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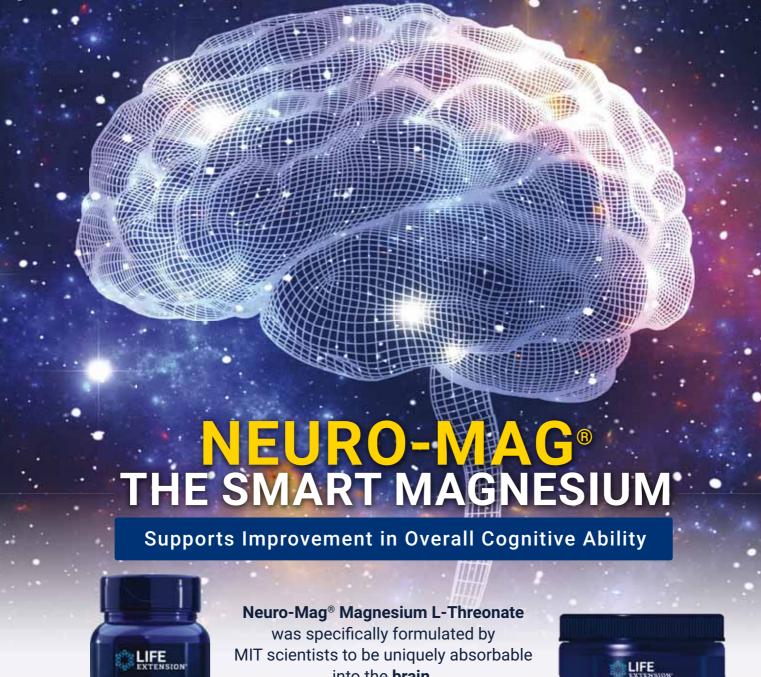
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May 2024

FEATURE ARTICLES

- 7 Rationale for Combination Approaches
- 22 Human Brain Scores and Benfotiamine
- 32 Promoting Healthy Longevity
- 44 GLP-1 Agonist Drugs: Good and Bad
- 52 Clinical Trials: Cancer & Vitamin D
- 65 What is PQQ?
- 75 Heart Attack Risk-Reducing Food







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References: 1. Neuropharmacology. 2016 2016/09/01/;108:426-39. 2. Journal of Cellular Physiology. 2016;231(9):1903-12. 3. J Alzheimers Dis. 2016;49(4):971-90.

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May 2024



with reduced mortality and greater healthspan by modulating multiple mechanisms of aging.

REPORTS

22 HUMAN BRAIN AGING AND BENFOTIAMINE

Benfotiamine has been shown to help defend against damage inflicted by blood sugar and slow measures of human cognitive decline.



44 GLP-1 FOR OBESITY AND CARDIAC FUNCTION

GLP-1 agonist drugs have emerged as game changers in controlling blood sugar and body weight. Clinical trials show that they also result in a 20% reduction in cardiovascular events and death.



52 CLINICAL TRIALS: VITAMIN D AND CANCER

Observational studies have shown an association between low vitamin D and increased risk for many cancers. Clinical trials indicate who is most likely to benefit by supplementing with more vitamin D.



65 WHAT IS PQQ?

By promoting the generation of new cellular mitochondria, **PQQ** helps promote full-body health.

DEPARTMENTS

7 RATIONALE FOR **COMBINED INTERVENTIONS**

In a massive review published in 2023, scientists stressed the urgent need for combination approaches to target degenerative aging.

Interventions that favorably affect differing aging mechanisms are widely available to consumers today.

15 IN THE NEWS

Resveratrol boosts endothelial function in diabetics; melatonin improves recovery factors in male soccer players; and more.

69 ASK THE DOCTOR: CHECK YOUR HEALTH WITH LAB TESTS

Dr. Fogle discusses the latest musthave lab tests available to help you and your doctor understand your current health status.



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CAUTION: Individuals consuming more than 50 mcg (2000 IU)/day of vitamin D (from diet and supplements) should periodically obtain a serum 25-hydroxy vitamin D measurement. Vitamin D supplementation is not recommended for individuals with high blood calcium levels.

† Ratings based on results of the 2023 ConsumerLab.com Survey of Supplement Users. More information at www.consumerlab.com/survey.

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Rationale for Combined Interventions



WILLIAM FALOON



A massive scientific review published in **2023** describes "aging" as a top global concern.¹

The authors emphasize the **urgent** need for <u>effective</u> interventions to <u>decrease</u> late-life disorders and improve **human** healthspans.

The authors note that the number of compounds that have extended **lifespan** in laboratory models has nearly **tripled** in recent years.

These compounds largely function via differing biological mechanisms.

This prompted the authors to propose a rationale for <u>combinations</u> of **anti-aging** interventions, something that readers of this magazine attempt to do today.

In recent years, <u>low</u>-cost **nutrients** have been identified that are associated with *longer* **lifespans**.

One challenge is to ingest **optimal potencies** that target specific age-accelerating pathways. None of us wants to omit **affordable** nutrients that have life-extending potential.

A nutrient that has garnered attention in recent years is **spermidine**. Found in wheat and other foods, **spermidine** has been shown to <u>increase</u> median **lifespan** in different species, including:²⁻⁶

300% in yeast
30% in flies
10% in older, aged mice

Observational studies in people associate <u>increased</u> **spermidine** ingestion with <u>lower</u> risk of **cardiovascular**⁷ and **cognitive** disorders,^{8,9} along with reduced **mortality**.¹⁰⁻¹²

Spermidine functions via several mechanisms including boosting **cellular housekeeping** (autophagy).^{4,13-15} Some scientists describe **spermidine** as a **calorie-restriction** mimetic.¹⁶

We welcome randomized controlled trials that attempt to validate the lifespan-enhancing effects of differing interventions in people. For those seeking to delay degenerative aging now, we don't have time to wait for results from multi-decade clinical trials.



Taurine and Lithum

A landmark study published in the journal **Science** described the amino acid **taurine** as an "anti-aging intervention."¹⁷

The authors of this study demonstrated how **taurine** levels **plummet** as **people** grow older. They showed *higher* levels of taurine are associated with <u>lower</u> BMI and lower prevalence of **abdominal obesity**, **inflammation**, and **type II diabetes**.

When taurine was supplemented in middle-aged mice, life expectancy <u>increased</u> by **18%-25%**. Taurine-fed mice appeared healthier, having <u>less</u> body fat, <u>more</u> bone mass, and better **insulin sensitivity**.

The dose of **taurine** found to be effective at supporting cardiovascular health in **human** trials is about **5,000 mg** a dav. ¹⁸⁻²⁰

It requires **1,500 mg** of wheat germ to obtain the desired potency of **spermidine**.^{2,4,9-11,21}

A mineral we've talked about for many years is **lithium**. In regions with *higher* lithium in drinking water, people tend to **live longe**.²²⁻²⁴

Managing the bottles and swallowing the capsules needed to obtain these potencies can be challenging.

This prompted the formulation of a <u>combination</u> of **nutrients** into a daily scoop of powder. The savings and ability to easily ingest *higher* potencies make compliance simpler than it has been.

The article on page 32 of this month's issue outlines the science that supports nutrients associated with *longer* life and enhanced healthspans.

The box below summarizes a massive review emphasizing the need for <u>combination</u> approaches to target **aging.**

What Can You Do Today?

The most important steps to avoid premature morbidities and mortality are to optimize **blood pressure** and have comprehensive **blood tests**.

Blood tests can identify <u>correctable</u> risk factors long *before* they manifest outwardly as serious illness or death.

The most popular test our supporters order is the **Male** or **Female Blood Panels** described on the next page.

Commercial labs charge around \$2,500 for <u>all</u> these tests, but they are available once a year from **Life Extension**® for only \$224—a savings of around 90%.

Upon receiving your order, we'll send you a list of blood draw stations in your area where you can walk in at your **convenience** for a blood draw and then receive **rapid reporting** of the results.

Blood tests can be ordered 24-hours-a-day by calling **1-800-208-3444** or visiting **www. LifeExtension.com/sales/lab-tests**

For longer life,

William Faloon, Co-Founder

Life Extension®

(References for this editorial can be found on page 10.)

Compound Combinations Targeting Longevity: Challenges and Perspectives

"Anti-aging drug therapy is one of the most promising strategies to combat aging."

"Most geroprotective compounds affect only a few biological targets."

"There is a great rationale for use of combinations of anti-aging interventions."

"Aging is currently at the top of the major global concerns, urgently requiring effective, large-scale interventions to decrease the number of late-life disorders and improve human healthspan."

Published January 13, 20231

COMPREHENSIVE BLOOD TESTS AT LOW LAB SALE PRICES





Commercial labs charge around \$2,500 for blood tests needed to evaluate cardiac, inflammatory, immune, and other degenerative risk factors.

Life Extension offers these same tests for \$224 when the Male or Female Panel is ordered during the annual Lab Test Sale. This represents a savings of about 90% compared to commercial labs.

Regular Price: \$299

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MALE PANEL

GLUCOSE PROFILE

Insulin Hemoglobin A1c Glucose

CARDIAC MARKERS

C-Reactive Protein (high sensitivity)
Apolipoprotein B (ApoB)
Homocysteine
Total Cholesterol
LDL (low-density lipoprotein)
HDL (high-density lipoprotein)
Triglycerides

MINERALS

Magnesium (serum)
Ferritin (measure of iron status)
Calcium
Potassium
Phosphorus
Sodium
Iron
Chloride

ORGAN FUNCTION

Kidney: creatinine,
BUN, uric acid, BUN/creatinine ratio
Liver: AST, ALT, LDH,
GGT, bilirubin, alkaline phosphatase

COMPLETE BLOOD COUNT (CBC)

Blood proteins: albumin, globulin, total protein, albumin/globulin ratio Immune Cells including: lymphocytes, monocytes, eosinophils, neutrophils, basophils

Red Blood Cells including: hemoglobin, hematocrit, MCV, MCH, MCHC, RDW

Platelets

HORMONES

Free and Total Testosterone
DHEA-S
Estradiol (an estrogen)
TSH (thyroid function)
Vitamin D (25-hydroxyvitamin D)

CANCER MARKER

PSA (Prostate Specific Antigen)

FEMALE PANEL

GLUCOSE PROFILE

Insulin Hemoglobin A1c Glucose

CARDIAC MARKERS

C-Reactive Protein (high sensitivity)
Apolipoprotein B (ApoB)
Homocysteine
Total Cholesterol
LDL (low-density lipoprotein)
HDL (high-density lipoprotein)
Triglycerides

MINERALS

Magnesium (serum)
Ferritin (measure of iron status)
Calcium
Potassium
Phosphorus
Sodium
Iron
Chloride

ORGAN FUNCTION

Kidney: creatinine, BUN, uric acid, BUN/creatinine ratio Liver: AST, ALT, LDH, GGT, bilirubin, alkaline phosphatase

COMPLETE BLOOD COUNT (CBC)

Blood proteins: albumin, globulin, total protein, albumin/globulin ratio Immune Cells including: lymphocytes, monocytes, eosinophils, neutrophils, basophils

Red Blood Cells including: hemoglobin, hematocrit, MCV, MCH, MCHC, RDW

Platelets

HORMONES

Progesterone
Free and Total Testosterone
DHEA-S
Estradiol (an estrogen)
TSH (thyroid function)
Vitamin D (25-hydroxyvitamin D)

To obtain these extensive Male or Female Panels at these low prices, call 1-800-208-3444 or visit www.LifeExtension.com/sales/lab-tests to order your requisition forms.

After you order and receive our form, you can visit a blood-draw facility we suggest at your convenience in your area or the **Life Extension Nutrition Center** in Ft. Lauderdale.

Lab tests are available in the continental United States and Anchorage, AK, only. Not available in Maryland.

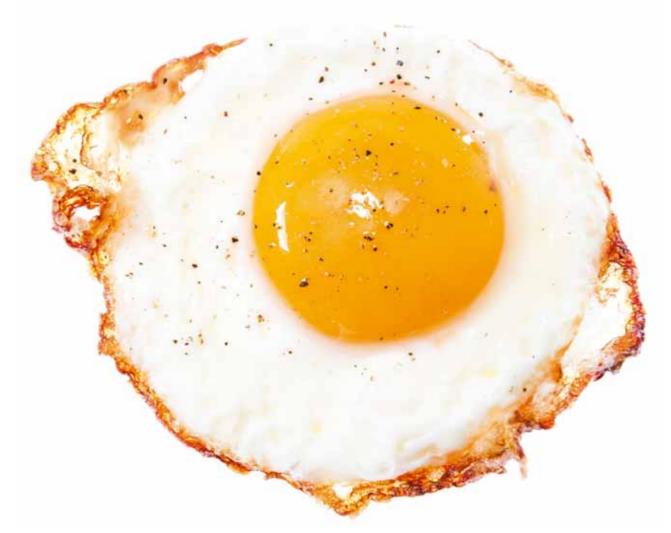
Restrictions apply in MA, NY, NJ, and RI. Kits not available in PA. The Lab Test Sale expires on **July 8, 2024**.



