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Volume Twenty Three / Number One • January 2017

REPORTS

26 KILL DANGEROUS BACTERIA IN YOUR INTESTINES

Probiotics have become enormously popular supplements. When bacteria-killing **phages** were added to **probiotics** in the animal model, there was a 40-100-fold increase in beneficial intestinal flora with huge reductions pathogenic *E.coli*. Combining **probiotics** with **phages** may represent a new standard of intestinal balance.



Apigenin is a polyphenol found in vegetables such as parsley and celery. Data shows that apigenin can effectively starve cancer cells, guard DNA against toxins, and block malignant cell propagation.

58 ACHIEVING OPTIMAL SELENIUM STATUS

Selenium has long demonstrated cancer prevention potential, but controversy exists over what are ideal forms of this mineral. A continuous stream of data reveals how different selenium compounds exert their own unique effects in impeding malignant transformation.

68 OUERCETIN BLOCKS DAMAGING EFFECTS OF PESTICIDES

Pesticides are absorbed through our air, food, and water. These chemicals can lead to neurodegeneration, cancer, and other disorders. Researchers have found that the flavonoid quercetin offers protection against some of the dangers inflicted by pesticides.

79 RESEARCH UPDATE: POO REDUCES ARTHRITIS INFLAMMATION

PQQ, or **pyrrologuinoline guinone**, can improve heart and brain health, and possibly slow the progression of aging. Researchers have found that **PQQ** has potential to decelerate the deterioration of joints in rheumatoid arthritis and osteoarthritis, exerting a protective effect in the joints.

86 METFORMIN REDUCES THE INCIDENCE OF OPEN-ANGLE GLAUCOMA

Glaucoma is the second leading cause of blindness in the world. Scientists have discovered persuasive data that the AMPKactivating drug metformin may help protect against glaucoma. **Life Extension**° encourages those at risk to speak to their doctor about these new findings.



36 NATURAL PREVENTION OF DEEP VEIN THROMBOSIS

Those who sit more than four hours a day have a 48% increased risk of potentially lethal blood clots, known as deep vein thrombosis (DVT). A human study found that those taking a dual plant extract experienced zero cases of deep vein thrombosis. This and other data underscore a remarkable opportunity to protect against a leading cause of disability and death in adults, which is an abnormal **clot** that forms inside a **blood vessel**.



DEPARTMENTS

7 AS WE SEE IT: **GREATEST THREAT TO LONGEVITY**

According to the Surgeon General, deep vein thrombosis may cause up to 180,000 deaths each year. Those who spend time sitting are at the highest risk. Researchers have developed a nondrug approach for the prevention of deep vein thrombosis, utilizing two natural compounds that drastically reduce platelet aggregation-and fibrin-induced clots.

19 IN THE NEWS

Vitamin D shortens hospital stays; omega-3 reverses heart damage; drug doubles melanoma survival time; and taurine and magnesium inhibit cardiovascular disease.









Volume Twenty Three / Number One

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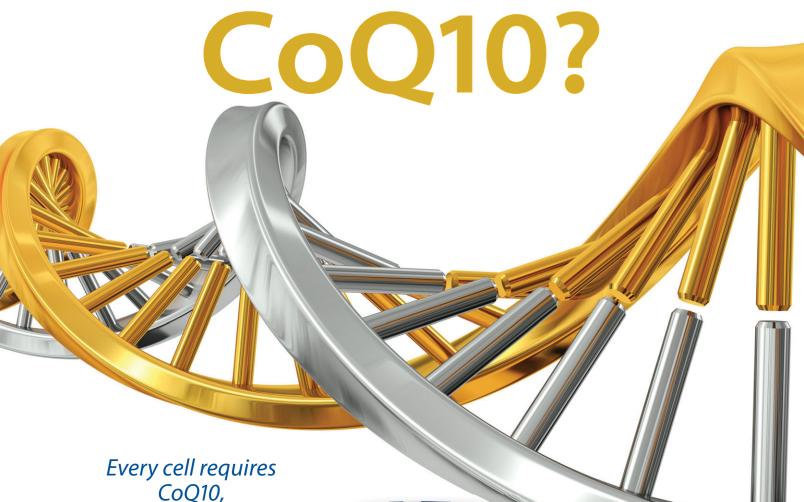
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References

- 1. Regul Toxicol Pharmacol. 2007;47(1):19-28.
- 2. Exp Neurol. 2004;188(2):491-4.
- 3. Pharmacologyonline. 2009;1:817-25.

