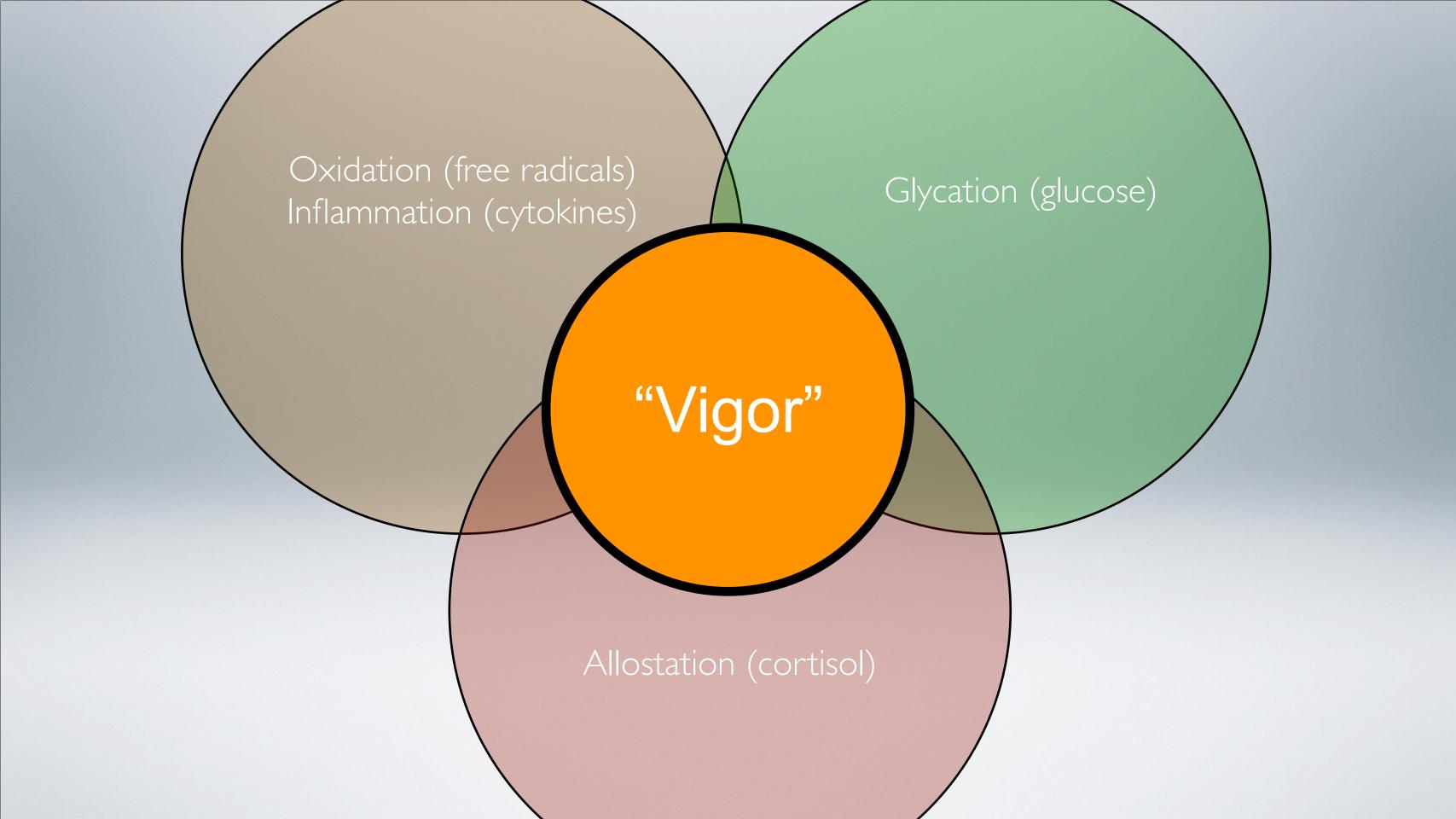




## What does

# Energy mean to





## Just an Antioxidant?



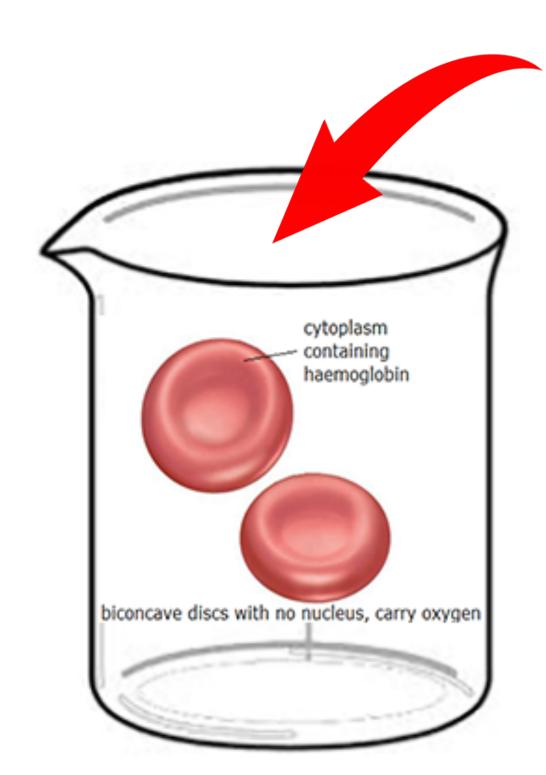






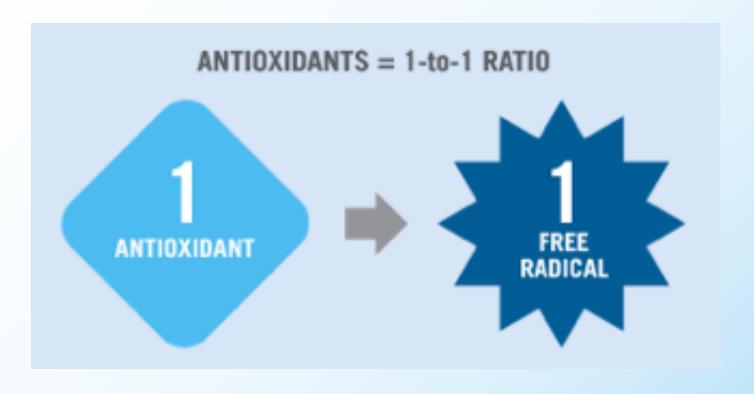


## CAP-e = Cell-based Antioxidant Protection in Erythrocytes ORAC = Oxygen Radical Absorbance Capacity

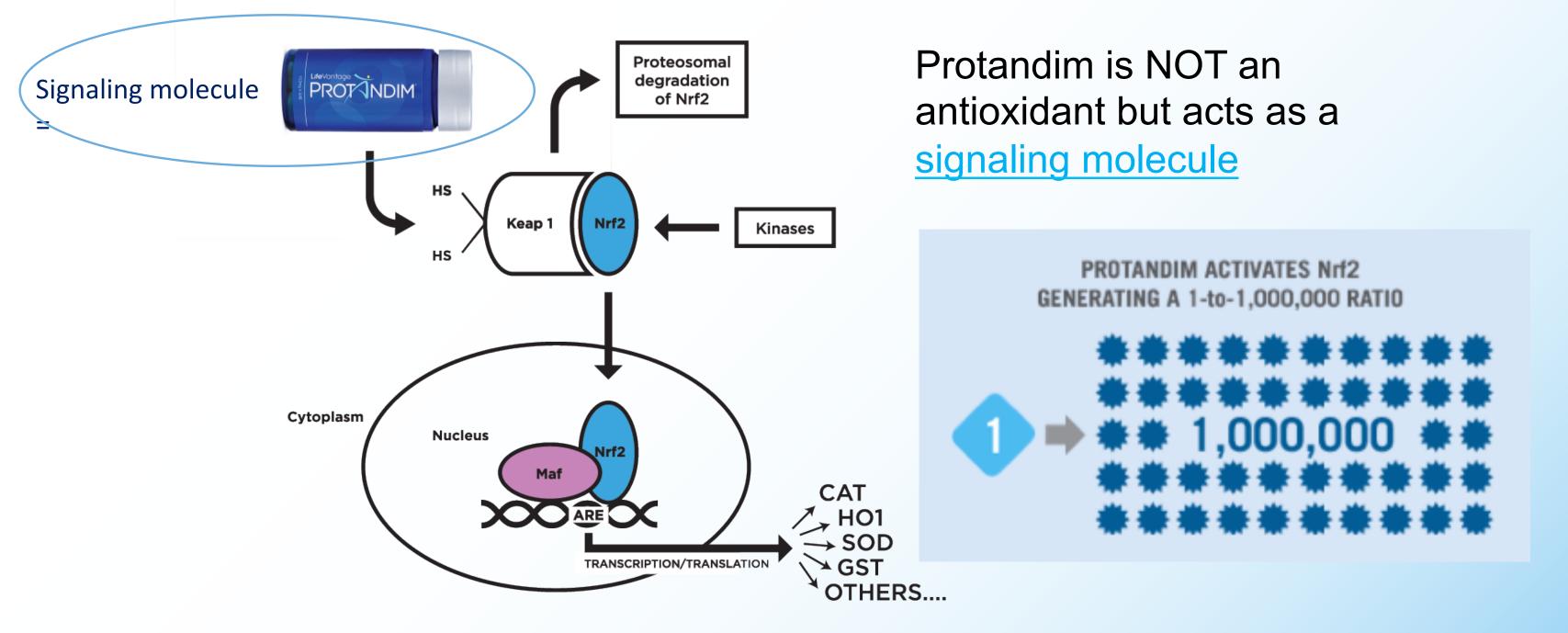


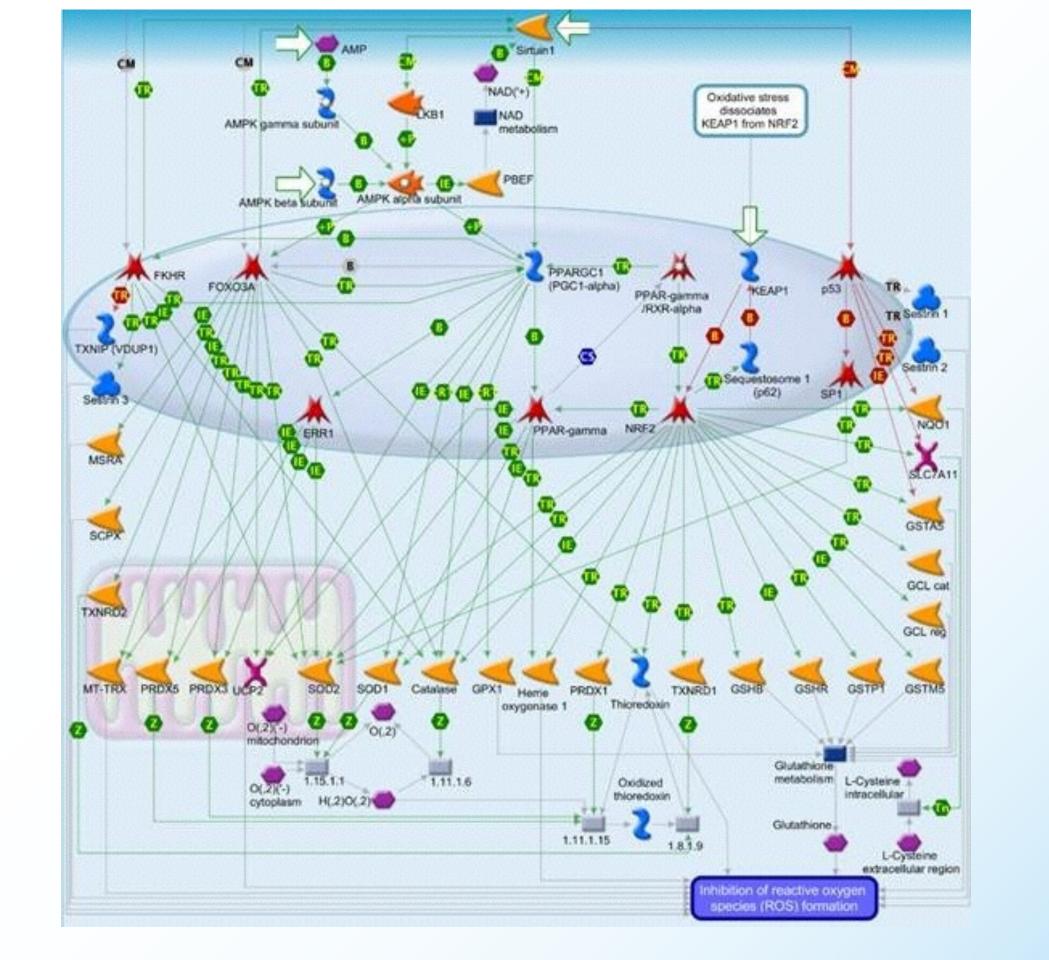
Add anti-oxidants to the RBC in beaker

- Let the anti-oxidant diffuse into the RBC
- Measure how oxidants are neutralized
- OLD TECHNOLOGY