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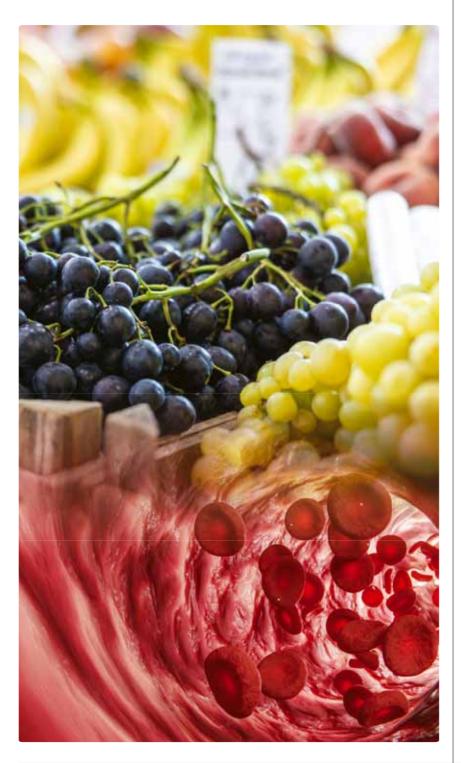
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In the News



Resveratrol Associated with Improved Endothelial Function in Diabetes Patients

A randomized, double-blind, crossover study revealed improvement in endothelial function (health of the inner arterial lining) in people with chronic kidney disease and diabetes who received resveratrol compared to those given a placebo.*

The study included 28 diabetics with stage 3 chronic kidney disease. Participants received 400 mg resveratrol per day or a placebo for six weeks. Endothelial function was measured by brachial artery flow-mediated dilation at the beginning and end of the study. Other factors, including estimated glomerular filtration rate, hemoglobin A1c, blood pressure, and nitroglycerin-mediated dilation, were also measured at these time points.

At the end of the study, participants who received resveratrol had significantly greater flow-mediated dilation compared with the placebo group. Other measured factors remained unchanged in both the resveratrol and placebo groups, which suggests that the improvement in flowmediated dilation occurred independently of changes in traditional cardiovascular risk factors.

Editor's Note: Compounds known as polyphenols, which include resveratrol, may help combat atherosclerosis because of their ability to limit oxidative stress and support a healthy inflammatory response in endothelial cells.

* Clin J Am Soc Nephrol. 2023 Oct 16.

Melatonin May Improve Athletic Recovery in Male Soccer Players

A systematic review found a decrease in oxidative stress, muscle damage, and inflammatory markers in association with melatonin use among professional male soccer players.*

Researchers reviewed eight interventional studies that compared the effects of melatonin to a placebo or no melatonin among 152 male soccer players. The studies evaluated the hormone's effect on physical performance, blood markers, and other factors. Melatonin dosages ranged from 5 to 8 mg.

Five of the studies found a reduction in oxidative stress after sports practice in association with melatonin. Exercise can cause an increase in oxidative stress.

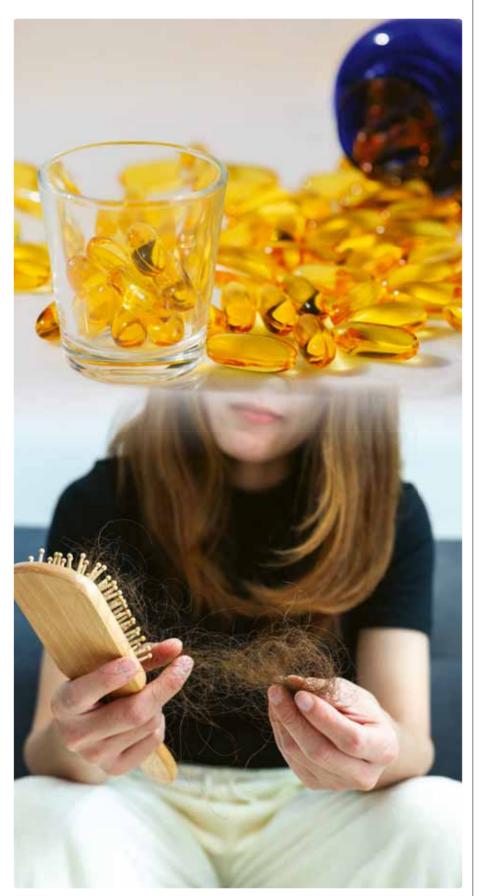
Three studies found improvement in inflammation, which can also be generated by exercise.

Muscle damage as assessed by creatine kinase and lactate dehydrogenate levels, was reduced in three of the studies.

Editor's Note: None of the studies included women in their samples.

* Nutrients. 2023;15(20):4467.





Hair Loss Linked to Vitamin D Insufficiency

The results of a systematic review and meta-analysis of 23 casecontrol studies showed an association between non-scarring alopecia (hair loss) and low serum levels of vitamin D.*

Non-scarring alopecia includes androgenetic alopecia, female pattern hair loss, alopecia areata, and other types of hair loss.

The research team found an association between having non-scarring alopecia and decreased serum 25(OH)D concentrations.

Vitamin D deficiency, defined as a 25(OH)D level of less than 20 ng/mL, was associated with more than three times the risk of androgenic alopecia, four times the risk of female pattern hair loss and a 60% greater risk of diffuse hair loss compared with healthy controls.

Editor's Note: The studies included 3,374 individuals with non-scarring alopecia and 7.296 controls.

* J Cosmet Dermatol. 2023 Nov 27.

Exercise Enhances Memory and Cognitive Function in Older Adults

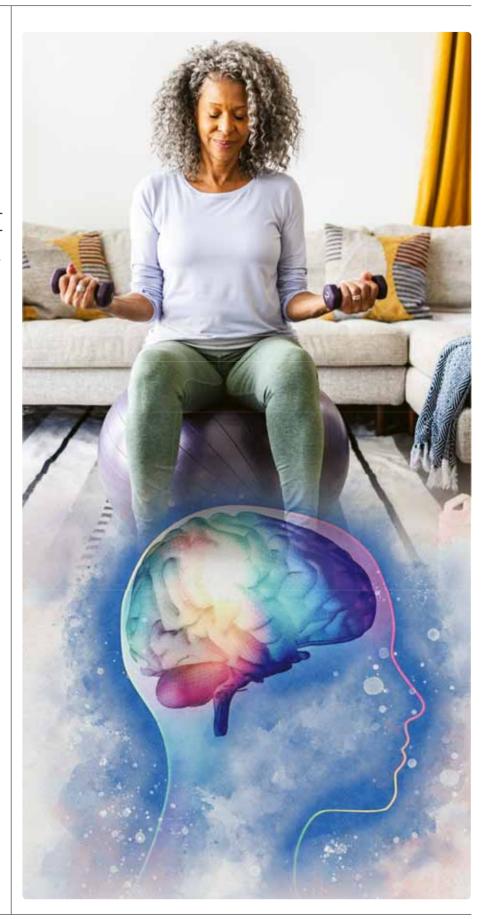
Both healthy older adults and those with mild cognitive decline benefit from improved cognitive function after exercising, according to a study published in the Journal of Alzheimer's Disease Reports.*

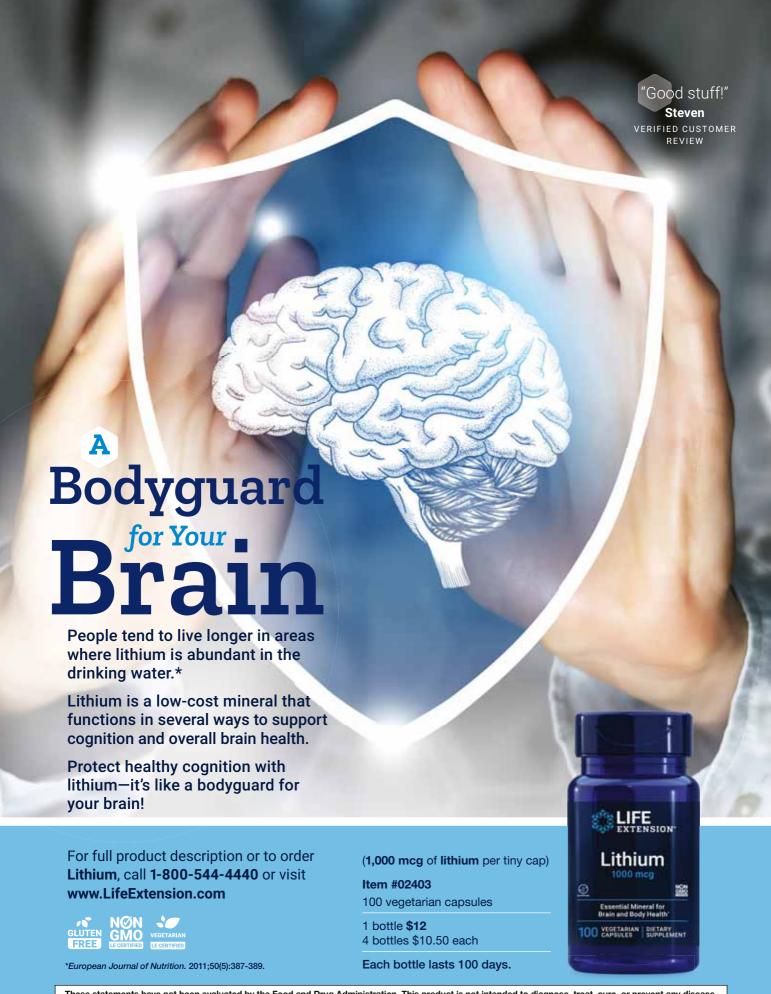
Researchers divided 33 older adults who had previously been sedentary into two groups. One group did brisk walking for 30 minutes, four times per week for 12 weeks. The other group did not exercise.

After 12 weeks, individuals in the exercise group performed better on the cognitive tests, regardless of whether they were healthy or had mild cognitive impairment.

Editor's Note: Brain scans also revealed that the exercise group had enhanced within-network and between-network connectivity of core brain networks. The researchers suggested that the increase in neural connections could explain the improvements in memory performance, both in healthy adults and in those with impaired cognitive function.

* Journal of Alzheimer's Disease Reports, vol 7, no. 1, pp. 399-413, 2023.







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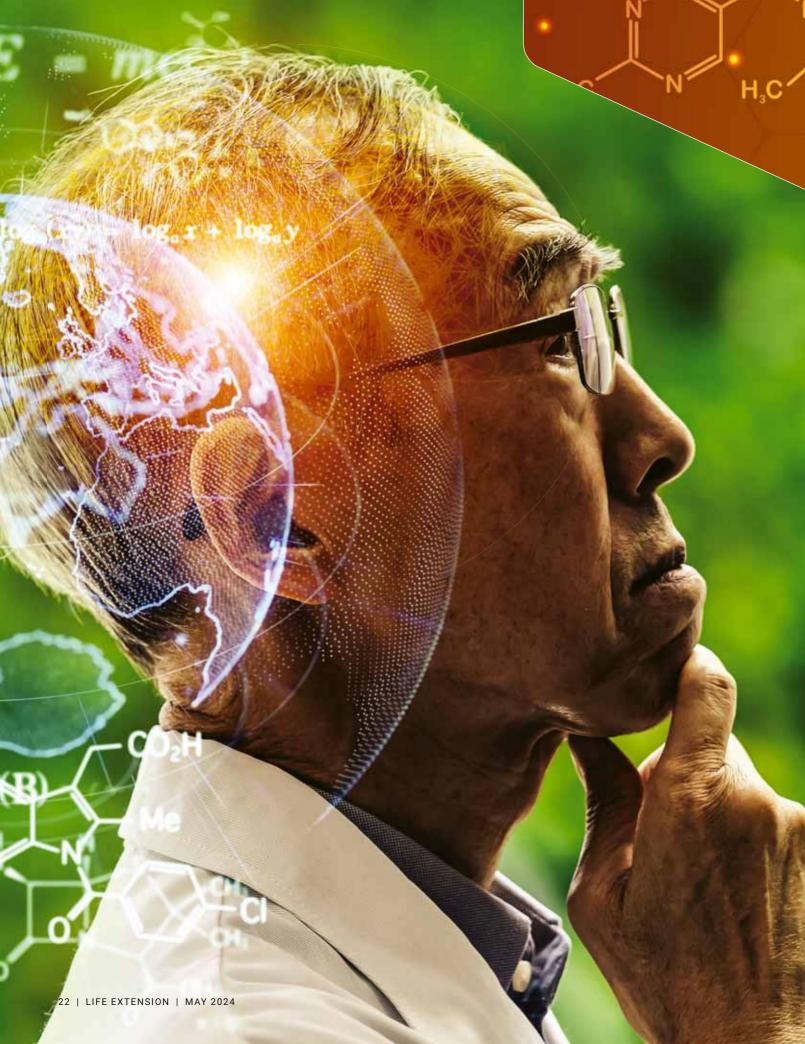
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Benfotiamine's Effects on Measures of BRAIN AGING

Many people know how damaging **high blood sugar** can be to the heart, nerves, and kidneys.