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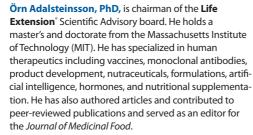
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Greatest Threat to Longevity

BY WILLIAM FALOON

Twenty years ago I was leaving a medical conference when one of our ardent supporters rushed up and handed me a huge textbook.¹ She begged I take it home to read.

She was adamant that *Life Extension*® make a greater effort to enlighten its readers about the underlying <u>cause</u> of most disability and death in persons over age 50.1

The threat described in the textbook occurs when an **abnormal blood clot** forms inside an artery or vein. The medical term is **thrombosis**.

Two disorders involving **arterial** thrombosis are:

- Heart Attack
- Ischemic Stroke

Two disorders involving **venous** thrombosis are:

- Deep Vein Thrombosis (DVT)
 - Pulmonary Embolism

Those stricken with **cancer** are particularly susceptible to **venous thrombosis**.² Chemotherapy patients are up to **6-times** more vulnerable.³

One reason we've recommended **low-dose aspirin** since **1983** is its ability to <u>inhibit</u> platelet aggregation, a major factor involved in **arterial thrombosis**, leading to a **heart attack** or **ischemic stroke**.⁴

Recent studies show that **arterial thrombosis** occurs <u>more</u> frequently than previously thought.^{5,6} Minor thrombotic events seldom display outward symptoms and, over time, predispose us to a host of degenerative illnesses including mind-robbing **ministrokes**.^{6,7}

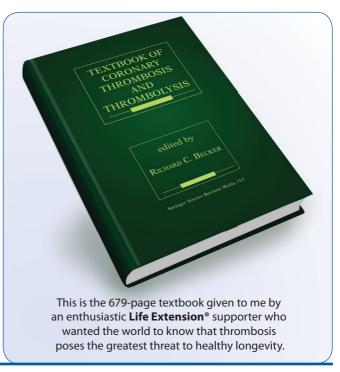
Many of the **nutrients** you take have <u>diverse</u> **antiplatelet** effects. This is important in protecting against **arterial** thrombosis, but far less so for **venous** thrombosis.

The Surgeon General published a report showing that deep vein thrombosis (and subsequent pulmonary embolism) may cause 100,000-180,000 deaths each year in the US.8 To put this in perspective, pancreatic cancer is estimated to kill more than 41,000 Americans in 2016.9 Pancreatic cancer has a decidedly deadly reputation, yet the public is largely unaware that deep vein thrombosis (and subsequent pulmonary embolism) poses a greater overall health risk.

The Surgeon General was highly critical of mainstream medicine for <u>not</u> recognizing patients at risk for **deep vein thrombosis** and taking appropriate preventive measures.

Longtime readers of this magazine should be comforted with the knowledge that they are already taking steps to <u>reduce</u> their **arterial thrombotic risk**.

This issue describes startling new data about **deep vein thrombosis**, and what can be done to help prevent it.



As We See It

People often take for granted that blood effortlessly flows through their arteries and veins like water moves through a hose.

The reality is that **blood flow** is highly dependent upon a complex interplay of different mechanisms, including coagulation factors that regulate the tendency of blood to form a clot.

For example, **platelets** play an important role in "plugging" holes in our circulatory system, helping to reduce bleeding in conjunction with other clotting factors.