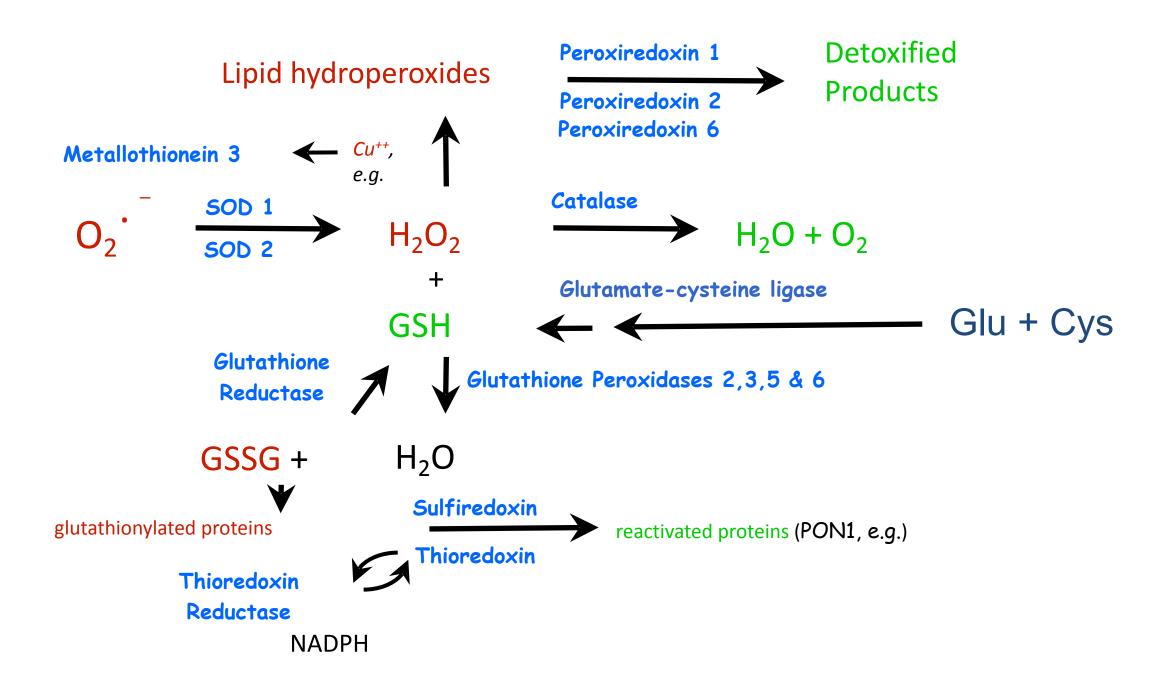


## Protandim is the New Way to deal with oxidative stress via Nrf2 pathway and expression of anti-oxidant enzymes





## The Internal System of Protective Antioxidant Enzymes Activated/Triggered by Nr2 Pathway



Free

journal home

### **Heart Failure**

Chronic Pulmonary Artery Pressure Elevation Is Insufficient to Explain Right Heart Failure

Harm J. Bornard, MD. PhD\*: Ramoh Natarajan, PhD\*: Scott C. Honderson, PhD

function, by of data on I

Ke Original Contribution

Protandim attenuates intimal ex vivo via a catalase-depend

Binata Joddar a,b,c, Rashmeet K, Reen b Jay L. Zweier b, Keith J. Gooch a.b.\*

- \* Department of Biomedical Engineering, The Ohio State Univers Davis Heart & Lung Research Institute. The Ohio State Univer-
- <sup>6</sup> BIKEN Nanomedical Engineering Laboratory, Wako-shi, Saitan
- Department of Surgery, The Ohio State University, Columbus, 6 Department of Cardiothoracic Surgery, The Ohio State Univers
- Division of Pulmonary and Critical Care Medicine, Department

### ARTICLE INFO

Received 26 August 2010 Revised 7 December 2010 Accepted 8 December 2010 Available online 15 December 2010

-3.6-fold increase Keywords. Protandim, a nuti Free radicals enzymes in severa Scavenging enzymes isolated HSV. Prof Catalase Human suphenous veins respectively, and o Ex vivo culture Protandin and proliferation.

Billion I Baharil Code Back CO.

### United States Patent Myhill et al.

- (56) COMPOSITIONS FOR ALLEVIATING INFLAMMATION AND OXIDATIVE STRESS
- (25) Inventors: Paul R. Myhill, Castle Rock, CO (US): William J. Driscoll, Englewood, CO.
- (23) Assignor: Lifeline Nutracenticals Corporation Englewood, CO (US)
- Subject to any disclaimer, the term of this (\*) Notice:

(10) Patent No.: (45) Date of Patent: Al-Shart, "C-Reactive Protein and

Hum

endogenous antio

model of HSV IH.

American F Learn

The Dietary Suppleme

Plasma Osteopontin a

Muhammad Mudda

Oxidative Stress in Mus

OPEN & ACCESS Freely available online

Associa Protandim, a Fundamentally New Antioxidant Approach in Chemoprevention Using Mouse Two-Stage Skin Carcinogenesis as a Model

Jianfeng Liu', Xin Gu', Delira Robbins', Guohong Li', Runhua Shi', Joe M. McCord', Yunfeng Zhao'\*

Eppartment of Pharmacology, Touloslogy & Neuroscience, Louisiana State University Health Sciences Certer, Streveport, Louisiana, United States of America 2 Department of Pathology, Louisiana State University Health Sciences Center, Shreveport, Louisiana, United States of America, 3 Department of Neurosurgery, Louisiana State University Health Sciences Center, Shreveport, Louisiana, United States of America, 4 Felst-Wellier Cancer Center, Louisiana State University Health Sciences Cente Symptom, Lautiana, Linked States of America, \$-Copportment of Medicine, University of Colorado Health Sciences Carter, Carter, Carter, Colorado, United States of America

### Abstract

Warren C. McClure, MS

Nicole L. Arevalo, MA

Rick E. Rabon, BA

Benjamin Mohr

Swapan K. Bose, BS, BPharm

Joe M. McCord, PhD

Brian S. Tseng, MD, PhD

patency. To evaluate the role of reactive oxygen species (ROS) signaling in intima hyperplasia (IH), an early

stage pathology of vein-graft disease, and to explore the potential therapeutic effects of up-regulating

OPEN BACCESS Freely available on line

Oxidative stress is an important contributor to cancer development. Consistent with that, antioxidant enzymes have been demonstrated to suppress tumorigenesis when being elevated both in vitro and in vivo, making induction of these enzymes a more potent approach for cancer prevention. Protandim, a well-defined combination of widely studied medicinal plants has been shown to induce superoxide dismutase (SOD) and catalase activities and reduce superoxide generation and lipid peroxidation in healthy human subjects. To investigate whether Protandim can suppress tumor formation by a dietary approach, a two-stage mouse skin carcinogenesis study was performed. At the end of the study, the mice on a Protandimcontaining basal diet had similar body weight compared with those on the basal diet, which indicated no overt toxicity by Protandim. After three weeks on the diets, there was a significant increase in the expression levels of SOD and catalase, in addition to the increases in SOD activities. Importantly, at the end of the carcinogenesis study, both skin tumor incidence and multiplicity were reduced in the mice on the Protandim diet by 33% and 57% respectively, compared with those on basal diet. Biochemical and histological studies revealed that the Protandim diet suppressed tumor promoter-induced oxidative stress (evidenced by reduction of protein carbonyl levels), cell proliferation (evidenced by reduction of skir hyperplasia and suppression of PKC/INK/Jun pathway), and inflammation (evidenced by reduction of ICAM-1/VCAM-1 expression, NF-x8 binding activity, and nuclear p65/p50 levels). Overall, induction of antioxidant enzymes by Protandim may serve as a practical and potent approach for cancer prevention.

ournal of the American College of Cardiology O 2004 by the American College of Cardiology Foundation

Serum Levels of Thiobarbituric Acid Reactive Substances Predict Cardiovascular

Beverly and Boston, Massachusetts; New York, New York; and Groton, Connecticut



Contents lists available at ScienceDirect

### Free Radical Biology & Medicine

journal homepage: www.elsevier.com/locate/freeradbiomed



Original Contribution

Synergistic induction of heme oxygenase-1 by the components of the antioxidant supplement Protandim

Kalpana Velmurugan a,b, Jawed Alam c, Joe M. McCord d, Subbiah Pugazhenthi a,b,\*

ELSEVIER

- Division of Endocrinology, Department of Medicine
- Section of Endocrinology, Veterans Affairs Medical tment of Molecular Genetics, Ochsner Medica
- Division of Pulmonary Sciences, Department of Me



Article history: Received 8 July 2008 Revised 23 September 2008 Accepted 31 October 2008 Available online 17 November 2008

Keywords: Oxidative stress Heme orggenase-Synergy Curcumin Protandim

ESSN 07

Available online at www.sciencedirect.com



Free Radical Biology & Medicine 40 (2006) 341 - 347



Original Contribution

The induction of human superoxide dismutase and catalase in vivo: A fundamentally new approach to antioxidant therapy

Sally K. Nelson a,b, Swapan K. Bose a, Gary K. Grunwald C, Paul Myhill d, Joe M. McCord a,b,d,\*

4 Lifeline Therapeutics, Dancer, CO, USA

\* Webb-Waring Institute for Cancer, Aging and Antioxidant Research, University of Colorado Denver Health Sciences Center, Denver, CO 89262, USA Department of Medicine, University of Colorado Donver Health Scionces Center, Donver, CO 80262, USA Department of Preventive Medicine and Biometrics, University of Colorado Denver Health Sciences Center, Denver, CO 80262, USA