What most don't know is that it can be *especially* harmful to the brain.¹⁻⁴

Metabolic disorders, including **insulin resistance** and **high blood glucose**, are major risk factors for the deterioration of **brain function** and development of **dementia**.^{1,5-8}

Scientists have found that a compound called **benfotiamine** (fat-soluble form of vitamin B1) can not only help against the destructive effect of high blood sugar and metabolic disease but also help slow **neurodegeneration**.⁹⁻¹¹

Human studies show that benfotiamine intake can slow the progression of **Alzheimer's disease** and boost **cognitive function**. 12,13

In one **clinical study**, individuals with mild cognitive impairment or early Alzheimer's disease who took **benfotiamine** experienced **77%** *less decline* in their clinical dementia rating over a year than those receiving a **placebo**.¹²

Metabolic Abnormalities Harm the Brain

Advanced glycation end products are toxic compounds formed when the body's proteins, fats, and nucleic acids irreversibly bind with sugars.

They can potentially impact several aspects of health, including blood sugar levels that can provoke and worsen diabetes, cardiovascular, and neurodegenerative diseases. ¹⁴ Glycation happens in normal individuals but in diabetics the process is *accelerated*. ¹⁵

Not only does glycation damage brain structures, but accumulation of advanced glycation end products (AGEs) in the brain is a mechanism that has linked diabetes to cognitive impairment.^{1,16,17}

The link between metabolic disease and brain disease is so close, doctors often refer to Alzheimer's disease as **type** III **diabetes**.¹ Individuals with **diabetes**, for example, have **double the risk of developing dementia**.^{18,19}

The *higher* the blood glucose levels, the *higher* the risk. But diabetics are not the only ones in danger.

Alarmingly, even *slight elevations* in glucose increase the risk of **cognitive decline**.^{5,20-22} Individuals with **prediabetes** have an **18%** greater chance of developing dementia than those with normal glucose.⁵

Thiamine and Healthy Brain Function

Thiamine is an essential nutrient also known as **vitamin B1**. It is required for cellular metabolism.⁹⁻¹¹

Thiamine is particularly crucial in the **brain**, where it is neuroprotective and helps mitigate glycation and AGE-related tissue damage.^{9,23}

Without adequate thiamine, the brain's metabolism is compromised, and the damage done by glycation is accelerated. Studies have found that thiamine function is significantly *diminished* in those with neurodegenerative disorders such as **Alzheimer's** and **Parkinson's** disease. 9,10,24

Benfotiamine is a **fat-soluble** form of thiamine that is far *more* **bioavailable** than regular thiamine and provides additional mechanisms of brain protection.¹⁰

Brain Benefits of Benfotiamine

Benfotiamine is converted into **thiamine** in the body. Oral intake of benfotiamine has been shown to rapidly restore body levels of thiamine.⁹⁻¹¹

But benfotiamine isn't *just* a more **bioavailable** thiamine. It has been shown to offer benefits to **brain health** that are not seen with thiamine alone.^{9,10}

Animal models of neurodegenerative disease have found that benfotiamine helps prevent **brain damage** and improve **cognitive function**. 9-11,25-27

It does this in multiple ways, including by:

- Restoring thiamine levels. Benfotiamine converts into thiamine, which ensures that there is enough available for thiaminedependent enzymes to function optimally.¹⁰
- Anti-glycation activity. Glycation in the brain diminishes the function of vital proteins, contributing to disease. Benfotiamine is a potent anti-glycation nutrient. 10-12,23
- Anti-inflammatory activity. Oxidative stress and chronic inflammation in the brain accompany metabolic disease and cognitive dysfunction. Benfotiamine shields the brain from both of these threats.⁹⁻¹¹
- Improving metabolic health. Excess activity of the enzyme GSK-3 (glycogen synthase kinase-3) has been implicated as a potential target in insulin resistance and metabolic abnormalities in the brain. ^{28,29} Preclinical studies suggest benfotiamine reduces GSK-3 activity, which could help protect brain function. ^{9-11,25}

Human Studies of Benfotiamine

The neuroprotective properties of **benfotiamine** have been demonstrated in studies of adults with early cognitive impairment or Alzheimer's disease.^{12,13}

In one **placebo-controlled trial study**, individuals with a diagnosis of mild cognitive impairment or early Alzheimer's disease were randomized to receive either **300 mg** of **benfotiamine** twice daily for a year or a **placebo**. Patients were assessed on the *Alzheimer's Disease Assessment Scale-Cognitive Subscale* and *Clinical Dementia Rating score*.¹²

Those in the **benfotiamine** group experienced better preservation of **cognitive function** as well as a **77**% *less decline* in the Clinical Dementia Rating Scale than those receiving the **placebo**.

The Alzheimer's Disease Assessment Scale-Cognitive Subscale score was **43**% <u>lower</u> in the **benfotiamine** group as compared to placebo.¹² (Lower by this measure means improved cognitive function.)



In addition, while **glycation** increased by almost **10%** in the **placebo** group, it *decreased* by more than **5%** in those receiving **benfotiamine**. ¹²

An earlier pilot study recruited five patients with mild-to-moderate Alzheimer's disease, assessing their cognitive status at baseline with the Mini Mental Status Examination (MMSE).

Prior to treatment, participants also underwent brain scans to detect the toxic protein *beta-amyloid*, and its presence was confirmed in all of them. Study participants received **300 mg** of **benfotiamine** per day over the course of 18 months.¹³

At the end of the study period, they retook the MMSE and underwent another PET scan to see if the benfotiamine supplementation led to any improvements. The results were impressive. Four out of five patients demonstrated cognitive improvement on the MMSE. Over that time, the five subjects *improved* by an average of 3.2 points on the Mini-Mental State Exam (MMSE), a common screening tool for cognitive impairment.¹³

Even a one-point change in **MMSE** is considered meaningful, though the usual expected change is a <u>decrease</u>.³⁰ That makes this 3.2 point average improvement not only highly clinically meaningful but also remarkable.

These studies indicate that benfotiamine can slow cognitive deterioration and even improve cognitive function in those with early signs of dementia.

Guard Against Cognitive Decline

- Elevated blood sugar and the resulting toxic glycation increase the risk for cognitive decline and dementia. Even slightly elevated blood glucose significantly increases the risk of dementia.
- **Thiamine** is a vitamin that is crucial for energy metabolism in cells.
- While thiamine absorption is limited, its precursor benfotiamine has superior bioavailability and rapidly increases thiamine levels in the body.
- Benfotiamine improves brain cell metabolism, fights harmful glycation, and defends against cognitive decline in multiple other ways.
- In human studies, benfotiamine slows the progression of, or even reverses, cognitive decline in those with early signs of dementia.

Summary

Elevated blood sugar and toxic glycation contribute to decline in cognitive function and risk for dementia.

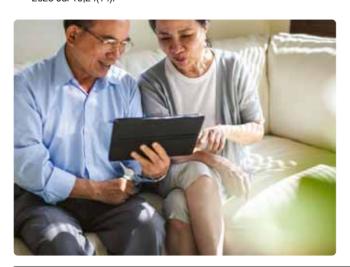
Thiamine is a vitamin that is crucial to normal metabolism in the brain. The thiamine precursor benfotiamine protects the brain against damage done by metabolic disease and high blood sugar.

Human studies of benfotiamine have demonstrated its ability to slow or reverse cognitive decline in individuals with early dementia. •

If you have any questions on the scientific content of this article, please call a Life Extension Wellness Specialist at 1-866-864-3027.

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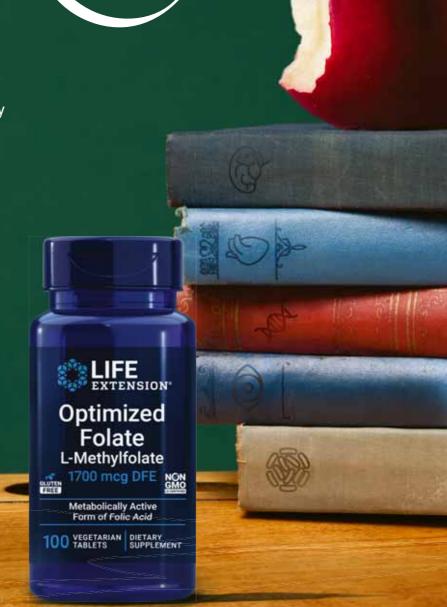
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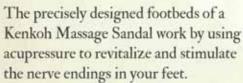




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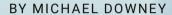
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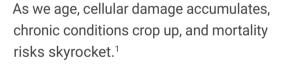
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Three NUTRIENTS that Drive HEALTHY AGING





In recent years, <u>three</u> nutrients were independently shown to **delay aging** processes and support **healthy lifespan** in cell and animal studies:

- · The amino acid taurine,
- The polyamine spermidine from wheat-germ, and
- The mineral lithium.

Seeking confirmation in **humans**, scientists turned to large epidemiological observational studies.

They found that *higher* dietary intake correlates with <u>lower</u> **mortality** and <u>reduced</u> **risk** of several **age-related conditions** (also known as increased **healthspan**).²⁻⁷

These findings suggest that supplementation with **taurine**, **lithium**, and **spermidine** may promote **healthy aging** and longevity.





Taurine

Taurine is an amino acid found in nearly all tissues.^{8,9} It is produced in small amounts by the body, but levels drop with age. In epidemiological studies, by human adulthood, **taurine** production may be *inadequate* to maintain optimal health.^{9,10}

In the large, over 25-year-long epidemiological study EPIC-Norfolk, researchers found a correlation between *higher* blood **taurine** levels and <u>lower</u> risks of various age-associated pathologies.^{11,12}

Another large population study, coordinated by the **World Health Organization** and spanning 24 populations in 16 countries, showed that *higher* urinary levels of taurine were associated with **lower mortality** rates from ischemic **stroke**.^{2,13}

In human randomized controlled studies, oral taurine intake has been shown to improve markers of health implicated in **rapid-onset aging**,¹⁴⁻¹⁷ along with supporting multiple aspects of cardiovascular health, like blood pressure,¹⁸⁻²⁰ lipid profile,¹⁸ fasting glucose,^{15,21,22} athletic performance,^{19,23} and cognition.^{17,24}

In animal studies, taurine extends lifespan. Taurine intake:11

- Increased median lifespan in worms by 23%.
- Increased median lifespan in mice by up to 12%,
- Increased life expectancy in elderly mice by up to 25%, and
- Improved multiple biomarkers of aging in middle-aged monkeys.

Spermidine

Spermidine is a polyamine (a compound having two or more amino groups) found in all living organisms and many foods, including mushrooms, legumes, corn, and whole grains.²⁵ It is commonly extracted from **wheat germ**.

Spermidine is also known to mimic the anti-aging effects of **caloric restriction**, such as support of **autophagy** (the recycling of old or damaged cell parts, often referred to as "cellular housekeeping").²⁶

In humans, levels of spermidine decline with aging.²⁷ This is a potential problem. Human epidemiological observational studies show *higher* dietary intake of spermidine is correlated with <u>lower</u> risk of chronic diseases and reduced mortality.

Spermidine is considered an age-delaying agent that helps protect against chronic disorders and reduce mortality. 6,26

Several **human** observational studies found a correlation between *higher* dietary intake of spermidine and:^{5,7,28-30}

- Lower mortality risk,
- Decreased risk of cardiovascular issues and related mortality,
- Reduced cognitive impairment risk, and
- Lower blood pressure.

Pre-clinical studies have demonstrated that spermidine may help dissolve **amyloid plaques** in the brain by **autophagy**. This may help with age-related memory decline.

A **clinical trial** of participants aged 60-100 years found that oral spermidine supplementation at doses of **0.9 - 3.3 mg/d** (wheat germ extract) improved **memory** scores as compared to **placebo**.³¹

Pre-clinical and clinical evidence has shown that functions aided by **spermidine** include:^{6,27,30,32,33}

- Metabolism of lipids,
- Regulation of cell growth,
- Lowering of inflammation levels,
- Neuroprotection,
- Cardioprotection,
- Enhanced anticancer immune response (in rodent models), and
- Autophagy.

In animal studies, **spermidine** intake <u>increases</u> median **lifespan** by:^{28,33-37}

- 3-fold in yeast,
- 30% in flies,
- 15% in worms, and
- 10% in old mice.

Lithium

Lithium is a mineral found in rocks and subsoil in some geographical areas. Some natural water sources contain small amounts of this element.