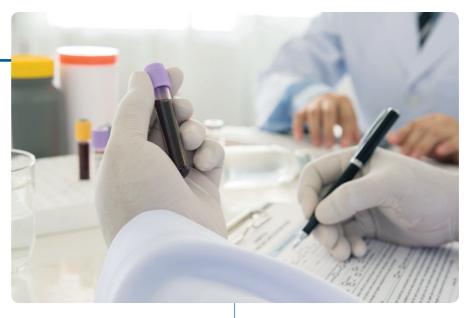
Conversely, when platelets abnormally aggregate (clot) inside a blood vessel in response to arterial plaque and/or endothelial damage, the result is stoppage of blood flow to the affected part of the anatomy. An abnormal blood clot in a cerebral artery can lead to **ischemic stroke**, whereas a thrombus (clot) that forms in a coronary artery can result in a heart attack.

As humans age, mini-thrombotic events can occur to small arteries in the brain. This includes transient ischemic attacks (TIAs) in the brain that, over time, can cause damage to our cognitive abilities.

Preventing the development of these minor and major thrombotic episodes is critical for healthy longevity. The good news is that we know a lot about what causes pathologic **blood clotting** inside arteries and veins, and how to prevent it.

Role of Inflammation in **Both Arterial and Venous Thrombosis**

Inflammation sets in motion a sequence of events that can lead to arterial and venous thrombosis. Normal aging results in increasing levels of vascular inflammation, often without outward symptoms.



Readily obtainable blood markers that can reveal systemic inflammation are **homocysteine**, ¹⁰⁻¹² C-reactive protein, 13,14 and fibrinogen. 15-17 Heightened levels of these inflammatory biomarkers are correlated with arterial thrombosis and subsequent risk of cardiovascular disease.18,19

Fish oil,²⁰ vitamin D,^{21,22} cur**cumin**,^{23,24} and other plant extracts inhibit many underlying inflammatory factors that increase **C-reactive protein**. The biologically active form of folic acid (5-MTHF)^{25,26} along with vitamins B12²⁷ and B6²⁷ can slash elevated homocysteine through two distinct detoxification pathways.28

In contrast to the association between risk of arterial thrombosis and C-reactive protein elevation, **C-reactive protein** is not very useful in predicting future **venous** thrombosis or pulmonary embo**lism**. Recent evidence does point to the role of proinflammatory cytokines like IL-8 and tumor necrosis factor-alpha in venous thrombosis risk.29

Nattokinase is an enzyme extracted from a Japanese food (natto) prepared from fermented soybeans.30 Venous inflammation tends to raise fibrinogen levels, and fibrinogen is an important factor involved in **inflammation** as well as venous thrombosis formation.31

Nattokinase has been shown to decrease levels of **fibrinogen** along with clotting factors VII and VIII. which are involved in the formation of venous thrombosis.32

Epidemic of Deep Vein Thrombosis

Deep venous thrombosis and pulmonary embolism are major causes of disability and death.³⁹

Each year, as many as 900,000 Americans may be affected by venous thromboembolism. Of those diagnosed, up to 30% will die within one month, and the first symptom will be sudden death in about 25% of those who have a pulmonary embolism.⁴⁰

Venous thrombosis is the formation of a blood clot inside a vein that can obstruct flow in the localized affected part of the venous circulatory system. When a venous blood clot dislodges from its primary location and travels to block circulation in another body part, this is referred to as a venous throm**boembolism.** When a deep vein thrombosis dislodges and travels to the lungs, this worrisome and potentially life-threatening condition is called a **pulmonary** embolism.

Conventional Arterial Thrombotic Risk Factors

LDL cholesterol is a common factor involved in development of atherosclerotic plaque in arteries.33

Elevated LDL contributes to deposits of plaque that cause arterial pathways to gradually narrow until normal blood flow is disrupted. When this happens, there is a greater propensity for arterial clot formation (arterial thrombosis).33

Hypertension increases the velocity at which blood is thrust through the arterial system. As blood pressure elevates, platelets become more likely to aggregate and create a thrombotic event.34

Conventional cardiovascular and arterial thrombosis risk factors are diabetes, smoking, abdominal obesity, and hyperlipidemia (elevated LDL and triglycerides). 35-38 Some of these same factors are also associated with increased risk of deadly venous thrombotic events.