Received 22 June 2005; revised 24 August 2005; accepted 28 August 2005

A composition consisting of extracts of five widely studied medicinal plants (Protandim) was administered to healthy human subjects ranging in age from 20 to 78 years. Individual ingredients were selected on the basis of published findings of induction of superoxide dismutase (SOD) and/or catalase in rodents in vivo, combined with evidence of decreasing lipid peroxidation. Each ingredient was present at a dosage sufficiently low to avoid any accompanying unwanted pharmacological effects. Blood was analyzed before supplementation and after 30 and 120 days of supplementation (675 mg/day). Frythrocytes were assayed for SOD and catalase, and plasma was assayed for lipid peroxidation products as thiobarbituric acid-reacting substances (TBARS), as well as uric acid, C-reactive protein, and cholesterol (total, LDL, and HDL). Before

### Events in Patients With Stable Coronary Artery Disease

ABSTRACT. Therapeutic options for Duchenne muscular dystrophy (DMD), t A Longitudinal Analysis of the PREVENT Study

common and lethal neuromuscular disorder in children, remain elusive. O Mary F. Walter, PhD,\* Robert F. Jacob, PhD,\* Barrett Jeffers, PhD,\* Mathieu M. Ghadanfar, M. Chadanfar, M. Chada Gregory M. Preston, PHD, Jan Buch, MD, R. Preston Mason, PHD + an over-the-counter supplement with the ability to induce antioxidant enzymes

PLoS one

### The Chemopreventive Effects of Protandim: Modulation of p53 Mitochondrial Translocation and Apoptosis catalase activity by during Skin Carcinogenesis

cultured HSV and Delira Robbins<sup>1</sup>, Xin Gu<sup>2</sup>, Runhua Shi<sup>6</sup>, Jianfeng Liu<sup>1</sup>, Fei Wang<sup>3</sup>, Jacqulyne Ponville<sup>4</sup>, Joe M. McCord<sup>5</sup>, Yunfeng Zhao1\*

> 1 Department of Pharmacology, Toxicology and Neuroscience, Louisiana State University Health Sciences Center, Shreveport, Louisiana, United States of America, 2Department of Pathology, Louisiana State University Health Sciences Center, Streveport, Louisiana, United States of America, 3 College of Life Science, Jilin University, Changchun, Jilin Province, China, 4 Department of Chemistry, Nicholls State University, Thibodeux, Louisiena, United States of America, 5 Department of Medicin University of Colorado at Donver and Health Sciences Center, Aurora, Colorado, United States of America, & Feist Weiller Cancer Conter, Louisiana State University Health on Center, Showeport, Louisiana, United States of America

Protandim, a well defined dietary combination of 5 well-established medicinal plants, is known to induce endogen antioxidant enzymes, such as manganese superoxide dismutase (Mn500). Our previous studies have shown through the induction of various antioxidant enzymes, products of oxidative damage can be decreased. In addition, we have shown that tumor multiplicity and incidence can be decreased through the dietary administration of Protandim in the two-stage skin carcinogenesis mouse model. It has been demonstrated that cell proliferation is accommodated by cell death during DMBA/ TPA treatment in the two-stage skin carcinogenesis model. Therefore, we investigated the effects of the Protandim diet on apoptosis; and proposed a novel mechanism of chemoprevention utilized by the Protandim dietary combination. Interestingly, Protandim suppressed DMBA/TPA induced cutaneous apoptosis. Recently, more attention has been focused on transcription-independent mechanisms of the tumor suppressor, p53, that mediate apoptosis. It is known that cytoplasmic p53 rapidly translocates to the mitochondria in response to pro-apoptotic stress. Our results showed that used the mitochondrial translocation of p53 and mitochondrial outer membrane proteins such as Bax. We examined the levels of p53 and MnSOD expression/activity in murine skin JB6 promotion sensitive (P+) and promotion resistant (P-) epidermal cells. Interestingly, pS3 was induced only in P+ cells, not P- cells; whereas MnSOD is highly expressed in P- cells when compared to P+ cells. In addition, wild-type p53 was transfected into J86 P- cells. We found that the introduction of wild type p53 promoted transformation in JB6 P- cells. Our results suggest that suppression of p53 and induction of MrSOD may play an important role in the tumor suppressive activity of Protandim



Contents lists available at SciVerse ScienceDirect

### Molecular Aspects of Medicine

journal homepage: www.elsevier.com/locate/mam



### Oxidative Stress in Health and Disease: The Therapeutic Potential of Nrf2 Activation

Brooks M. Hybertson, a,b Bifeng Gao, a Swapan K. Bose and Joe M. McCorda,b

Department of Medicine, Division of Pulmonary Science and Critical Care Medicine, University of Colorado at Denver, Aurora, CO 80045 \*LifeVantage Corporation, 10813 S. Riverfront Parkway, South Jordan, UT 84095



Anderson, et al., "Differential Rose ellis to Induction of Apoptosis by E Analogue, or TEA," Conver Res. Baker, et al., "Reduced RBC Very Due to Endotoxin," Circul, Shock, bosa, et al., "Decreased Oxidat tion, 2003; vol. 19: pp. 837-842. Buttacharya, et al., "Astioxidant

2004; vol. 291, No. 23; pp. 2818-2

from Withania somisfire," Ind. J. Esper. Biol., 1997; vol. 35: pp.

Frank L. Meyskens Jr., MD Laurence H. Baker, DO DETIME RISK OF PROST

cer in the United Sta rently estimated to be though most cases an an early, curable stage, tre costly and urinary, sexual, a related adverse effects are o Even men who choose activ lance as an initial managem egy face anxiety, uncertain p and a measurable risk of sepsi low-up biopsies,3 and more third of those who initially del are ultimately treated.43 Wi

Author Video Interview ava www.jama.com.

02011 American Medical Associa

Downloaded From: http://jama.jamanetwork.c

THE NEW ENGLAND JOURNAL OF MEDICINE

### EFFECTS OF A COMBINATION OF BETA CAROTENE AND VITAMIN A CARDIOVASCULAR DISEASE

GILBERT S. OMENS, M.D., PH.D., GARY E. GOODMAN, M.D., M.S., MARK JOHN BALMES, M.D., MARK R. CULLEN, M.D., ANDREW GLASS, M.D., JA FRANK L. MEYSKENS, JR., M.D., BARBARA VALANIS, DR.P.H., JAMES H. SCOTT BARNHART, M.D., M.P.H., AND SAMUEL HAMMAR,

Abstract Background. Lung cancer and cardiovascular disease are major causes of death in the United States. It has been proposed that carotenoids and retinoids are agents that may prevent these disorders.

Methods. We conducted a multicenter, randomized, double-blind, placebo-controlled primary prevention trial - the Beta-Carotene and Retinol Efficacy Trial - involving a total of 18,314 smokers, former smokers, and workers exposed to asbestos. The effects of a combination of 30 mg of beta carotene per day and 25,000 IU of retinol (vitamin A) in the form of retinyl palmitate per day on the primary end point, the incidence of lung cancer, were compared with those of placebo.

Results. A total of 388 new cases of lung cancer were diagnosed during the 73,135 person-years of followup (mean length of follow-up, 4.0 years). The active-treatment group had a relative risk of lung cancer of 1.28 (95 percent confidence interval, 1.04 to 1.57; P=0.02), as

UNG cancer is the leading cause of death from cancer in the United States, accounting for approximately 29 percent of deaths from cancer and 6 percent of all deaths. New approaches are essential to prevent lung cancer in persons who have smoked cigarettes or who have had occupational exposure to asbestos. Twenty-nine percent of men and 25 percent of women who are 45 to 64 years of age currently smoke,2 and at least 40 percent of men and 20 percent of women in this age group are former smokers.3 An estimated 4000 to 6000 deaths from lung cancer per year are attributed to ex-

On the basis of epidemiologic observations and laboratory studies, beta carotene and vitamin A have attracted wide interest as agents that may prevent lung cancer.<sup>63</sup> The Beta-Carotene and Retinol Efficacy Trial (CARET) is one of several recent trials to assess the

posure to asbestos. (1)

From the Division of Public Idealth Sciences, Fred Hundstrom Cancer Re-search Center, Seattle (G.S.O., G.E.G., M.D.T., S.B.); the Departments of Envi-connected Health and Medicine, University of Washington, Scattle (G.S.O., G.E.G., S.B., S.H.;; the Swedish Hospital Tumor Institute, Seattle (G.E.G.); the unt of Medicine, University of California at San Francisco, San Fran ciaco (J.R.); the Department of Medicine, Vale University, New Horen, Conn. (M.R.C.); Kaiser Permanento-Center for Health Research, Portland, Oreg. (A.G., B.V.); the Department of Medicine, University of Maryland, Baltimore (J.P.K.); and the Department of Medicine and Cancer Center, University of California at

data for Department of Indicated and Canter California, Canterland of Cambridge of Canterland Parkins, Crange (FE.M., 119%). Address regretal response to Dr. Chemin at the Fixed Postchiones. Cancer Research Center, Division of Public Illeath Sciences, 1124 Callination. AMPRID, Scalads, MA. 981005.

Supported by grants (URL CAMSTO, URL CAASSEL, URL CAAF989, URL CAAF980, URL CAAFS0, URL CAAFS0, CARSON, URL CAAFS0, TO CARSON, CAR

\*Other contributing authors were Carl Andrew Brookin, M.D. (University of Workington, Seattle, Martin G. Chevniack, M.D. (Tale University, New Haves, Conn.), James E. Grizzle, Ph.D. (Fod Hinkshimon Catzer Research Center, Seattle), Marjorie Perloff, M.D. (National Cancer Institute, Bethesda, Md.), and Linds Rosenstock, M.D., M.P.R. (University of Washington, South):

compared with the places tically significant different cancer. In the active-tree death from any cause w interval, 1.03 to 1.331; of c percent confidence interv from cardiovascular disea interval, 0.99 to 1.61). On randomized trial was st planned: follow-up will co

Conclusions. After an plementation, the combin min A had no benefit and on the incidence of lung. from lung cancer, cardiov in smokers and workers Med 1996;334:1150-5.)

chemopreventive efficac and related agents. (6-1)

This report presents CARET study, which coi of the steering committee to stop the trial's active ditional end points is exfive years.

### Study Design

The study's strategy, design where. 1044 Briefly, CARET w domination in Scattle in 1985 randomly assigned in a 1:1 m 15 mg of beta carotene per day tive treatment) or placebu; the carotene per day, 25,000 IU of additional study centers in 190

domly assigned in a 1:1 ratio The pilot groups receiving active agents were consolid into a single group receiving a standard daily regimen of 3 beta carotene plus 25,000 IU of retinol in the form of retin se. Thus, in the pilot study with the cohort of smokers, t jects were assigned to active treatment for every subject as scebo; therefore, the rates rather than numbers of end po be compared between active and placebo groups. The desig for active intervention until late 1997 (110,000 person-years) porting of results in 1998.