Pre-clinical evidence has shown that **lithium** can:³⁸⁻⁴³

- Inhibit GSK-3 (glycogen synthase kinase-3), an age-accelerating enzyme,
- Help maintain longer telomeres (protective caps on the ends of chromosomes tied to increased longevity),
- Regulate genes related to healthy DNA,
- Help protect against dysfunctional senescent cells that contribute to age-related disease and accelerated aging,
- · Increase activity of beneficial neurotransmitters,



Age the Healthy Way

- Animal and human studies show that higher dietary intake of the nutrients taurine, spermidine, and lithium is correlated with lower mortality, better health, and increased longevity.
- In animal studies, each of these compounds substantially increases median lifespan.
- Human data show that these nutrients inhibit a broad range of age-related diseases and promote healthy aging.
- Increase brain-derived neurotrophic factor, a signaling molecule that protects brain cells and augments their function, and
- Reduce buildup of beta-amyloid, a plaque associated with Alzheimer's disease.

Studies indicate that lithium can promote healthy overall **aging** and reduce **mortality**.

Lithium administration was shown to increase **lifespan** in both roundworms³⁸ and fruit flies.³⁹ In one study, lithium increased the lifespan of worms by an astonishing **46**%.⁴⁴

In **humans**, people taking high-dose lithium for medical reasons generally have lower **mortality** rates, including lower rates of death due to cardiovascular disease. 45,46

Other human observational data revealed that the rates of **death** from **Alzheimer's** are *higher* in areas with *low* levels of lithium in the water.⁴⁷

Two other observational studies showed that people living in areas with lithium in the drinking water tend to live longer.^{3,4}

How They Work

These three nutrients may beneficially influence the hallmarks of **aging** in multiple ways.

Taurine promotes **DNA** repair processes in the body, which help protect against mitochondrial dysfunction, inflammation, and cell senescence.^{48,49}

Spermidine helps regulate **autophagy**, helps promote anti-inflammatory activity, and suppresses harmful compounds produced by senescent cells.^{28,32,50-52}

Lithium may help prevent the shortening of **telomeres** associated with aging. By inhibiting the age-accelerating enzyme **GSK-3**, it supports cell defense and synthesis of the energy source glycogen.^{38,53}

Together, these actions may help reduce the risk of age-associated disorders and promote healthy aging.



Summary

The compounds **taurine**, **spermidine**, and **lithium** have been shown to potentially **slow the aging process**, inhibit development of age-related disease, and extend longevity.

Large observational **human** studies suggest that each of these nutrients prolongs healthy human **lifespan**.

Taken together, taurine, spermidine, and lithium may have significant mechanistic benefits on **healthy** aging. •

If you have any questions on the scientific content of this article, please call a **Life Extension**Wellness Specialist at 1-866-864-3027.

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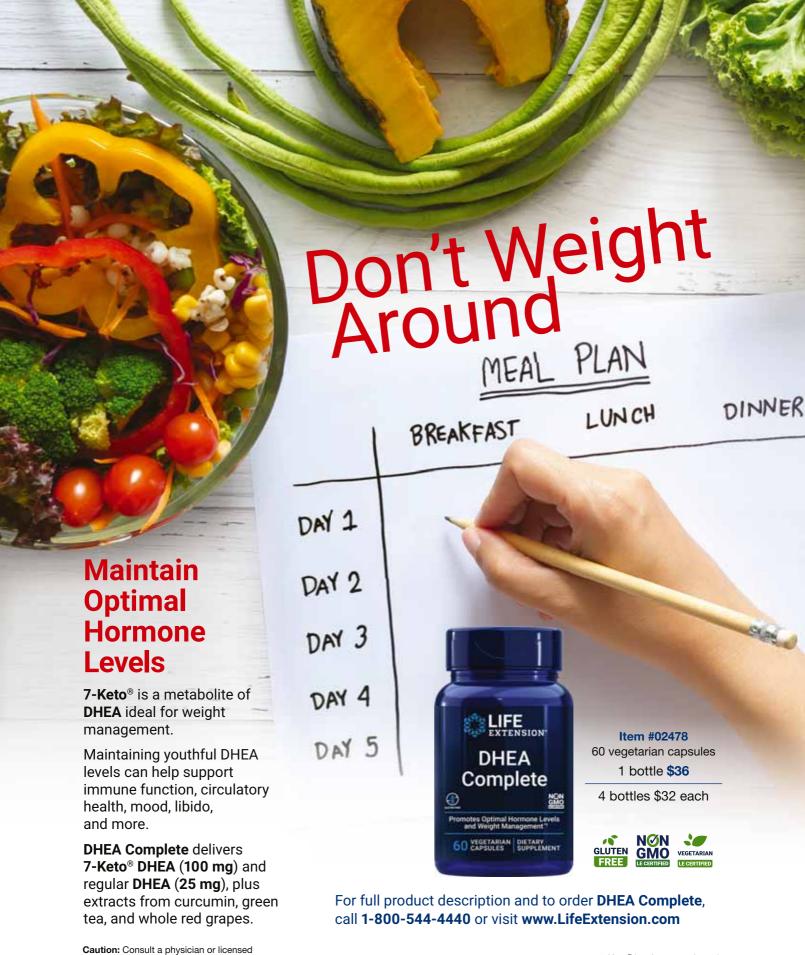
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In recent years, three nutrients have emerged as having healthy lifespanenhancing potential:

Taurine

Functions via a range of anti-aging mechanisms1-5

Lithium

People tend to live longer in areas where lithium is abundant in the drinking water.6-8

Spermidine

In epidemiological studies, higher intake correlates with longer healthspan.9-11 A clinical study showed spermidine supplementation improved memory scores.12

Consumers have used some of these nutrients for decades, albeit at lower potencies than what may be optimal.

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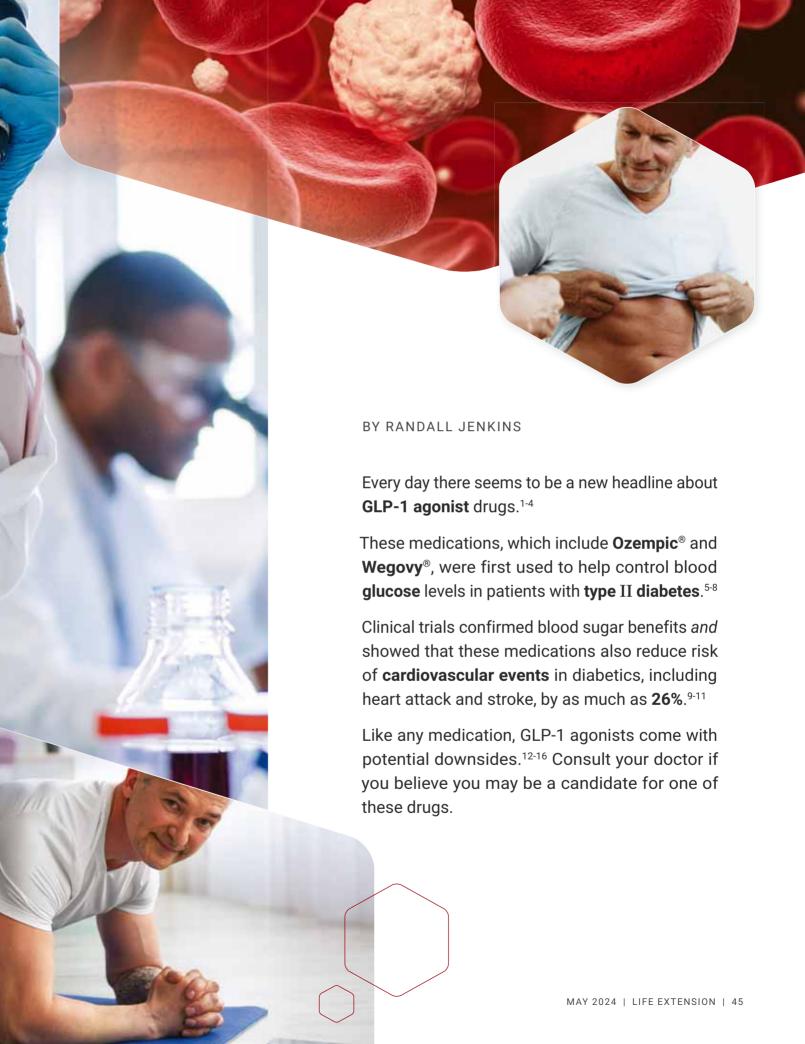
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What are GLP-1 Agonists?

GLP-1 agonists are a relatively new class of drugs. They bind to and <u>activate</u> a cellular **receptor** for the hormone **GLP-1** (glucagon-like peptide-1).

This cell **receptor** is most abundant in the insulinsecreting part of the pancreas. It is also present in the nervous system and throughout the gastrointestinal tract, as well as the cardiovascular system, kidneys, and lungs.^{17,18}

The formal name of this drug class is **GLP-1 receptor agonists**, but in common usage this has been shortened to **GLP-1 agonist**. These drugs act on the **GLP-1** hormone receptor.

First developed as **anti-diabetes** medications, in the pancreas these drugs stimulate production of **insulin**, the hormone that helps reduce blood glucose levels after a meal.^{3,19}

They also slow down the emptying of the stomach after eating and promote a feeling of fullness or satiety.¹⁹

Multiple meta-analyses of randomized controlled **clinical trials** have firmly established that GLP-1 agonist drugs are highly effective in type II diabetics for reducing blood sugar as well as hemoglobin A1C, a marker of longer-term glucose control. Some of these studies also found that these medications reduce systolic blood pressure.^{5,6,8}

What has received so much attention is that these and other studies have also found that patients taking this class of drugs have had **significant weight loss**.

In part, this is because GLP-1 agonists activate receptors in the area of the brain that controls hunger. They stimulate **satiety**, the sense of feeling full after a meal, reducing cravings and caloric intake. GLP-1 agonists are approved for use in some people with type II diabetes, and two of them are FDA-approved for weight loss in certain people and situations.

In such individuals starting a GLP-1 agonist drug, it is recommended to also adopt a reduced-calorie meal plan and to increase physical activity at the same time.

Impressive Clinical Results

GLP-1 agonists have been found to offer benefit for a variety of common chronic conditions. Here are a few highlights:

TYPE II DIABETES

Meta-analyses of placebo-controlled trials of GLP-1 agonists in patients with **type II diabetes**^{5,6,8} have consistently shown that they are effective at reducing blood glucose.⁵⁻⁸

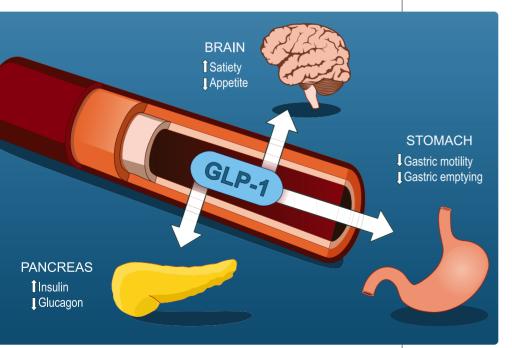
A **hemoglobin A1c (HbA1c)** test shows the average blood sugar level over two to three months. This value is higher in those with *poor* glucose control. In many clinical studies, use of a GLP-1 agonist has resulted in significantly lower HbA1c levels.

WEIGHT LOSS

The meta-analyses of randomized controlled trials in type II diabetics referenced above also found that, on average, subjects lost significant weight while taking these medications.^{5,6,8}

In a trial of <u>non-diabetic</u> adults with **obesity**,²¹ all patients received a lifestyle intervention and were randomized to receive either weekly injections of **semaglutide**—the drug in **Ozempic**® and **Wegovy**®—or placebo. The semaglutide group experienced an average of nearly *15% reduction in body weight* after 68 weeks, compared to **2.4%** in a placebo group.

This corresponds to an average loss of almost **34 pounds** in the treatment group, compared to less than **six pounds** in those who took a placebo.



REDUCTION IN **CARDIOVASCULAR EVENTS**

Both diabetes and obesity are risk factors for cardiovascular disease, which is damaging to both the heart and blood vessels. Rates of cardiovascular disease are much higher in those with poorly controlled diabetes and/ or excess body weight. It makes sense that by controlling blood glucose and reducing weight, GLP-1 agonists could also lower cardiovascular risk.

Recent analyses have started to look at these effects and here too the results are encouraging. A meta-analysis of randomized controlled trials that included over 56,000 subjects taking GLP-1 agonists for diabetes found that the drugs reduced major cardiovascular events (including stroke and heart attack) and death from cardiovascular and all causes by 12%.10

Two trials^{9,11} designed specifically to test cardiovascular effects of these drugs found that type II diabetics treated with semaglutide had a remarkable 21% and 26% reduction in risk for major cardiovascular events, including non-fatal stroke and heart attack, and death from cardiovascular causes.

To evaluate whether GLP-1 agonists could improve cardiovascular health in people without diabetes, the recent SELECT trial enrolled 17,604 nondiabetic overweight or obese adults with cardiovascular disease.²² They were randomized to receive **semaglutide** or a placebo.