### Eligibility, Recruitment, and Randomization

Workers exposed to asbestos were mon 45 to 74 years the pilot study and 45 to 69 years of age in the later per

α-Tocopherol and β-Carotene Supplements and Lung Cancer Incidence in the Alpha-Tocopherol, Beta-Carotene Cancer Prevention Study: Effects of Base-line Characteristics and Study Compliance

Demetrius Albanes, Olli P. Heinonen, Philip R. Taylor, Jarmo Virtamo, Brenda K. Edwards, Matti Rautalahti, Anne M. Hartman, Juni Palmgren, Laurence S. Freedman, Jaason Haapakoski, Michael J. Barrett, Pirjo Pietinen, Nea Malila, Eero Tala, Kari Liippo, Eija-Riitta Salomaa, Joseph A. Tangrea, Lyly Teppo, Frederic B. Askin, Eero Taskinen, Yener Erozan, Peter Greenwald, Jussi K. Huttunen\*

> The Journal of Nutrition **Nutritional Epidemiology**

### **Antioxidant Supplementation Increases the** Risk of Skin Cancers in Women but Not in Men<sup>1</sup>

Serge Hercherg, 2,74 Khaled Ezzedine, 2,4 Christiane Guinot, 3,6 Paul Preziosi, 2 Pilar Galan, 2 Sandrine Bertrais,2 Carla Estaquio,2 Serge Briançon,7 Alain Favier,8 Julie Latreille,1 and Denis Malvy9

UMR USST Inserm/U1125 Insa/EA3200 Cnam/Univ Paris 13, Bobigny, France 93017; \*Unité de Surveillance et d'Epidémiologie Natritioneelle, Centre de Rocherche en Natrition Humaine IIe-de-France UFR SMBH Paris 13, Bobigny, France 93017; \*Department of Dermanology, University Hospital Erasme, Université Libre de Bruselles, Bruselles, Belgium 1070; \*Biometrics and Epidemiology Unic, CER.LE.S., Neully au Seine, France 92521, \*Computer Science Laboratory, Ecole Polytechnique, Université de Toum, Tours, France 37200, \*EA 3444, Ecole de Santé Publique, Epidémiologie elinique, Faculté de Médecine, CHU Nancy, France 54035, \*Laboratoire Listons des Acides Nuclitiques, UMR CNRS-CEA-UJF 5046, Grenoble, Franco 38000; and \*EA 3677 and Centre Reni-Labusquière (Tropical Medicine and International Health Brunchi, Université Victor Segalon Bondraux 2 and Department of Intornal Medicine and Eropical Diseases, University Hospital Centes, Bordeson, France 33076

Background

suggest that

of vitamin E

other foods

precursor o

dark-green,

the risk of o

ings of th

Prevention

lung cancer

who receive

were recent

ficacy Trial

carotene an

of a-tocoph

cidence of I

the ATBC

age, number

status, and

in relation t

tologic type

whether the

could facilit

Study result

tion and r

total of 29

or more o

receive a-1

tocopherol

(median, 6.1

study entry.

tocopherol i

and death of

pendently o

able for 9

evaluated b

hazards mo

1560 ARTICLES

= 894) were

factors for 1

This research aimed to test whether supplementation with a combination of anticuidant ultamins and minerals could reduce the risk of skin cencers (SCI, it was performed within the framework of the Supplementation in Vitamins and Mineral Antioxidants study, a randomized, double-blinded, placebo-controlled, primary prevention trial testing the efficacy of nutritional doses of antioxidents in reducing incidence of cancer and ischemic heart disease in the general population. French adults (7876 women and 5141 men) were randomized to take an oral daily capsule of anticxidants (120 mg vitamin C, 30 mg vitamin E, 6 mg & canotione, 100 µg selenium, and 20 mg sincl or a matching placebo. The median time of followup was 7.5 y. A total of 157 cases of all types of SC were reported, from which 25 were melanomas. Because the effect of antioxidants on SC incidence varied according to gender, men and women were analyzed separately. In women, the incidence of SC was higher in the antioxidant group (adjusted hazard ratio (adjusted HR) = 1.68; P = 0.03). Conversely, in men, incidence did not differ between the 2 treatment groups ladjusted HR = 0.60; P = 0.11). Despite the small number of events, the incidence of melanoma was also higher in the antioxidant group for women ladjusted HR = 4.31; P = 0.03; The incidence of nonmelanome SC did not differ between the anticoldent and placebo groups ladjusted HR = 1.37; P = 0.22 for women and adjusted HR = 0.72; P = 0.19 for men). Our findings suggest that antioxidant supplementation affects the incidence of SC differentially in men and women. J. Nutr. 137; 2098-2105, 2007.

### Introduction

Melanoma and normelanoma skin cancers (5C),10 namely squamous cell carcinoma (SCC) and basal cell carcinoma (BCC), are the most common forms of malignancy in the Caucasian population (1) and sun exposure is thought to be the main established risk factor for all 3 types of tumor (2). An aging population, more intense exposure to UV rays due to depletion of the ozone layer, and sun exposure habits would appear to favor a higher incidence of skin malignancy (3).

Numerous studies have demonstrated the role of reactive oxygen species, also called free radicals, in skin carcinogenesis and the potential protective effect of antioxidants (4). Formation

nonmelanoma SC (NMSC) in 1312 individuals with an individual 0022-3166/07 \$8.00 © 2007 American Society for Nutrition Manuscript received 28 February 2007. Initial review completed 21 March 2007. Revision accepted 22 June 2007

of free radicals in the skin can be enhanced by UV ra

cutaneous system has a very efficient interlinked

for their supposed photoprotective properties.

However, excessive exposure to sunlight or other st

light can overwhelm the skin's antioxidant capacity. A poten-

tially interesting strategy for preventing UV exposure damage

could be to boost the endogenous antioxidant system by oral

intake of antioxidant vitamins and minerals. Although clinical

trials have showed contradictory findings (5-7), oral antioxidan

pills have been recommended for the prevention of sunburns and

In particular, it has been suggested that nutrients such as

β-carotene, ascorbic acid, vitamin E, selenium, and zinc may

prevent such harmful effects of UV exposure because of their

antioxidant ability (8). Clinical trials testing the impact of

supplementation with high doses of antioxidants over long

periods have, however, failed to reveal beneficial effects on the

of Cancer trial, a double-blind, randomized clinical trial, was

designed to test whether selenium (200 µg/d) could prevent

incidence of SC (9.10). For example, the Nutritional Prevention

Annals of Internal Medicine

### NIH CONFERENCE

### The Efficacy and Safety of Multivitamin and Mineral Supplement Use To Prevent Cancer and Chronic Disease in Adults: A Systematic Review for a National Institutes of Health State-of-the-Science Conference

Han-Yao Huang, PhD, MPH; Benjamin Caballero, MD, PhD; Stephanie Chang, MD; Anthony J. Alberg, PhD, MPH; Richard D. Semba, MD, MPH; Christine R. Schneyer, MD; Renee F. Wilson, MSc; Ting-Yuan Cheng, MSc; Jason Vassy, MPH; Gregory Prokopowicz, M.D. MPH: George J. Barnes H. BA: and Eric R. Bass, M.D. MPH

Background: Multivitamin and mineral supplements are the most commonly used dietary supplements in the United States.

Purpose: To synthesize studies on the efficacy and safety of multivitamin/mineral supplement use in primary prevention of cancer gastric cancer and the overall mortality rate from cancer by 13% to 21%, in a French trial, combined supplementation with vitamin C. vitamin E, \$-carotene, selenium, and zinc reduced the rate of cancer by 31% in men but not in women. Multivitamin and mineral supplements had no significant effect on cardiovascular disease

### ORIGINAL INVESTIGATION

### Dietary Supplements and Mortality Rate in Older Women

The Iowa Women's Health Study

Jaakko Mursu, PhD; Kim Robien, PhD; Lisa J. Harnack, DrPH, MPH, Kyong Park, PhD; David R. Jacobs Jr, PhD

Buckground: Although dietary supplements are commonly taken to prevent chronic disease, the long-term health consequences of many compounds are unknown. States (1). Examinari Methods: We assessed the use of vitamin and mineral supplements in relation to total mortality in 38 772 older recent use

61.6 years at baseline in 1986. Supplement use was selfquate intareported in 1986, 1997, and 2004. Through December monly us 31, 2008, a total of 15594 deaths (40.2%) were identisupplemen fied through the State Health Registry of Iowa and the Many National Death Index. duding ci style, and

Results: In multivariable adjusted proportional hazards regression models, the use of multivitamins (hazard ratio, 1.06: 05% CL 1.02-1.10; absolute risk increase, 2.4%), vitamin B. (1.10: 1.01-1.21: 4.1%). folic acid (1.15: 1.00-1.32; 5.9%), iron (1.10; 1.03-1.17; 3.9%), magnesium (1.08; 1.01-1.15; 3.6%), zinc (1.08; 1.01-1.15; 3.0%), and cop-

women in the Iowa Women's Health Study; mean age was

per (1.45; 1.20-1.75; 18.0%) were associated with increased risk of total mortality when compared with corresponding nonuse. Use of calcium was inversely related (hazard ratio, 0.91; 95% confidence interval, 0.88-0.94; absolute risk reduction, 3.8%). Findings for iron and calcium were replicated in separate, shorter-term analyses (10year, 6-year, and 4-year follow-up), each with approximately 15%-of the original participants having died, starting in 1986, 1997, and 2004.

Conclusions: In older women, several commonly used dietary vitamin and mineral supplements may be associated with increased total mortality risk; this association is strongest with supplemental iron. In contrast to the findings of many studies, calcium is associated with decreased risk

Arch Intern Med. 2011;171(18):1625-1633

these facto

damage, i

Data Source

EMBASE, A

hand-seard

Study Sele

viewed to

observation

Data Extra

dently asser

**Data Synth** 

trials and 3

quality was

disease, cat

for the stu

population.

evol, and set

use multiv

safety.

defense system for counteracting UV-induced exis Downlanded From ht

Department of Health Sciences, Institute of Public Health and Clinical Nutrition, University of Eastern Finland, Kuopio Campus, Kuopio, Finland (Dr Mursu); Division of Epidemiology and Commu Health, School of Public Health University of Minnesota Minneapolis (Drs Mursu Robten, Harnack, and Jacobs') Department of Food and Nutrition, Yeungnam University, Gyeongbuk Republic of Korea (Dr Park): and Department of Nutrition. School of Medicine, University of Oslo, Oslo, Norway (Dr Jacobs).

Author Affiliations

\$20 billion.13 Sixty-six percent of women participating in the Iowa Women's Health Study1 used at least 1 dietary supplement daily in 1986 at an average age of 62 years; in 2004, the proportion increased to 85%. Moreover, 27% of women reported using 4 or more supplemental products in 2004. At the population level, dietary supplements contributed substantially to the total intake of several nutrients, particularly in

elderly individuals.12 Supplemental nutrient intake clearly is beneficial in deficiency conditions.4 However, in well-nourished populations, supplements often are intended to yield benefit by preventing chronic diseases. Results of epidemiologic studies54 assessing supple ment use and total mortality risk have been. inconsistent. Several randomized controlled trials (RCTs), 20,11 concentrating mainly on calcium and vitamins B, C, D and E, have not shown beneficial effects of

### See Invited Commentary and Editor's Note at end of article

dietary supplements on total mortality rate; in contrast, some<sup>13,13</sup> have suggested the possibility of harm. Meta-analyses14,37 concur in finding no decreased risk and potential harm. Supplements are widely used, and further studies regarding their health effects are needed. Also, little is known about the long-term effects of multivitamin use and less commonly used supple ments, such as iron and other minerals

ARCH INTERN MEDIVOL 171 (NO. 18), OCT 10, 2011 WWW.ARCHINTERNMED.COM

IN THE UNITED STATES, THE USE OF

dietary supplements has in-

creased substantially during the

past several decades,15 reaching

approximately one-half of adults

in 2000, with annual sales of more than

**02011** American Medical Association, All rights reserved.

Downloaded From: http://wrchinte.jamanetwork.com/ on 81/26/2014

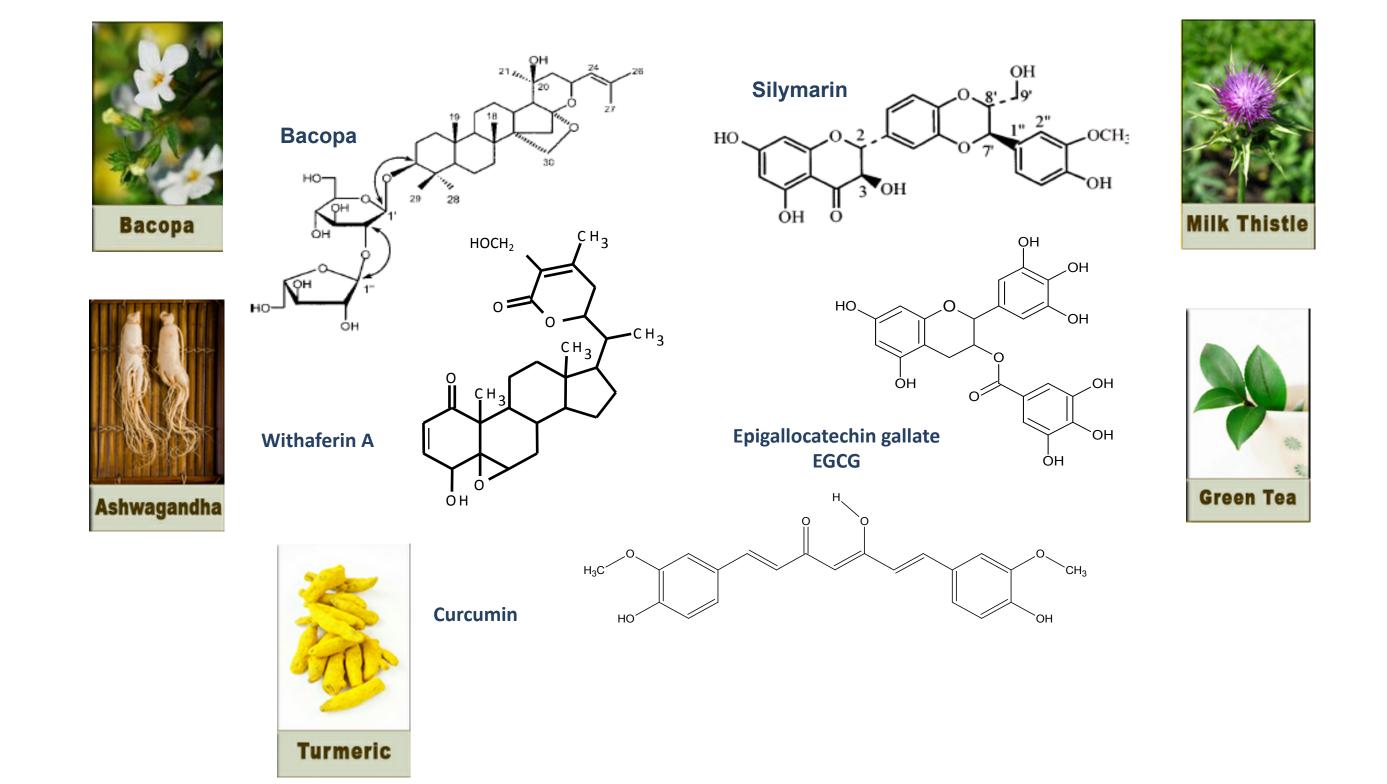
Author disclosures: S. Hersberg, K. Ezzedine, P. Proziosi, P. Galan, S. Bertrais, Estaquio, S. Briangon, A. Favier, and D. Malvy, no conflicts of interest, C. Guinot and J. Latreite, the CERLES, is a research center on human skin.

Abbreviations used BCC, bessi cell cercinome; HR, hazard ratio; MSC. melanoma skin cancer; NMSC, nonmelanoma skin cancer; SC, skin cancer; SCC, squamous cell carcinoma: SUVI.MAX, Supplementation on Vitamines et Minéraux Antioxydents study.

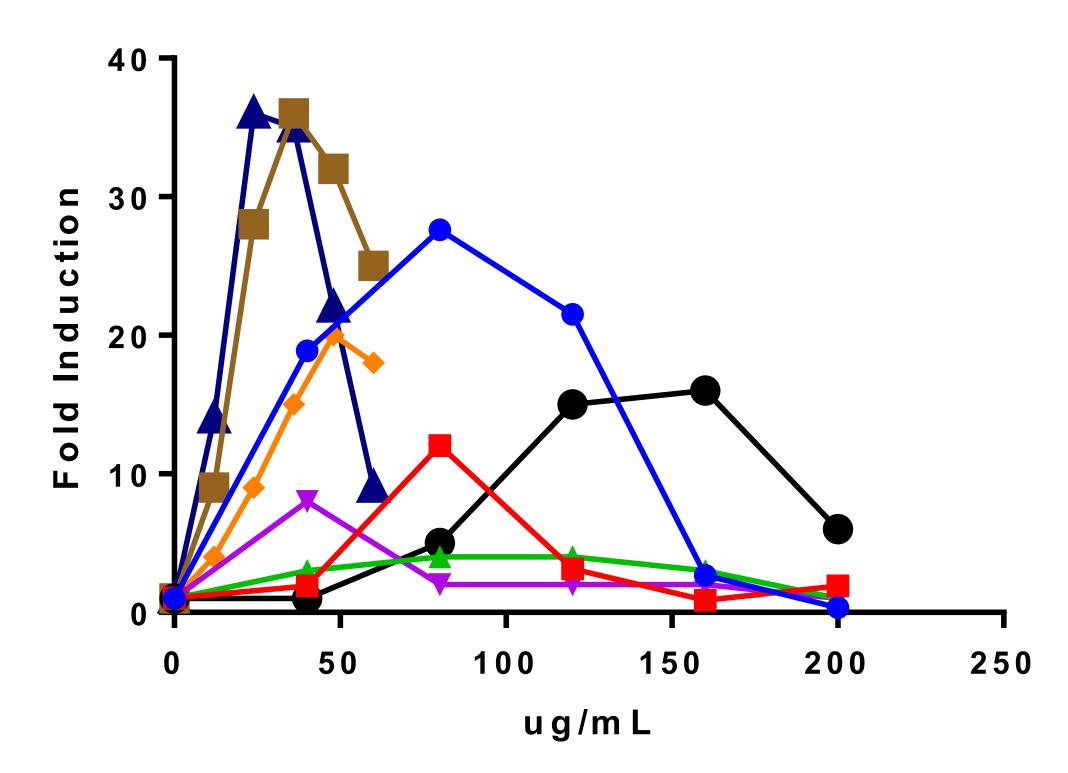
<sup>\*</sup> To whom correspondence should be addressed. E-mail: heroberg@cnam.fr.



### Nrf2 = a powerful "master regulator" of antioxidant enzymes and survival genes



### ARE reporter assay

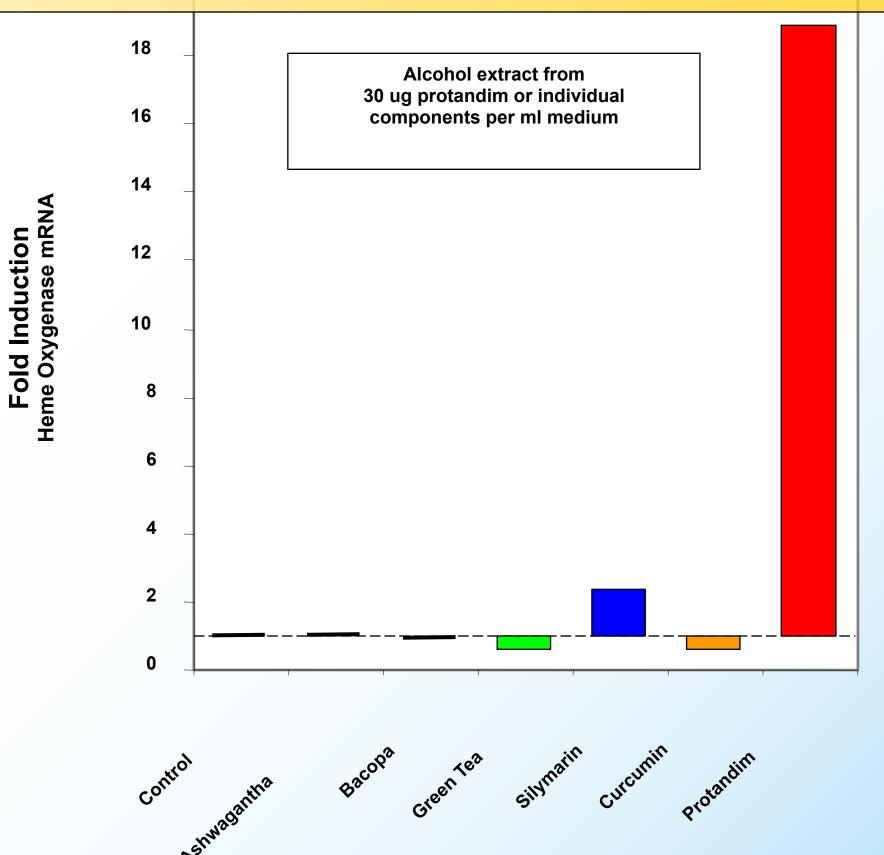


- Ashwaghanda
- Black Pepper
- → Milk Thistle
- Tumeric
- → Green Tea
- Bacopa
- Protandim US
- Protandim Japan

### SYNERGY = Action greater than the sum of the parts

All five ingredients together produced an 18-fold increase in the expression of this antioxidant gene.

Protandim BLEND works 18 times more effectively than the sum of its parts.



### Protandim Antiinflammatory and Antifibrotic Proteins Kinase Kinase Kinase Stress-related Enzymes Keap1 Nrf2 DNA Human cell Nucleus



### Available online at www.sciencedirect.com





Free Radical Biology & Medicine 40 (2006) 341 - 347

www.elsevier.com/locate/freeradbiomed

### Original Contribution

## The induction of human superoxide dismutase and catalase in vivo: A fundamentally new approach to antioxidant therapy

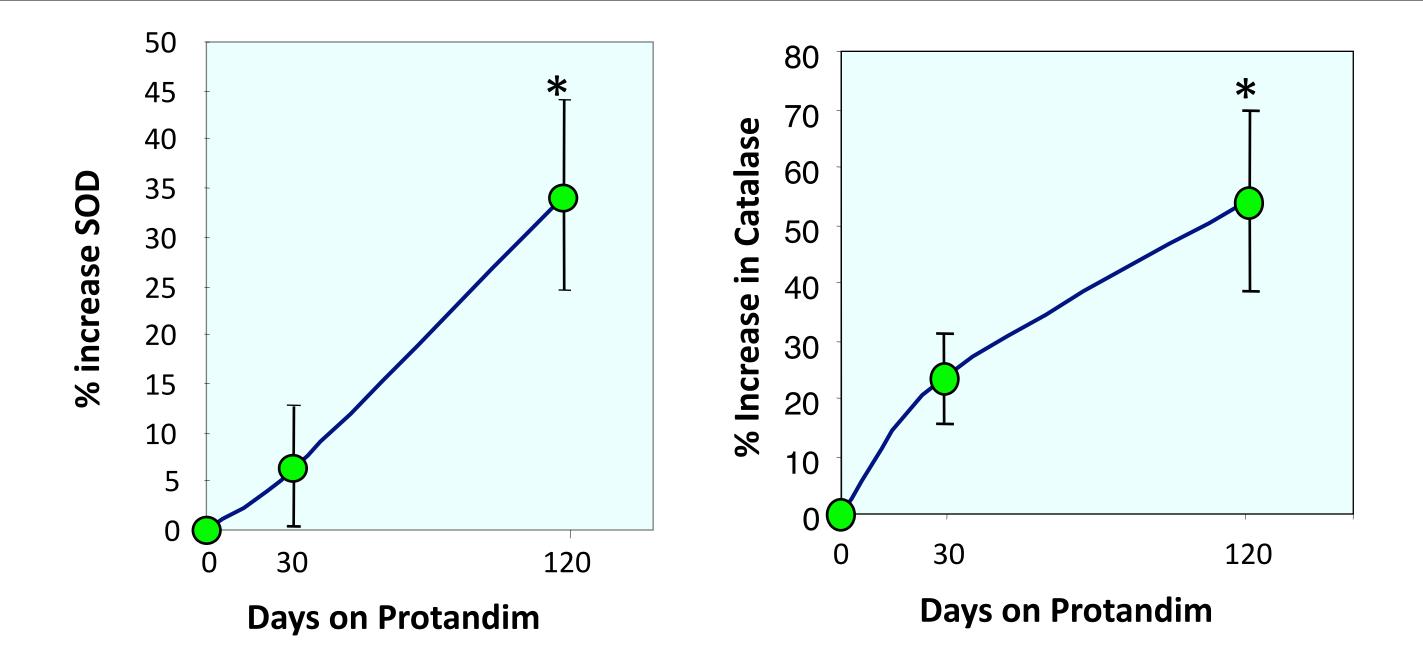
Sally K. Nelson a,b, Swapan K. Bose a, Gary K. Grunwald c, Paul Myhill d, Joe M. McCord a,b,d,\*