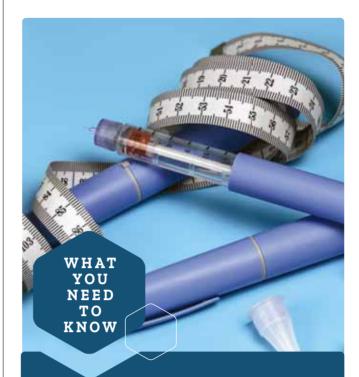
The results of this trial have not been published in the medical literature yet, so they should be assessed with caution. But the preliminary results reported in the news and on the drug company's website1,23 have exceeded expectations. Even in those without diabetes, use of this drug led to a 20% reduction in cardiovascular events and death.

These findings show that GLP-1 agonists can be game changers in improving blood glucose control and body weight and in reducing risk of cardiovascular events and death.

Need for Caution

GLP-1 agonists have been generally well tolerated in trials. However, there are potential concerns that should be considered before starting these medications. 4,14,16,24

Gastrointestinal side effects, including nausea and vomiting at higher doses, are by far most common,25 and a reason for discontinuation of treatment.^{25,26} Therefore. starting with lower doses and a slower dose titration may help minimize this risk.²⁷



The Skinny About **GLP-1** Agonists

- Drugs called **GLP-1 agonists**, which include Ozempic® and Wegovy®, have caused a buzz in the medical community and the media.
- Originally designed to reduce blood glucose levels in people with type II diabetes, GLP-1 agonists also promote significant weight loss in obese or overweight adults.
- Clinical trials have also found that GLP-1 agonists can reduce risk for major cardiovascular events, including cardiovascular-related death, heart attack, and stroke, by 20% or more.
- Gastrointestinal problems are common side effects, and safety concerns remain about long-term use. Consult a doctor if you think GLP-1 agonists may benefit you.

For those patients using GLP-1 agonist drugs for weight loss, various side effects have been reported including nausea, vomiting, abdominal pain, diarrhea, constipation, gastroenteritis, and certain incidents of pancreatitis and bowel obstruction.²⁸⁻³¹

Anyone considering using a GLP-1 agonist should consult with a doctor and weigh potential side effects against the many benefits.

Summary

GLP-1 agonists are the subject of tremendous "buzz" and exuberance in the medical community and the media.

Originally designed to help manage **type** II **diabetes**, these medications have also been found to induce significant **weight loss**.

Clinical trials show that these drugs also reduce cardiovascular events and death by 20% or more in those with type II diabetes and in overweight or obese adults without diabetes.

GLP-1 agonists can cause gastrointestinal symptoms, and concerns remain about their long-term effects, but they have tremendous potential to control blood sugar, promote weight loss, and protect cardiovascular health. •

If you have any questions on the scientific content of this article, please call a **Life Extension**Wellness Specialist at 1-866-864-3027.

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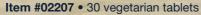
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Vitamin D plays a variety of roles in cells throughout the body.^{1,2}

One of its functions is to help <u>prevent</u> abnormal growth that could transform into **cancer**.^{1,3-5}

Some observational studies have noted a clear association of cancer incidence and vitamin D <u>deficiency</u>. 5-7

In a subset analysis of a six-year clinical trial, people taking vitamin D3 had a <u>lower</u> risk of developing metastatic or fatal cancer than those who received placebo.8

Dangers of Low Vitamin D

In a study based on the well-known **NHANES** survey of Americans and their health, more than **40%** of Americans are estimated to have <u>insufficient</u> levels of **vitamin D** in their blood.⁹

In this study, vitamin D deficiency was seen in all age groups, and was higher in women and individuals with darker skin color. In people aged **20-29 years** the rates of deficiency were even greater.⁹

The damaging effects of these kinds of <u>low</u> **vitamin D** levels are underappreciated. Vitamin D is more than a vitamin. It is also a very important **prohormone**—a kind of hormone precursor.¹⁰

Adequate **vitamin D** levels are critical for bone health, immune function, glucose metabolism, and more.¹¹ Vitamin D can also regulate the growth, development, and death of certain cells—in ways that may slow the development and progression of malignancies.¹

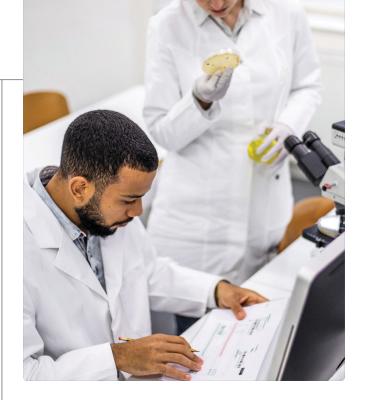
Anti-Cancer Activity

Cancer is caused by mutations that allow cells to grow and spread uncontrollably.

Vitamin D can help limit the ability of cancer cells to reproduce, grow, and spread. 1,5,12

Here are just a few of vitamin D's known anti-cancer mechanisms:1,3,5,12

- Impeding Cancer Cell Growth. Vitamin D
 helps prevent the abnormal, excessive growth of
 cancer cells by arresting its cell cycle, the process that can allow cancer cells to divide
 out of control.
- Promoting Cancer Cell Death. Apoptosis is a process by which cells shut down and die, ridding the body of aged or damaged cells.
 Cancer cells can evade apoptosis and propogate out of control. Vitamin D reverses this, promoting beneficial apoptosis.
- Starving Cancer of Nutrients. Vitamin D helps block angiogenesis, the formation of blood vessels that deliver nutrients to cancers. This reduces the nutrient supply to cancer cells and limits their growth.
- Reducing Metastasis. Aggressive cancers can invade surrounding tissue and eventually metastasize, spreading through the body.
 Vitamin D interferes with cancer cells' ability to invade and metastasize.



The Cancer, Low Vitamin D Link

Human observational studies have shown an association between <u>low</u> **vitamin D** levels and increased risk for many **cancers**. 5,13,14

Vitamin D deficiency is typically defined by the **World Medical Association (WMA)** as having a 25-hydroxyvitamin D blood level below **20 ng/mL**. ¹⁵ This is quite common in cancer patients. ^{6,7,16} Multiple observational studies have found a high rate of vitamin D deficiency in cancer patients, often in the range of **62%-67%** of patients. ¹⁷⁻¹⁹

In studies evaluating *specific* cancer types, the same pattern is seen: Those with lower circulating levels of vitamin D have the highest rates of cancer, including breast, lung, head and neck cancers, melanoma, colorectal cancers, and bladder cancers.²⁰⁻²⁸

Here are a few examples:

- In a meta-analysis of 11 observational studies, individuals with the highest circulating vitamin D levels had a 39% lower risk of developing colorectal cancer than those with lower vitamin D levels.²¹
- A combined analysis of three clinical studies found that a 25-hydroxyvitamin D blood level greater than 60 ng/mL was associated with an 80% lower risk of breast cancer compared to individuals who had vitamin D deficiency (<20ng/ml).²³
- Vitamin D may not only help reduce the risk of cancer development, but it may also impact how aggressively cancers behave.

 In one observational study of men with prostate cancer, their levels of vitamin D were found to be inversely associated with a tumor proliferation marker. This lends credence to the hypothesis that vitamin D may protect against tumor progression.²⁶

Clinical Trials

Many experts, including those at **Life Extension**, recommend daily intake of **5,000 to 8,000 IU** of **vitamin D3**. It is widely available to Americans in high potency capsules at modest cost.

Unfortunately, clinical trials evaluating vitamin D's effect on cancer have mostly used doses of **2,000 IU** or below. In people with low levels, these doses are unlikely to bring their blood levels into what many consider an optimal range (over **50 ng/mL**).

Researchers often give the <u>same</u> dose of vitamin D to <u>all</u> study participants. This is an error, because overweight/obese people and those with absorption challenges need *higher* doses of vitamin D.^{29,30} A dose that might properly elevate serum vitamin D in a normal weight person can have little effect on someone with a high body mass index (BMI). In many cases, this may mean researchers are failing to give adequate vitamin D to people who need it the most.

Even so, benefits have been shown.

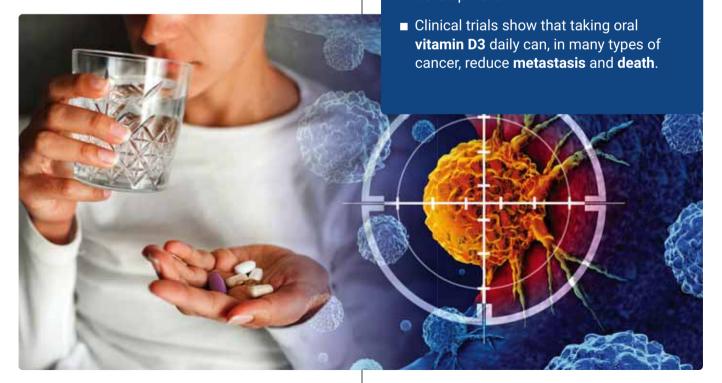
A **six-year**, multicenter clinical trial called VITAL enrolled more than **25,000** adults who were randomized to receive **2,000 IU** of **vitamin D** and / or **1 gram omega–3** fish oil supplementation, or placebo.

Participants were randomized into four groups:

- Active vitamin D and fish oil
- Active vitamin D and placebo instead of fish oil
- Placebo instead of vitamin D and active fish oil
- Placebo pills instead of either vitamin D or fish oil

Vitamin D's Role in the Fight Against Cancer WHAT YOU NEED TO KNOW

■ Low levels of vitamin D are extremely common. Observational studies have often shown that <u>low</u> vitamin D status is associated with <u>increased</u> risk of cancer development.





A significant **17**% <u>reduction</u> in advanced cancer (metastatic or fatal) incidence was found in those who received vitamin D.

When analyzed by BMI (body mass index) categories of normal weight, overweight, and obese, a significant **38%** risk <u>reduction</u> was seen in the vitamin D group that had <u>normal</u> BMI **(25-<30)**.8 This may indicate that study participants with a *higher* BMI should have received <u>more</u> vitamin D than **2,000 IU/day**.

Another double-blind, placebo-controlled trial evaluated patients with **digestive tract cancers**, including esophagus, stomach, small intestine, colon, and rectum, administering oral vitamin D supplementation at a dose of **2,000 IU/day** beginning two to four weeks after cancer surgery.³¹

This trial did not show a change in five-year relapse-free survival. But a later analysis of the trial data examined whether mutations in the **p53** tumor suppressor gene, in the tumors themselves, predicted the likelihood of a beneficial response to vitamin D supplementation.³² **P53** is the most commonly mutated gene in human cancers³³

When considering the **p53** mutation status, the results of vitamin D supplementation were quite remarkable. In people with **p53 mutations**—meaning their cells had lost an innate ability to suppress tumor cell proliferation—relapse or death was **73%** <u>less</u> likely to occur in those supplemented with vitamin D compared to placebo.

The follow-up period in this study was over **five years** and results indicate that cancer patients with **p53 mutations** may be <u>more</u> likely to respond favorably to vitamin D.

In a meta-analysis of five randomized controlled trials, vitamin D supplementation conferred a **13**% <u>lower</u> risk of cancer death. This benefit was largely attributable to daily vitamin D dosing rather than bolus dosing.³⁴

This evidence shows that vitamin D supplementation reduces cancer death, that it is especially effective in those whose tumors carry the most common cancerpromoting mutations, and that overweight and obese individuals probably require *higher* doses of vitamin D to gain these benefits.

Summary

Vitamin D has been shown to defend against the development and progression of **cancer** in multiple ways.

In many observational studies, people with *low* levels of vitamin D have often been found to have *higher* rates of many types of cancer.

Several clinical trials show that taking daily oral **vitamin D3** can help reduce the risk of cancer **metastasis** and **death**. •

If you have any questions on the scientific content of this article, please call a **Life Extension**Wellness Specialist at 1-866-864-3027.