Webb-Waring Institute for Cancer, Aging and Antioxidant Research, University of Colorado Denver Health Sciences Center, Denver, CO 80262, USA
 Department of Medicine, University of Colorado Denver Health Sciences Center, Denver, CO 80262, USA
 Department of Preventive Medicine and Biometrics, University of Colorado Denver Health Sciences Center, Denver, CO 80262, USA
 Lifeline Therapeutics, Denver, CO, USA

Received 22 June 2005; revised 24 August 2005; accepted 28 August 2005

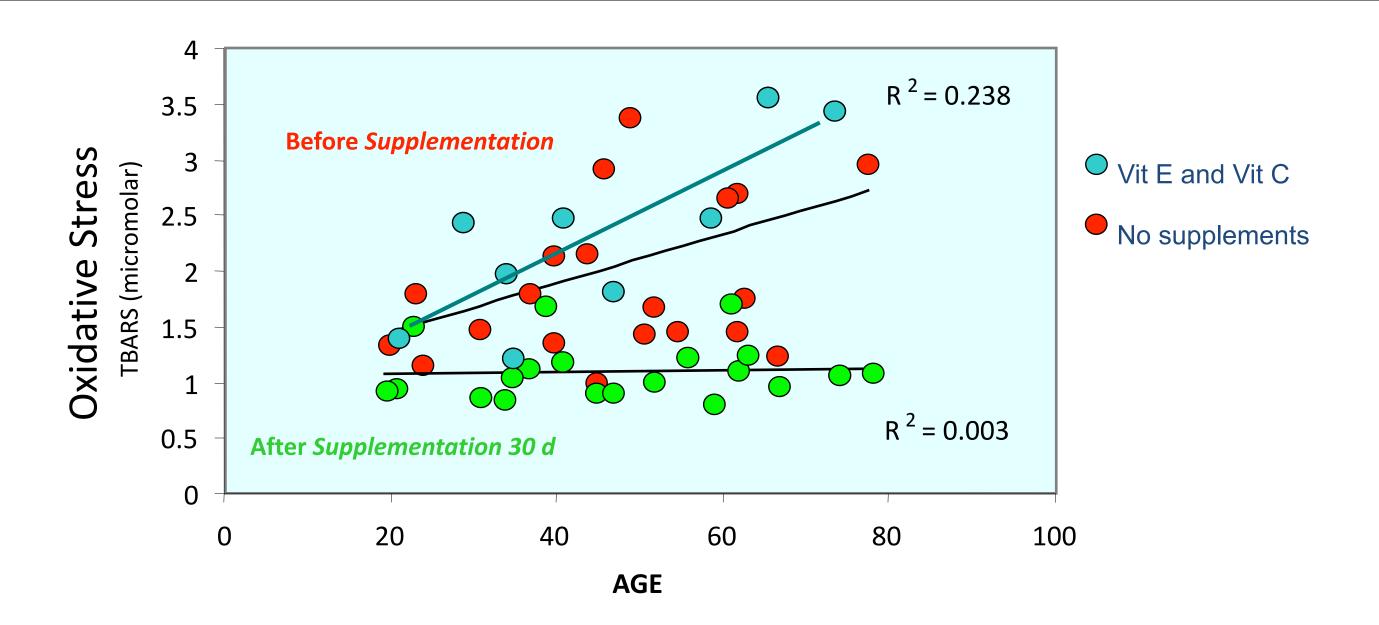
### Abstract

A composition consisting of extracts of five widely studied medicinal plants (Protandim) was administered to healthy human subjects ranging in age from 20 to 78 years. Individual ingredients were selected on the basis of published findings of induction of superoxide dismutase (SOD) and/or catalase in rodents in vivo, combined with evidence of decreasing lipid peroxidation. Each ingredient was present at a dosage sufficiently low to avoid any accompanying unwanted pharmacological effects. Blood was analyzed before supplementation and after 30 and 120 days of supplementation (675 mg/day). Erythrocytes were assayed for SOD and catalase, and plasma was assayed for lipid peroxidation products as thiobarbituric acid-reacting substances (TBARS), as well as uric acid, C-reactive protein, and cholesterol (total, LDL, and HDL). Before supplementation, TBARS showed a strong age-dependent increase. After 30 days of supplementation, TBARS declined by an average of 40% (p = 0.0001) and the age-dependent increase was eliminated. By 120 days, erythrocyte SOD increased by 30% (p < 0.01) and catalase by 54% (p < 0.002). We conclude that modest induction of the catalytic antioxidants SOD and catalase may be a much more effective approach than supplementation with antioxidants (such as vitamins C and E) that can, at best, stoichiometrically scavenge a very small fraction of total oxidant production.



After 120 days...

SOD increased by 34% Catalase increased by 54%



After 30 days...

"Remarkably, this age-dependent increase in TBARS was almost completely abolished by Protandim treatment (Fig. 1D), with an overall average reduction of the oxidative stress marker by 40%."

Exotic Ingredients + Proven Science + Exclusive IP = World's finest products that help you Feel / Look / Perform Your Best

Protandim/Nrf2 – Fundamentally different approach to cellular protection



Coordinated Product Platform



## LOOK Your Best...

### TrueScience Regimen is equal/better compared to top "prestige" brands

Competitor		28 Days	56 Days
89% Perricone MD Cold Plasma at 4 weeks 75% Jeunesse Global Luminesce Cellular Rejuvenation at 8 weeks	Smoother looking skin	89%	94%
80% Nu Skin TruFace at 3 months	Firmer looking skin	81%	85%
78% L'Oreal Youth Code at 8 weeks	Younger looking skin		87%
70% Lancôme Dream Tone at 8 weeks 79% SkinMedica Lytera at 12 weeks	More even skin tone		83%
79% Clarins Double Serum at 4 weeks [Hydric + Lipidic System] 80% Nu skin 180°System at 8 weeks	Less noticeable fine lines and wrinkles	78%	82%

Cold Plasma is a trademark of Perricone MD
Youth Code is a trademark of L'Oreal
Dream Tone is a trademark of Lancôme
Lytera is a trademark of SkinMedica
Double Serum [Hydric + Lipidic System] is a trademark of Clarins

\*Competitive advertising details available upon request



## Clinical Study: What Users Said\*

28 Days		56 Days
94%	Loved the fragrance	99%
90%	More hydrated skin	95%
89%	Will buy the regimen	91%
89%	Smoother looking skin	94%
88%	Softer skin	90%
84%	More luminous skin	88%
83%	Younger looking skin	87%
81%	More even skin tone	83%
81%	Firmer looking skin	85%
78%	Less noticeable fine lines and wrinkles	82%
74%	Felt younger-looking	80%
74%	Better than what I usually use	80%



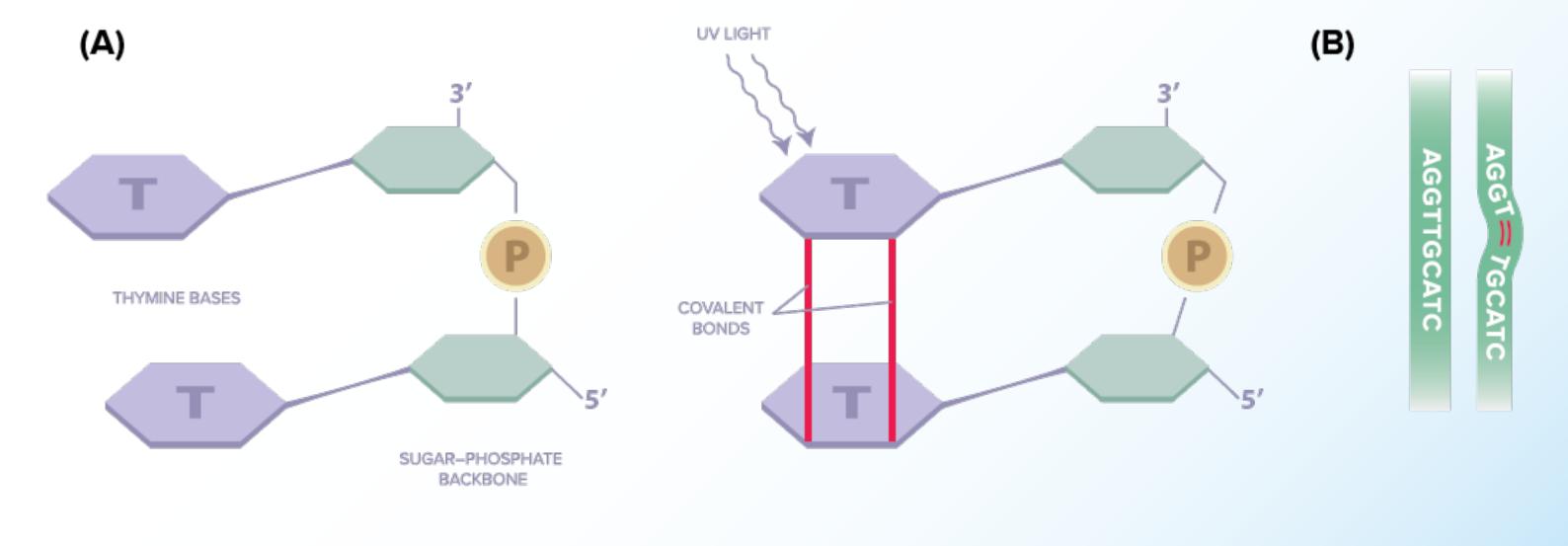
<sup>\*</sup>Satisfaction test, 86 women self-reported, 4 weeks and 8 weeks

- TrueScience™ facial cream with Advanced Nrf2
   Technology applied 2 times per day for 5 days
   (Nrf2 Technology: Bacopa, Milk Thistle, Turmeric, Green tea & Black Pepper extracts + Brassicas extract + Plantain extract)
- Non Nrf2 cream applied 2x per day for 5 days
- Then expose the samples to UV light
- Quantify DNA damage and amount of Nrf2 Protein
- Then collect DNA, run gene expression & identify biological pathways





### Following UV exposure, the cellular DNA is damaged (Thymine Dimers)



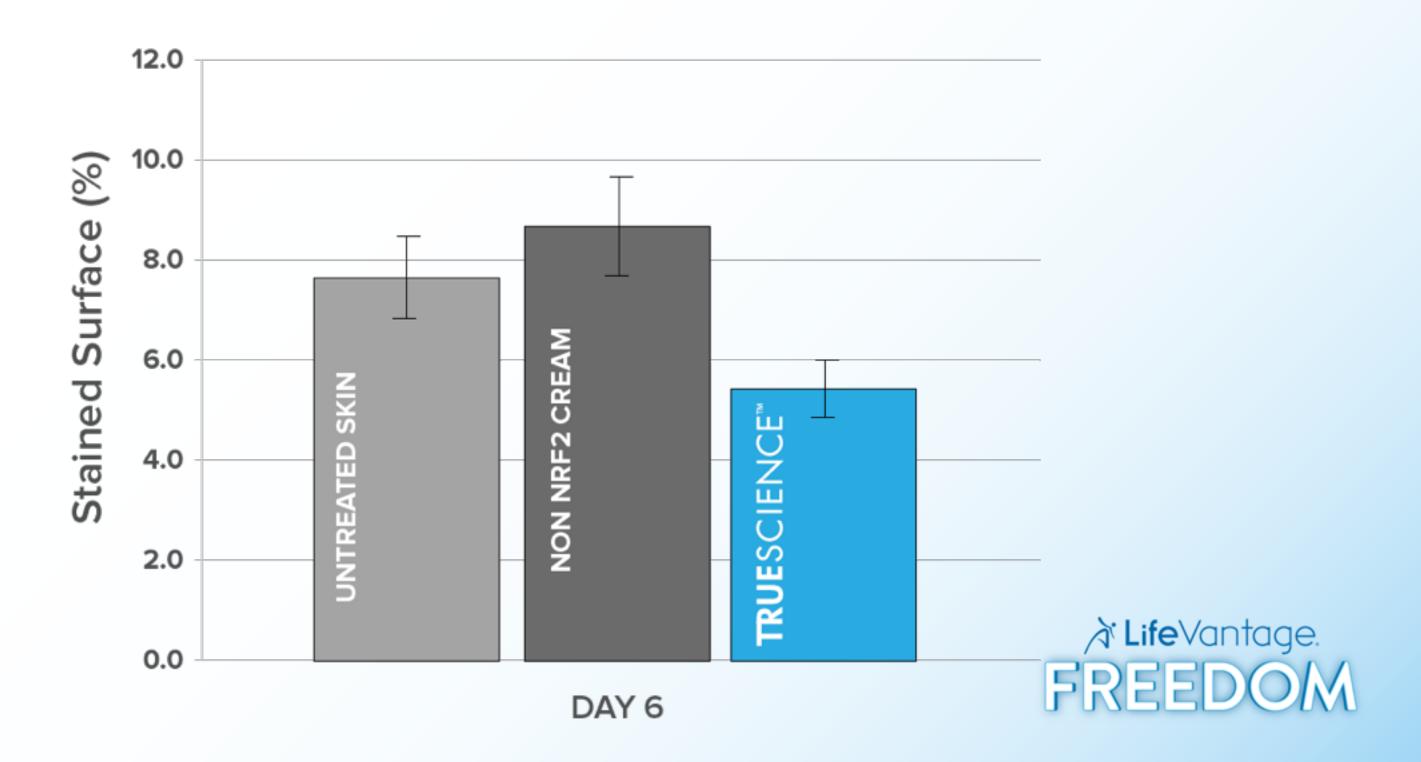
Affects DNA duplication

**Causes genome instability** 



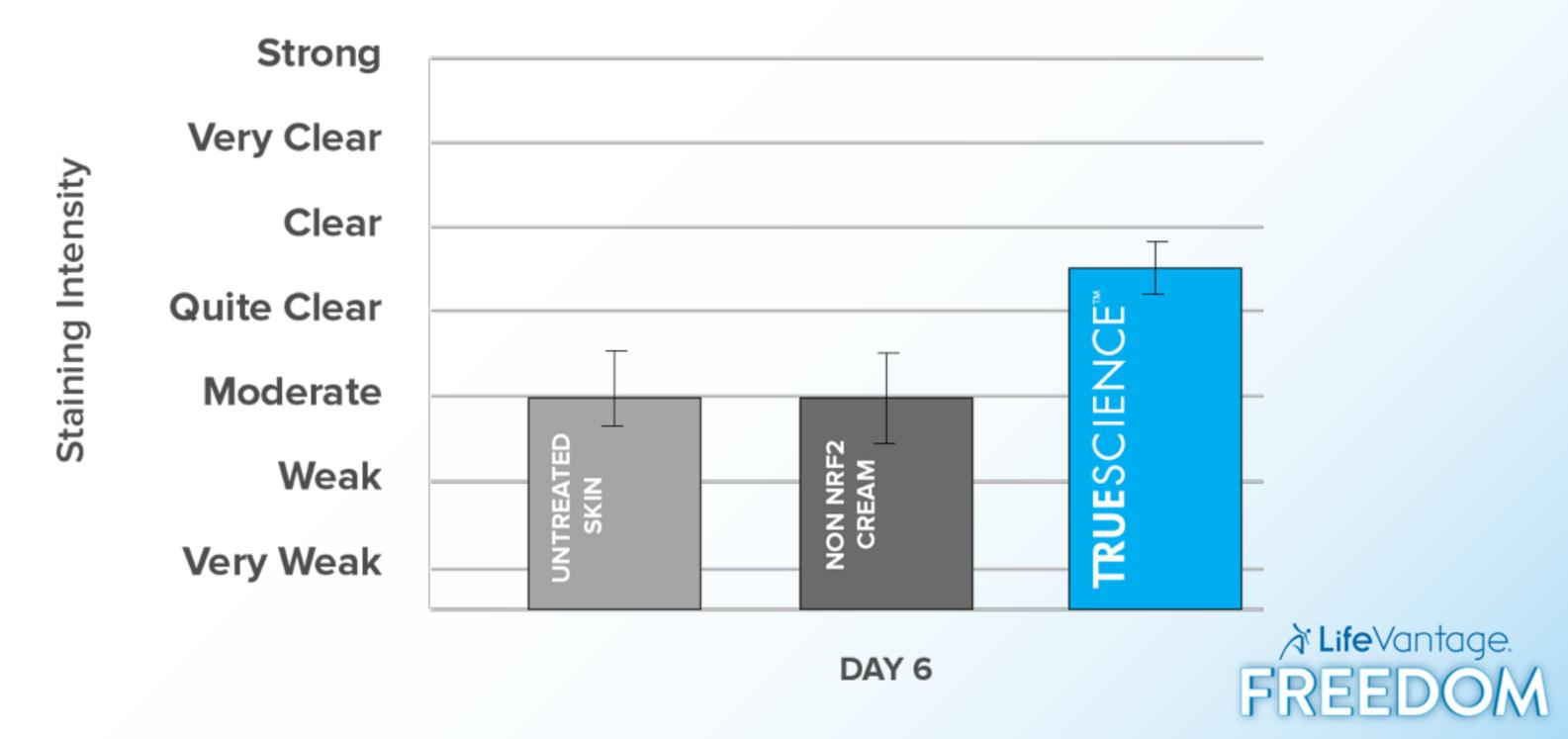
### Less Thymine Dimers with TrueScience

TrueScience™ Facial Cream with Nrf2 protects the cell DNA!



### Nrf2 Staining Intensity 24 hours after UV Exposure

TrueScience™ Facial Cream results in more Nrf2 in skin, thus more protection!



### Cascade Effect = All Skin Layers

healthier skin tissue homeostasis balance and maintain serine protease activities SPINK6 6 KLK12 6 in the skin cholesterol synthesis DHCR724 NUS124 intracellular cholesterol trafficking **SOX21<sup>24</sup> DOK2<sup>24</sup>** BMP6<sup>6</sup> **Cell differentiation** CEACAM16 CEACAM5<sup>6</sup> **Cell proliferation** CCAR1<sup>6</sup> **mRNA DNA** repair ARID3A<sup>24</sup> EVL<sup>24</sup> structural integrity potential marker of KRT19 6 of epithelial cells epidermal stem cells TIMP4<sup>24</sup> MMP7<sup>6</sup> tissue remodeling

Cornified

Granular

layers

**Spinous** 

layers

Basal

layer

**Dermis** 

layers

TS Facial cream initiates signals that cascade all the way though the deep layers of the skin

Resulting in better moisturization, protection, & younger looking skin



## TrueScience<sup>tm</sup> Facial Cream with Advanced Nrf2 Technology has been shown to:

- Boost skin protection from UV exposure by reducing DNA damage
- Increase Nrf2 Protein amount, thus improve resistance to oxidative stress
- Fight the signs of aging though all layers of the skin





## FEEL Your Best...



Tired, Stressed, Depressed... "Off"











































#### SUPPLEMENT FACTS

Serving Size: 1 Packet Serving Per Pouch: 30

Amount Per Serving		% DV
Calories	10	
Total Carbohydrate	2 g	<1%*
Niacin (as Nicotinic Acid)	24 mg	120%
Vitamin B6 (as Pyridoxine HCl	_) 1.60 mg	80%
Vitamin B12 (as Methylcobala	min 6 mcg	100%
Magnesium (as Magnesium As	spartate) 10 mg	2%
Caffeine	100 mg	t
Proprietary Blend	500 mg	10.70
DMAE Bitartrate		†
Green Tea Extract (Camellia sinensis) (Aerial)		†
Quercetin Dihydrate	- 18 18 18 18 18 18 18 18 18 18 18 18 18	†
Monterey Pine Extract (Pinus radiata) (Bark)		†
L-Theanine	., .	Ť

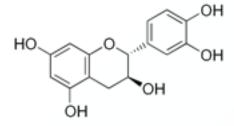
<sup>\*%</sup> Daily Value are based on a 2,000 calorie diet.

### SUPPLEMENT FACTS

Serving Size: 1 Packet Serving Per Pouch: 30

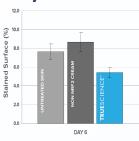
	Amount Per Serving	% DV
Calories	15	
Total Carbohydrate	3 g	1%*
Niacin (as Nicotinic Acid)	20 mg	100%
Vitamin B6 (as Pyridoxine HCL	) 1.60 mg	80%
Vitamin B12 (as Methylcobalar	min 6 mcg	100%
Magnesium (as Magnesium As	partate) 10 mg	2%
Proprietary Blend	280 mg	
Green Tea Extract (Camellia sinensis) (Aerial)		†
Monterey Pine Extract (Pinus radiata) (Bark)		†
L-Theanine		†
Quercetin Dihydrate		†

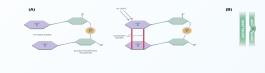
<sup>\*%</sup> Daily Value are based on a 2,000 calorie diet.