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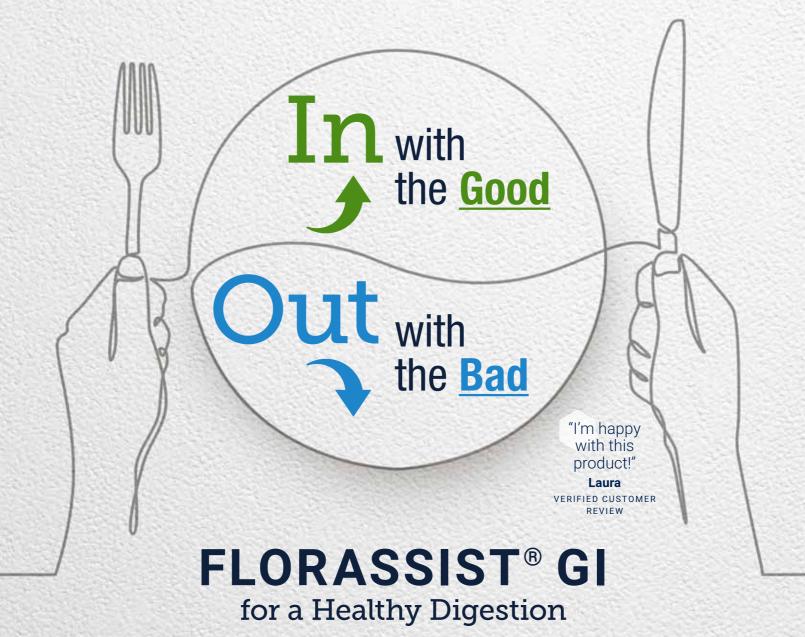
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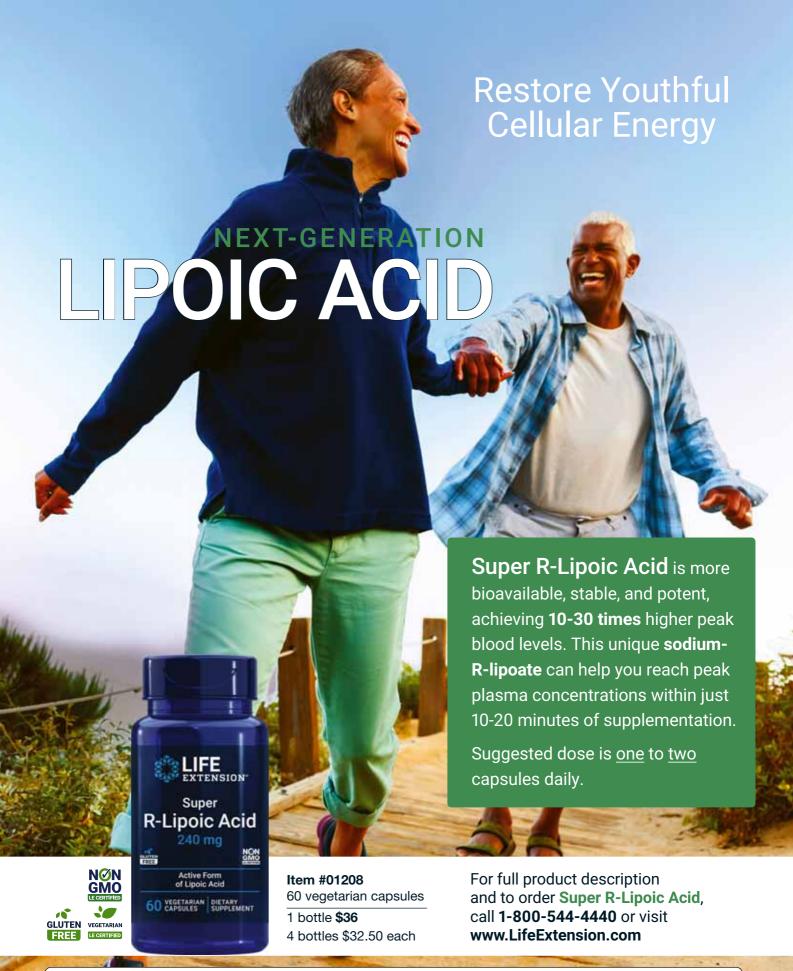
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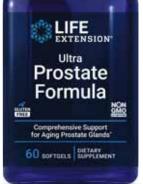


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What is PQQ?

BY LAURIE MATHENA



Mitochondria are tiny powerplants that supply energy to the cells throughout your body.¹

With age, your body produces less mitochondria, and existing ones can become damaged and dysfunctional.^{1,2}

This results in a cellular "energy shortage" that contributes to age-related problems like insulin resistance,^{2,3} cognitive decline,⁴ and macular degeneration.^{5,6}

A compound called **pyrroloquinoline quinone (PQQ)** has the potential to ward off health issues by stimulating the creation of healthy, **new mitochondria**.⁷⁻¹⁰ Additionally, animal studies have shown that boosting **PQQ** levels improves eye health, while guarding against metabolic disease, liver damage, and kidney damage.

One animal model showed that PQQ increased **lifespan** by an average of **30**%. 11,12

In a **human trial** of individuals with age-related cognitive decline, supplementing with **21.5 mg** of **PQQ** for 12 weeks resulted in improvement of composite memory, reaction time, and cognitive flexibility compared to placebo.¹³

Another study showed that supplementing with PQQ leads to improvements in attention and working memory.¹⁴

Aging and Mitochondrial Dysfunction

The body is a complex machine that requires an incredible amount of energy to run efficiently. That energy comes from energy factories called mitochondria that convert food into energy the body can use.

Healthy mitochondria divide on their own to replenish their numbers. This is a remarkable process known as mitochondrial biogenesis. Mitochondrial biogenesis is critical for protecting cells from premature aging.^{2,15,16}

But with age, this process slows. When mitochondria become damaged and die off, it creates a cellular energy shortage that accelerates aging and disease.

Mitochondrial dysfunction is especially damaging to the organs and tissues that have the greatest energy requirements, like our brain, heart, retina, pancreas, and liver.^{17,18}

Fortunately, with the right boost, mitochondria can continue to grow, repair, and replenish themselves even in later life.

That's where **PQQ** comes in.

PQQ Creates New Mitochondria

PQQ (pyrrologuinoline guinone) is a water-soluble, vitamin-like compound found in plants, yeast, and certain bacteria.

PQQ helps mitochondria in three important ways:

- By promoting the creation of healthy, new mitochondria, 16,19-24
- By clearing out defective mitochondria,19,25 which helps ensure cell survival,15 and
- By boosting the energy production of existing mitochondria. 20,26,27

In a human study, a single dose of PQQ (equaling 13 mg for an average-sized adult) led to improved measurements of urinary oxidant levels (indicators of mitochondrial efficiency).

This team of scientists also evaluated the impact of a higher daily dose of PQQ supplementation after three days (equaling about 20 mg per day for an average-sized adult) and found that measurements

of inflammation (such as C-reactive protein and interleukin-6) were decreased.8

In a study of healthy men who completed a six-week aerobic training program, those given 20 mg of PQQ daily more than doubled their PGC-1 alpha levels (mitochondrial-generating protein) as compared to placebo.7

Brain Protection

The brain uses more energy than any other organ in the body,28 so it is one of the hardest hit when mitochondrial function declines.

Aging, dysfunctional mitochondria are known as contributors to agerelated brain disorders, including Alzheimer's and Parkinson's disease. 15,23,29

By helping mitochondria perform more efficiently and by promoting the development of new mitochondria, PQQ may help prevent these and other disorders. 19,23,29,30

In a human study, participants aged 20-65 years received 20 mg of PQQ or placebo for 12 weeks. Supplementing with PQQ led to improvements in memory test scores after eight weeks. In older adults, at the end of 12 weeks, further improvements in complex and verbal memory were observed.31

Researchers have found preclinical evidence that PQQ may slow the progression of both conditions—or possibly even prevent them altogether. 30,32-37

That's because, in addition to boosting mitochondrial function, PQQ has been found to prevent the accumulation of abnormal proteins (such as beta-amyloid and alpha-synuclein) associated with neurodegeneration.32-34





Prevent Metabolic Issues

Mitochondrial dysfunction contributes to metabolic issues like obesity, insulin resistance, and type II diabetes, as well as cardiovascular disease.24,38,39

PQQ helps improve many factors that contribute to metabolic syndrome. PQQ affects insulin signaling via multiple pathways. That is why studies suggest that PQQ may be beneficial in insulin resistance and type II diabetes.41

In an animal study of metabolic syndrome, obese rats treated with PQQ for five weeks experienced the following benefits (compared to the untreated group):24,40

- Significant improvements in blood sugar control, insulin levels, and insulin sensitivity,
- Reductions in harmful inflammatory cytokines, and
- Healthier blood lipid levels, including total cholesterol, triglycerides, LDL ("bad") cholesterol, and HDL ("good") cholesterol.

Kidney and Liver Health

Two serious side effects of metabolic issues like obesity and diabetes are kidney and liver damage. PQQ has considerable potential for remedving metabolic diseases including diabetes, nonalcoholic fatty liver disease (NAFLD), and chronic kidney disease.41

In rodent models of diabetes. the kidneys show signs of significant damage and fibrosis (scarring). Kidney function is decreased and there are signs of oxidative stress and inflammation seen in the tissues.42,43

In an animal model of diabetes, giving animals PQQ significantly reduced the structural damage to their kidneys, while improving kidney function.42,43

And in a preclinical study of NAFLD (nonalcoholic fatty liver disease), PQQ protected the liver from fat accumulation by supporting the creation of new mitochondria, improving lipid metabolism, and boosting antioxidant protection.44

Eve Protection

Abnormal mitochondrial function is often found in three of the leading causes of blindness: macular degeneration, glaucoma, and diabetic retinopathy.6,9

Cell studies from eyes of human donors with macular degeneration show under-functioning mitochondria in the retinal pigment epithelium. Researchers used PQQ to treat retinal pigment epithelial cells from human organ donors who had agerelated macular degeneration.9

In half of the tissue samples, administering PQQ led to a 50%-350% improvement in the mitochondrial function of these critical retinal cells.

The treated cells also had a 59% increase in the production of ATP, which is the form of energy that the cells use

Summary

PQQ helps promote healthy aging by promoting the production of healthy, new mitochondria.

Enhancing the body's cellular energy with PQQ has the potential to improve memory and eye health, while protecting against metabolic, liver, and kidney disease.

PQQ also has the potential to increase lifespan.

Taking 10 to 20 mg of PQQ daily is a practical way to support healthy aging.

If you have any questions on the scientific content of this article, please call a Life Extension Wellness Specialist at 1-866-864-3027.

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Check Your Health with Lab Tests

DR. SCOTT FOGLE, ND



Lab tests can assess a wide array of health concerns and point the way to potential solutions.

In this interview, Dr. Scott Fogle, ND, discusses some of the best tests available to help you and your physician understand your current health status.

LE: With the annual lab test sale happening, people are wondering: What are some must-have tests Life Extension offers?

Dr. Fogle: We work with nine different labs, which provide us with amazing options. Our most popular tests remain our Male and Female Panels. If you got those tests from a commercial lab, you would pay around \$2,500, but during our sale we offer them at \$224. These are valuable, comprehensive tests.

LE: What's one of *your* favorite tests?

Dr. Fogle: The GI360 Stool Analysis for gastrointestinal (GI) issues looks for dysbiosis of your gut flora and the major pathogens that cause trouble in your gut. It also offers an excellent assessment of your digestive abilities.

LE: Is that test different from the popular **SIBO Home Breath Kit**?

Dr. Fogle: Yes, the Gl360 is a stool test that takes a broad and comprehensive look at your Gl health. The SIBO kit is a breath test looking for bacteria that have overgrown and proliferated in your small intestine. This bacterial overgrowth can cause gas, bloating, diarrhea, constipation, and cramping.

LE: Food allergies also affect your GI health. Can you test for those?

Dr. Fogle: Food allergies and sensitivities can cause systemic inflammation, headaches, migraine, foggy brain, musculoskeletal problems, fatigue, skin issues, mood disturbances, and more. We offer both food allergy profiles and food sensitivity panels. Food allergies are tested via IgE antibody reactions, while sensitivities are tested via IgG and IgA antibodies.

Our Advanced Food Sensitivity Panel tests for IgG sensitivities to 240 foods and is our most popular choice during the sale. For those suspecting numerous possible food reactions, our Elite Food Sensitivity Panel adds IgA testing for a closer look at sensitivities to those foods.

LE: More news keeps coming out about environmental pollutants and toxins. What tests can help identify them in our bodies?

Dr. Fogle: We offer an **Environmental Pollutants Profile**, a convenient at-home urine test that reveals levels of harmful toxins such as xylene, toluene, benzene, trimethylbenzene, styrene, phthalates, parabens, and MTBE. For harmful heavy metals we offer the **Toxic Metals Panel** fecal test, which looks at the worst offenders including arsenic, lead, mercury, uranium, and many more. Annual testing for toxin exposure is a good idea.

LE: What **new tests** are being offered during the annual sale?

Dr. Fogle: Let me start with hair loss. It's not just a genetic issue; you can test for medical conditions that contribute to or accelerate it. We created two panels, a Basic and Comprehensive Hair Loss Panel. The basic panel looks at the most common causes, such as excessive DHT (dihydrotestosterone), which has a negative effect on hair follicles. It also measures thyroid-stimulating hormone to check thyroid health, low ferritin (an iron marker) for anemia, C-reactive protein for excessive inflammation, and low or high cortisol related to acute or chronic stress, all of which can be factors in hair loss.

LE: What does the Comprehensive Hair Loss Panel offer?

Dr. Fogle: It includes everything in the basic panel plus additional tests that delve deeper. It checks antinuclear antibodies (ANA) as a general autoimmune screening test and has more advanced thyroid testing to look for hormone imbalances or significantly high or low levels.

LE: You mentioned that **stress** can contribute to hair loss?

Dr. Fogle: Yes. The good news is that often that type of hair loss can be recovered, and tests provide clues of current or ongoing stress. The stress hormone cortisol rises during acute stress, and low levels can point to chronic stress and implication in hair loss. Also, stress can cause an increase in what's known as reverse T3 hormone levels. The Reverse T3 blood test and Cortisol blood or urine tests can indicate various stages and degrees of stress.



LE: Do you have other **new tests**?