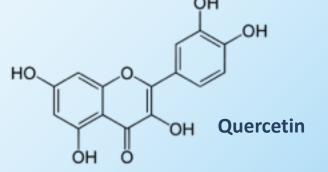
Catechin

#### Proanthocyanidin





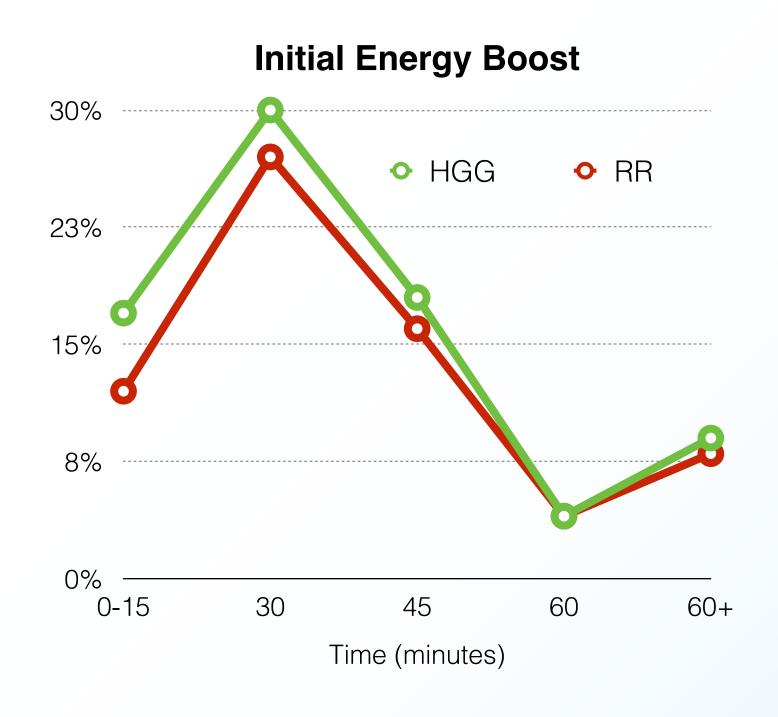
L-Theanine

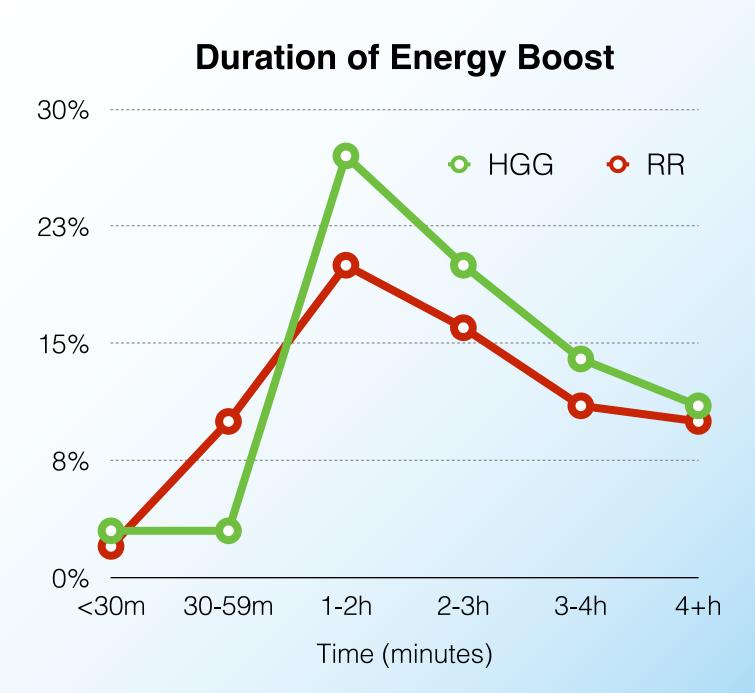


<sup>†</sup> Daily Value not established

<sup>†</sup> Daily Value not established

## Axio Usage Survey





# Axio Usage Survey



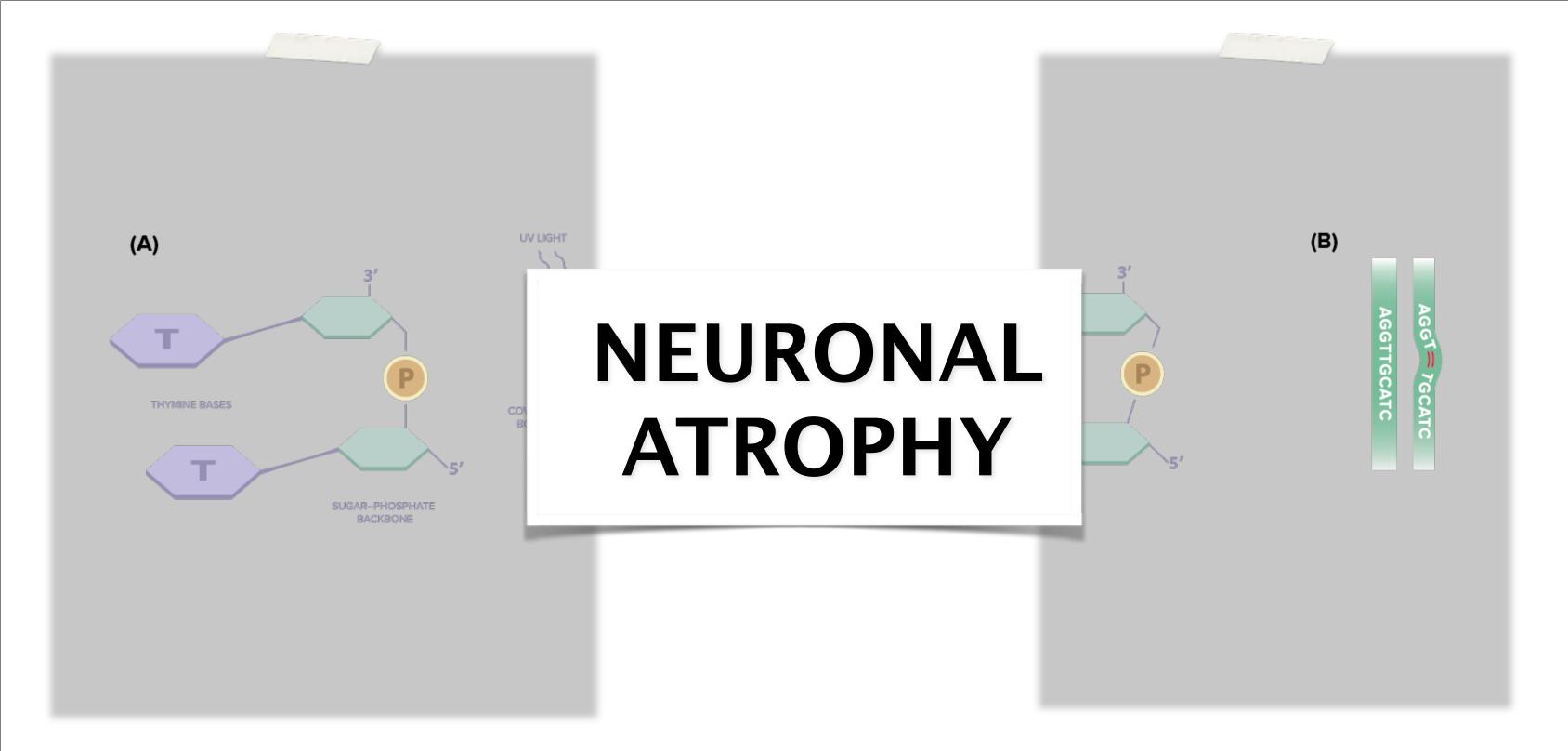
# Axio Usage Survey

- "It felt like a morning cup of coffee with the energy it gave me, but more than that it improved my focus."
- "I liked the energy that it gave me, I would say it lasted pretty good maybe 4 or 5 hours."
- "I was pleasantly surprised at this product's ability to keep me energized, awake and focused without harmful stimulants and without feeling nervous, jittery or having rapid heart rate. I would take this over caffeine any day."
- "It was a very subtle transition to having energy, just like I naturally had the energy. It was a good amount of energy too. Not too wired, not too draggy. The perfect amount. I didn't feel like it wore off halfway through the day or that I needed more energy. I also did not have a difficult time falling asleep at night because of it. I would definitely buy this instead of many other energy drinks or supplements."



## PERFORM Your Best...





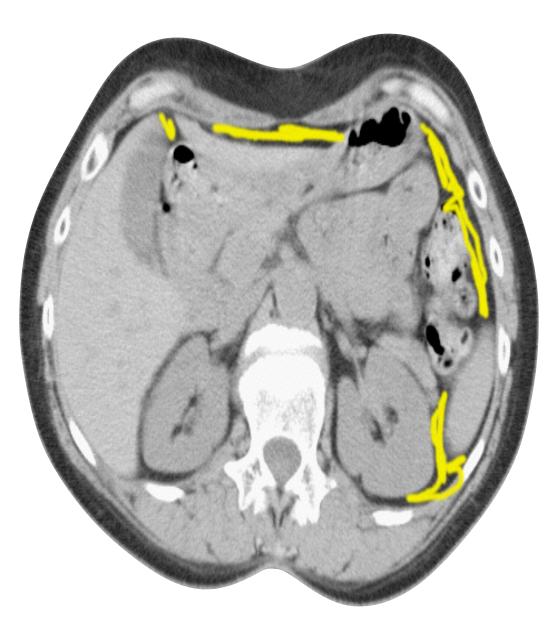
#### **NORMAL STRESS**

Healthy, Large, Many Projections, Optimal Function

#### **HIGH STRESS**

Small, Thin, Disrupted, Structural Damage, Poor Function

## ABDOMINAL FAT ACCUMULATION



Normal Stress



High Stress

## Research Study Update

Completed<sup>1</sup>, Ongoing<sup>2</sup>, Planned<sup>3</sup>

- 19 studies<sup>1</sup> (U Colorado, Ohio State U, Louisiana State U, Virginia Commonwealth U, Colorado State U, Texas Tech U...)
- "20th study" (Mayo Clinic, 2014)<sup>1</sup> anecdotal patient report prompts series of translational cell culture and rodent studies
  - Translational research = aims to make findings from basic science useful for practical applications that enhance human health and well-being
- Montreal, Canada (skin)<sup>1</sup>
- National Institutes of Health (longevity)<sup>2</sup>
- Nashville, TN (heart health)<sup>2</sup>
- Melbourne, Australia (brain function)<sup>3</sup>
- Okinawa, Japan (lung function)<sup>3</sup>
- Research Institutions<sup>3</sup> (Salt Lake, Miami, Louisville, Fort Collins, Boston, NYC...)
   Life Vantage.
  - energy/mood/focus, performance, antioxidant metabolism, eye health...
  - canine health, periodontal health, blood sugar balance...











































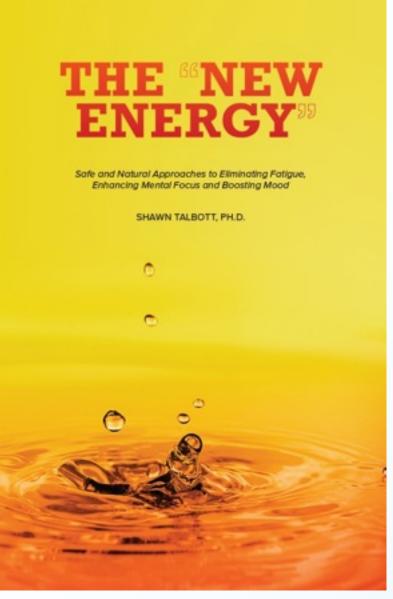
# 



Why Your Daily Vitamins May Be Causing Cancer and Shortening Your Life and How Nrf2 Can Turn on Your Body's Own Antioxidants for Optimal Health







## ProductQuestions@LifeVantage.com

www.ShawnTalbott.com

www.ShareCare.com

FaceBook / Twitter / Amazon