Dr. Fogle: People are concerned about mold at home, work, or school. We finally have a lab that can deliver good testing for mold and mycotoxins (the toxins molds secrete). We are now offering a Mold Exposure Mycotoxin urine test to see what mycotoxins are present in the urine, indicating the body's exposure to mycotoxins from food and environment. We also offer an EMMA (Environmental Mold & Mycotoxin Analysis) test that checks for mold and mycotoxins in a suspected environment by swabbing dust from any area of concern.

LE: How dangerous are mycotoxins?

Dr. Fogle: They are associated with a wide range of symptoms and diseases including immune dysfunction and even cancer. Common complaints are respiratory and sinus concerns, cognitive dysfunction, neurological issues, significant fatigue, changes in mood, and more.

LE: Testing urine can identify which mycotoxins may be causing these symptoms?

Dr. Fogle: Correct. The **Mold Exposure Mycotoxin** test checks for exposure to **16** of the worst-of-theworst mycotoxins. Knowing which one is most prevalent in your body is helpful to identify foods to avoid or which mold may be lurking behind the walls in your home.

LE: Are foods or moldy environments more likely sources of mycotoxins?

Dr. Fogle: Both can be, but most issues stem from water-damaged buildings. This makes it important



to use the **EMMA** test along with the mycotoxin urine test. Dust is collected from a suspected location to directly examine the mold and mycotoxins in that environment. EMMA results can be compared to urinary mycotoxin results to see if your body's mycotoxin exposure aligns with the mold and mycotoxins in a building. Doing both tests provides a more complete picture of mold exposure, which can help confirm the source of mycotoxins in the urine.

LE: Do different mycotoxins cause different health issues?

Dr. Fogle: Yes, some specific symptoms are related to different mycotoxins. For example, ochratoxin A from the common **Aspergillus** mold family is associated with fatigue, dermatitis, and

irritated bowel issues. Satratoxins from **black mold** are associated with fatigue, breathing issues, and nervous system dysregulation. Black mold is difficult to directly test for because it is heavy and sticky and doesn't circulate in the air easily. But what do circulate are the satratoxins it produces, and now we can test for those.

Dr. Scott Fogle is Executive Director of Clinical Information and Laboratory Services at **Life Extension**.

If you have any questions on the scientific content of this article, please call a **Life Extension** Wellness Specialist at 1-866-864-3027.



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MYELOPEROXIDASE (MPO)* (LC123006)

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The myeloperoxidase (MPO) test measures levels of an enzyme that oxidizes low-density lipoprotein (LDL) cholesterol, which could lead to increased arterial plaque formation.

OMEGA-3 INDEX COMPLETE* (LC100066)

Beneficial for everyone! People not taking omega-3/fish oil should check their baseline Omega-3 Index to see if it is in the desirable or concerning range. Those taking Omega-3/fish oil supplements should take the test to see if they need to adjust their dosage. You want to target a range of 8%-12% for your Omega-3 Index score.

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PERSONALIZED AMINO ACID HEALTH ASSESSMENT* (LC100090) \$299 \$224,25

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Know your numbers



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including hormones, thyroid, inflammation, and more! • E2 • Testosterone DHEA-s • DHT • TSH • fT4 • fT3 • rT3 • cortisol • insulin • ferritin • CRP

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MYCOTOXIN URINE TEST* (LC900007)

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Concerned about mold exposure? This convenient at-home urine test detects exposure to 16 common mycotoxins (toxins produced by mold).

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MYCOTOXIN ANALYSIS* (LC900008)

\$399 \$299.25

Worried about mold in your home or office? Check for mold and mycotoxins by swabbing a dusty area in your home!

BODY AND HOME MOLD &

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Combine the two tests above for additional savings to check if the mycotoxins in your body are the same as those in your home! WHOLE-BODY HEALTH

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- Reverse T3 IGF-1 SHBG Free and Total PSA Vitamin D 25-OH hs-CRP Ferritin
- TIBC Homocysteine Insulin Hemoglobin A1c Cortisol ApoB Magnesium

MALE BASIC HORMONE PANEL (LC100012)

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\$56.25

FEMALE ELITE PANEL (LC100017)

\$431.25 \$575 CBC/Chemistry/Lipids Panel • Free and total Testosterone • Estradiol • Total Lipids

- Estrone DHEA-S Progesterone Pregnenolone DHT FSH LH TSH Free T3
- Free T4 Reverse T3 IGF-1 SHBG Vitamin D 25-OH hs-CRP Ferritin TIBC
- Homocysteine
 Insulin
 Hemoglobin A1c
 Cortisol
 ApoB
 Magnesium

FEMALE BASIC HORMONE PANEL (LC100013)

\$56.25

DHEA-S • Estradiol • Total and Free Testosterone • Progesterone

WEIGHT LOSS PANEL-COMPREHENSIVE (LC100028) CBC/Chemistry/Lipids Panel • DHEA-S • Free and Total Testosterone

- Estradiol Progesterone Cortisol TSH Free T3 Free T4 Reverse T3 Insulin
- Hemoglobin A1c Vitamin D 25-hydroxy Ferritin C-reactive protein (high sensitivity)
- **HEALTHY AGING PANEL-COMPREHENSIVE (LC100026)** \$249 \$186.75 CBC/Chemistry/Lipids Panel • C-reactive protein (high sensitivity) • Vitamin B12
 - Folate Homocysteine Vitamin D 25-hydroxy Hemoglobin A1c TSH Free T3
 - Free T4 Ferritin Urinalysis ApoB Insulin

COMPLETE HORMONE &

METABOLITES PROFILE (HUMAP)*† (LC900001)

Comprehensive assessment of hormones and their metabolites. This urine test helps assess how your body metabolizes hormones, providing insight into endogenous hormone secretion, supplemental hormone utilization, enzyme activity, oxidative stress,

CBC/CHEMISTRY/LIPIDS PANEL

These CBC/Chemistry/Lipids Panel tests are included in the popular Male and Female Panels, and other panels on this page so you don't have to order them separately.

CARDIOVASCULAR RISK

Total Cholesterol • HDL Cholesterol • LDL Cholesterol Triglycerides Cholesterol/ HDL Ratio • Estimated CHD Risk • Glucose

LIVER FUNCTION

AST (SGOT) • ALT (SGPT) • LDH • Total Bilirubin • Alkaline phosphatase **KIDNEY FUNCTION**

BUN • Creatinine • BUN/Creatinine Ratio • Uric Acid **BLOOD PROTEINS**

Total Protein • Albumin • Globulin • Albumin/Globulin Ratio **BLOOD COUNTS**

Red Blood Cell Count • White Blood Cell Count • Eosinophils Neutrophils (Absolute) • Lymphs (Absolute) • Eos (Absolute) Baso (Absolute) • RDW • Monocytes (Absolute) • Monocytes Lymphocytes • Platelet Count • Hemoglobin • Hematocrit

MCV • MCH • MCHC • Neutrophils **BLOOD MINERALS**

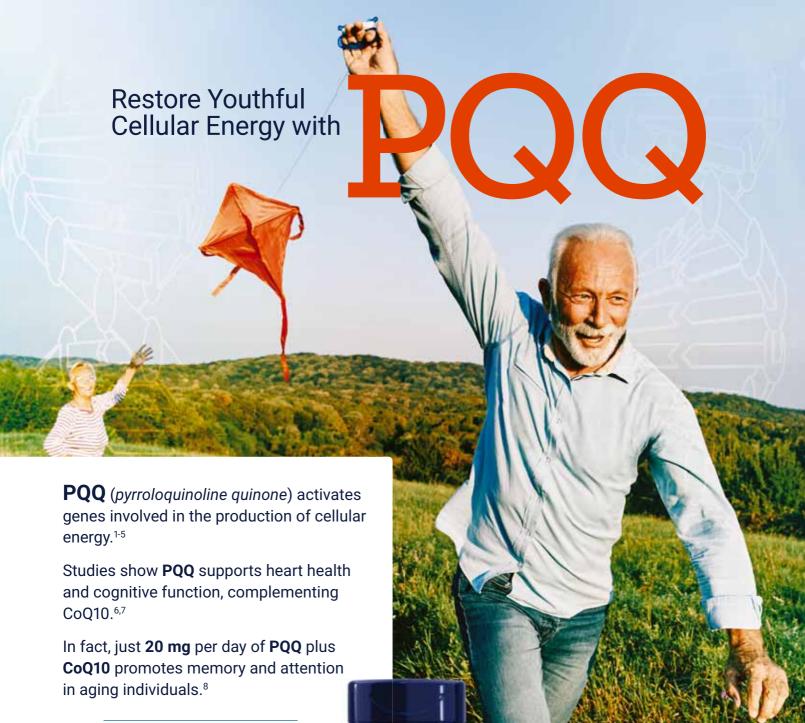
Calcium • Potassium • Sodium • Chloride • Iron

The sale price for the CBC/Chemistry/Lipids Panel alone is \$26.25. (LC381822)

This is NOT a complete listing of LE lab test services. Call 1-800-208-3444 for additional information.

- This test is packaged as a kit.
- Not available in NY.

Lab tests available in the continental United States and Anchorage, AK only. Restrictions apply in NY, NJ, RI, and MA. Not available in MD. Kits not available in PA.



This formulation contains 20 mg of PQQ per capsule, which is the recommended daily dose.

For full product description and to order PQQ or any other **PQQ-containing formulas,** call 1-800-544-4440 or visit www.LifeExtension.com

Item #01647 • 30 vegetarian capsules

1 bottle **\$24** • 4 bottles \$18 each









Also available are **10 mg PQQ caps** (Item #01500) and 100 mg Super Ubiquinol CoQ10 with PQQ (Item #01733).

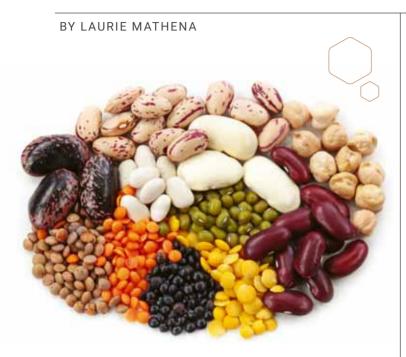
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30 VEGETARIAN DIETARY

Beans



Some foods deserve the title of superfood because they help keep you healthy. Beans deserve the title because they could help you live longer.

Beans come in a number of varieties. The most popular include garbanzo, navy, red, white and black beans.

Dan Buettner is an author who has spent years researching Blue Zones, which are areas of the world where people routinely live past 100 years old.

Buettner said, "In every Blue Zone I have visited, beans and other legumes were—and still are—a major component of the daily diet."

Beans are considered a legume plant.

Indeed, research has shown that every 20-gram increase in dietary intake of legumes per day reduces risk of mortality by 8%!1 This finding led the researchers to call beans "the most important dietary predictor of survival in older people of different ethnicities."

Every variety of bean has its own unique nutrient profile. so eating a variety is best. They are a good source of protein and fiber, while being a low glycemic-index carbohydrate.

Black beans, for example, provide 14.5 grams of protein and 16.6 grams of fiber in one cup.2

Because beans are high in fiber, they can help stabilize blood sugar levels.

In an animal study, consuming a variety of beans for three weeks was shown to improve gut health by boosting beneficial bacteria and promoting intestinal barrier function.3

In a study that followed 9,632 people for an average of 19 years, results showed that consuming legumes four times per week was associated with a 22% lower risk of coronary heart disease and an 11% lower risk of cardiovascular disease.4

Another study showed that consuming just ½ cup of beans per day was associated with a 38% lower risk of heart attack.5

In a randomized controlled trial of patients with metabolic syndrome participants were randomized to receive meals with black beans, fiber, or antioxidants. Blood samples were taken before and five hours after a meal. Insulin was lowered after a black bean meal as compared to meals with fiber or antioxidants.6

In a review of clinical trials, type II diabetics who consumed legumes on a regular basis showed reductions in fasting blood glucose, A1c, fasting blood insulin, and twohour post-prandial blood glucose.7

Beans and legumes—whether dry or canned—provide a low-cost way to include more superfoods in your diet. These versatile foods can be added to casseroles, tossed into salads, made into a soup, added to tacos, or served as traditional beans and rice. •

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02397

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ACTIVE LIFESTYLE & FITNESS		01540	DMAE Bitartrate
01529	Creatine Capsules	02006	Dopa-Mind™
02020	Super Carnosine	02413	Dopamine Advantage
02023	Tart Cherry with CherryPURE®	01658	Ginkgo Biloba Certified Extract™
02186	Wellness Bar - Chocolate Brownie	01527	Huperzine A
02518	Wellness Bar - Peanut Butter Chocolate Chip	00020	Lecithin
02246C	Wellness Code® Advanced Whey	02101	Memory Protect
	Protein Isolate Vanilla	00709	Migra-Eeze [™]
02221	Wellness Code® Muscle Strength & Restore Formula	01603	Neuro-Mag® Magnesium L-Threonate Caps
02127	Wellness Code® Plant Protein Complete &	02520	Neuro-Mag® Magnesium L-Threonate Gummies
	Amino Acid Complex	02032	Neuro-Mag® Magnesium L-Threonate Powder
02261C	Wellness Code® Whey Protein Concentrate	00888	Optimized Ashwagandha
	Chocolate	01676	PS (Phosphatidylserine) Caps
02260	Wellness Code® Whey Protein Concentrate Vanilla	02406	Quick Brain Nootropic
02243C	Wellness Code® Whey Protein Isolate Chocolate	01327	Vinpocetine
02242	Wellness Code® Whey Protein Isolate Vanilla	CHOL	ESTEROL MANAGEMENT
AMIN	O ACIDS	01828	Advanced Lipid Control
00038	Arginine Ornithine Powder	01910	CHOL-Support™
01253	Branched Chain Amino Acids	01304	Theaflavins Standardized Extract
01233	Carnosine	00372	Vitamin B3 Niacin Capsules
01623	D,L-Phenylalanine Capsules	DIGE	STION SUPPORT
01624	L-Arginine Caps	02412	Bloat Relief
01532	L-Carnitine	02412	Easy Fiber
00345	L-Glutamine	02314	Effervescent Vitamin C - Magnesium Crystals
00141	L-Glutamine Powder	02021	Enhanced Super Digestive Enzymes
01678	L-Lysine	02021	Enhanced Super Digestive Enzymes and Probiotics
01827	Taurine	02022	EsophaCool™
00133	Taurine Powder	02033	Esophageal Guardian
	D PRESSURE & VASCULAR SUPPORT	01706	Extraordinary Enzymes
		02100	Gastro-Ease™
01824	Advanced Olive Leaf Vascular Support		GY MANAGEMENT
02004	Arterial Protect		
02497	Endothelial Defense TM Pomegranate Plus		Adrenal Energy Formula • 60 veg capsules
56885	Natto Activ Cardiovascular & Respiratory Support NitroVasc [™] Boost		Adrenal Energy Formula • 120 veg capsules
02320		00972	D-Ribose Powder
01953	Pomegranate Complete	01473	D-Ribose Tablets
00956	Pomegranate Fruit Extract Triple Action Placed Procesure AM/PM	01900	Energy Renew
02024	Triple Action Blood Pressure AM/PM VenoFlow™	01544	Forskolin
02102		01805	Ginseng Energy Boost
BONE	HEALTH	01869	Mitochondrial Basics with PQQ
01726	Bone Restore Calcium Supplement	01868	Mitochondrial Energy Optimizer with PQQ
02123	Bone Restore Calcium Supplement Chewable Tablet	01904	NAD+ Cell Regenerator™ • 100 mg, 30 veg capsules
02416	Bone Restore Elite with Super Potent K2	02344	NAD+ Cell Regenerator™ 300 mg, 30 veg capsules
01727	Bone Restore Calcium Supplement w/ Vitamin K2	02348	NAD+ Cell Regenerator™ and Resveratrol Elite™
01725	Bone Strength Collagen Formula	01500	PQQ Caps • 10 mg, 30 vegetarian capsules
01963	Calcium Citrate with Vitamin D	01647	PQQ Caps • 20 mg, 30 vegetarian capsules Rhodiola Extract
01506C	Dr. Strum's Intensive Bone Formula	00889	
02417	Mega Vitamin K2	02003	Thyroid Support Complex
01476	Strontium Caps	EYE	HEALTH
BRAIN	N HEALTH	01923	Astaxanthin with Phospholipids
01524	Acetyl-L-Carnitine	02323	Digital Eye Support
01974	Acetyl-L-Carnitine Arginate	01514	Eye Pressure Support with Mirtogenol®
02419	B12 Elite	01992	MacuGuard® Ocular Support with Saffron
02510	Brain Fog Relief	01993	MacuGuard® Ocular Support with
01659	CDP Choline		Saffron & Astaxanthin
02321	Cognitex® Alpha GPC	01873	Standardized European Bilberry Extract
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Sesame Lignans & Olive Extract • 60 softgels	00302 Pregnenolone • 50 mg, 100 capsules
01988 Super Omega-3 Plus EPA/DHA Fish Oil,	00700 Pregnenolone • 100 mg, 100 capsules
Sesame Lignans, Olive Extract, Krill & Astaxanthin	01468 Triple Action Cruciferous Vegetable Extract
01982 Super Omega-3 EPA/DHA Fish Oil,	01469 Triple Action Cruciferous Vegetable Extract
Sesame Lignans & Olive Extract • 120 softgels	and Resveratrol
01985 Super Omega-3 EPA/DHA Fish Oil, Sesame Lignans	IMMUNE SUPPORT
& Olive Extract • 60 enteric coated softgels	02302 Bio-Quercetin®
01984 Super Omega-3 EPA/DHA Fish Oil, Sesame Lignans	02410 Black Elderberry + Vitamin C
& Olive Extract • 120 enteric coated softgels	02433 Echinacea Elite
01986 Super Omega-3 EPA/DHA Fish Oil, Sesame Lignans	01961 Enhanced Zinc Lozenges
& Olive Extract • 240 softgels	02425 Immune Packs with Vitamin C & D,
01812 Provinal® Purified Omega-7	Zinc and Probiotic
01640 Vegetarian DHA	02005 Immune Senescence Protection Formula™
FOOD	01681 Lactoferrin (Apolactoferrin) Caps
	02426 Mushroom Immune with Beta Glucans
02008 California Estate Extra Virgin Olive Oil	
02170 Rainforest Blend Decaf Ground Coffee	
02169 Rainforest Blend Ground Coffee	01394 Optimized Garlic
02171 Rainforest Blend Whole Bean Coffee	01309 Optimized Quercetin
GLUCOSE MANAGEMENT	01811 Peony Immune
01503 CinSulin® with InSea ^{2®} and Crominex® 3+	01708 Reishi Extract Mushroom Complex
01620 CoffeeGenic® Green Coffee Extract	01906 Standardized Cistanche
02122 Glycemic Guard™	01097 Ultra Soy Extract
00925 Mega Benfotiamine	01561 Zinc Lozenges
01803 Tri Sugar Shield®	INFLAMMATION HEALTH SUPPORT
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01066 Aspirin (Enteric Coated)	Turmeric Extract, Ginger & Turmerones
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01700 Cardio Peak™	02310 Black Cumin Seed Oil and Curcumin Elite™
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02508 Omega-3 Fish Oil Gummy Bites	30 veg capsules
02018 Optimized Carnitine	02407 Curcumin Elite™ Turmeric Extract
01949 Super-Absorbable CoQ10 Ubiquinone with	60 veg capsules
d-Limonene ∙ 50 mg, 60 softgels	01804 Cytokine Suppress® with EGCG
01951 Super-Absorbable CoQ10 Ubiquinone with	02223 Pro-Resolving Mediators
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01929 Super Ubiquinol CoQ10	01203 Specially-Coated Bromelain
01427 Super Ubiquinol CoQ10 with Enh Mitochondrial	00407 Super Bio-Curcumin® Turmeric Extract
Support™ • 50 mg, 30 softgels	JOINT SUPPORT
01425 Super Ubiquinol CoQ10 with Enh Mitochondrial	
Support™ • 50 mg, 100 softgels	02404 Arthro-Immune Joint Support
01437 Super Ubiquinol CoQ10 with Enh Mitochondrial	02238 ArthroMax® Advanced NT2 Collagen™
Support™ • 100 mg, 30 softgels	& AprèsFlex®
01426 Super Ubiquinol CoQ10 with Enh Mitochondrial	02138 ArthroMax® Elite
Support™ • 100 mg, 60 softgels	00965 Fast-Acting Joint Formula
01431 Super Ubiquinol CoQ10 with Enh Mitochondrial	02430 Fast Acting Relief
Support™ • 200 mg, 30 softgels	00522 Glucosamine/Chondroitin Capsules
01733 Super Ubiquinol CoQ10 with PQQ	02420 Glucosamine Sulfate
01859 TMG Liquid Capsules	02424 Joint Mobility
00349 TMG Powder	01600 Krill Healthy Joint Formula
HORMONE BALANCE	00451 MSM (Methylsulfonylmethane)
	02231 NT2 Collagen™
00454 DHEA • 15 mg, 100 capsules	
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00335

00882

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02304C Youthful Collagen 02252 Youthful Legs

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02029 Ultra Prostate Formula

01373

01940

		02075	Comma E Miyad Tooonharal Enhanced with
PET CARE		02075	Gamma E Mixed Tocopherol Enhanced with Sesame Lignans
01932	Cat Mix	02070	Gamma E Mixed Tocopherol & Tocotrienols
01931	Dog Mix	01913	High Potency Optimized Folate
02523	DOG Hip & Joints	01674	Inositol Caps
02524	DOG Multivitamin & Probiotics	02244	Liquid Vitamin D3 • 50 mcg (2000 IU)
02522	DOG Skin & Coat	02232	Liquid Vitamin D3 (Mint) • 50 mcg (2000 IU)
02525	DOG Stress & Behavior	01936	Low-Dose Vitamin K2
PROBIOTICS		00373	No Flush Niacin
01622	Bifido GI Balance	01939	Optimized Folate (L-Methylfolate)
01825	FLORASSIST® Balance	01217	Pyridoxal 5'-Phosphate Caps
02421	FLORASSIST® Daily Bowel Regularity	01400	Super Absorbable Tocotrienols
02125	FLORASSIST® GI with Phage Technology	02334	Super K
01821	FLORASSIST® Heart Health	01863	Super Vitamin E
02250	FLORASSIST® Mood Improve	02422	Vegan Vitamin D3
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02234	Glycine	02040	Vitamins D and K with Sea-Iodine™
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01722	L-Tryptophan	02479	7-Keto® DHEA Metabolite
01668	Melatonin • 300 mcg, 100 veg capsules		100 mg, 60 veg capsules
01083	Melatonin • 500 mcg, 200 veg capsules	02207	AMPK Metabolic Activator
00329	Melatonin • 1 mg, 60 capsules	02504	Body Trim and Appetite Control
02503	Melatonin • 3 mg, 60 gummies	02478	DHEA Complete
00330	Melatonin • 3 mg, 60 veg capsules	02506C	Mediterranean Weight Management
00331	Melatonin • 10 mg, 60 veg capsules	01432	Optimized Saffron
00332	Melatonin • 3 mg, 60 veg lozenges	02511	Thermo Weight Control
02201	Melatonin IR/XR	02509	Waistline Control™
01787	Melatonin 6 Hour Timed Release	WOM	EN'S HEALTH
	300 mcg, 100 veg tablets	01942	Breast Health Formula
01788	Melatonin 6 Hour Timed Release	01626	Enhanced Sex for Women 50+
	750 mcg, 60 veg tablets	01894	Estrogen for Women
01721	Optimized Tryptophan Plus	02204	Menopause 731™
01445	Quiet Sleep Melatonin • 5 mg, 60 veg capsules	02319	Prenatal Advantage
02502	Rest & Renew	01649	Soy Isoflavones
02526	Serene Sleep	02513	Women's Bladder Support
VITAMINS			
01533	Ascorbyl Palmitate		
01000	Ponfotioming with Thioming		

00920

01945

00102

00084

02229

Biotin

Benfotiamine with Thiamine

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Magnesium is a sweet way to prioritize whole-body healthespecially the heart and bones.

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"Great product." Roniele

VERIFIED CUSTOMER REVIEW



60 vegetarian capsules

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4 bottles \$8.75 each







For full product description and to order Extend-Release Magnesium, call 1-800-544-4440 or visit www.LifeExtension.com

Extend-Release Magnesium

6 Hour Extended Release Formula

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Fish oil is a popular supplement people use to boost omega-3 levels in the heart, brain, eyes, and other tissues.

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CAUTION: If you are taking anti-coagulant or anti-platelet medications, or have a bleeding disorder, consult your healthcare provider before taking this product.



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IN THIS EDITION OF LIFE EXTENSION MAGAZINE®













7 RATIONALE FOR COMBINED INTERVENTIONS

A **2023** review stressed the **urgent** need for <u>combination</u> approaches to target **aging**.

22 THE HUMAN BRAIN AND BENFOTIAMINE

Benfotiamine has been shown to improve **cognition** scores in **clinical studies** compared to **placebo**.

32 TARGET MULITIPLE AGING PATHWAYS

Scientists have identified nutrients that correlate with **reduced mortality** and **greater healthspan** by favorably modulating multiple *mechanisms of aging*.

44 GLP-1 FOR DIABETES, OBESITY, AND CARDIAC FUNCTION

GLP-1 agonist drugs are game changers in controlling **blood sugar** and **body weight**. Clinical trials show a **20**% *reduction* in cardiovascular events and death.

52 VITAMIN D AND CANCER

Clinical trials indicate <u>who</u> is most likely to benefit by supplementing with more **vitamin D**.

65 WHAT IS PQQ?

By promoting the generation of *new* cellular mitochondria, **PQQ** helps promote full-body health.