The encouraging news is that proven methods exist to control underlying causes of thrombosis and the vascular diseases that can develop acutely or chronically.

A variety of factors are implicated in the formation of venous thrombosis. Two major, related risks for the development of **deep** vein thrombosis are:

- **Hemostasis** (reduction/ stagnation of blood flow)
- Hypercoagulability (propensity of blood to clot inside veins due to lifestyle, cancer or genetics)

The good news is that steps can be taken to reduce the risk of deep vein thrombosis, as well as thrombotic risks throughout the circulatory system. This means that strategies to protect against deep vein thrombosis may also confer protection against stroke and heart attack.

What Causes Blood Vessel Clots in Arteries?

To sustain life, blood must remain in a fluid state so that it can freely circulate, while simultaneously being able to properly **clot** at the site of a vascular injury.

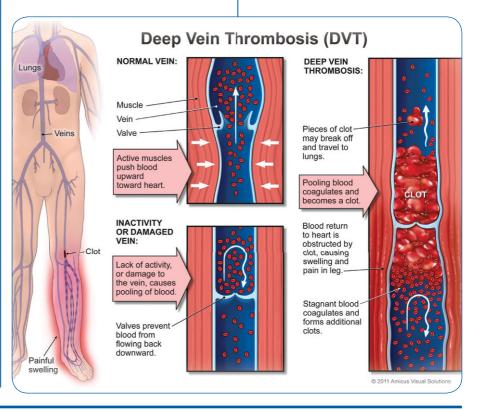
Any event that <u>activates</u> platelets can cause them to aggregate to form an occlusive thrombus.

As it relates to **aspirin**, **fish oil** and certain plant **polyphenols**. vou'll often read about their "antiplatelet" properties. What this describes is their ability to interfere with platelet activity, adhesion, and aggregation, thereby reducing arterial thrombotic risk.

Antiplatelet therapies, however, can be sabotaged by dysfunction of our endothelium (inner arterial lining). A healthy endothelium produces substances that stabilize platelets and impede their unwanted adhesion.

When the endothelial lining is lost, platelets are exposed to parts of the arterial wall that cause them to aggregate. Protecting against endothelial dysfunction is thus essential to maintain vascular health as we age. This is where **pomegranate** and other plant polyphenols play a critical

Antiplatelet strategies employed today, utilizing certain medications and nutrients, can greatly mitigate these arterial thrombotic factors.



How Arterial and Venous Clots Differ

There are distinctions between the processes that cause <u>arterial</u> and <u>venous</u> thrombosis.

Arterial thrombosis largely involves platelet aggregation forming around clogged/jagged points in the arterial system, or in response to irregular heart beat (atrial fibrillation) or an artificial heart valve.

Deep vein thrombosis typically occurs due to **hemostasis** (reduction in venous blood flow) and **hypercoagulability** (tendency of the blood in veins to clot due to genetic, cancer or lifestyle factors).

One major cause of reduction in venous blood flow is **chronic venous insufficiency**. This frequently occurs from obesity, lack of physical activity/sitting with the legs in a dependent position, and previous deep vein thrombosis, which injures or destroys one or more of the **valves** that are located in the deep veins of the leg.

In order to efficiently return blood to the heart when a person is sitting or standing, veins contain tiny **valves** that open and close. Properly functioning valves prevent blood from flowing backward while muscles surrounding the veins compress them, helping pump venous blood back to the heart.⁸

Veins contain valves while arteries do not. When veins are damaged by prior venous clot, or physical inactivity leads to pooling of blood in the deep veins of the legs, venous blood flow decreases, setting the stage for venous thrombosis.

In contrast with the venous system, platelets in the arterial system are adversely activated as they bump into buildups of **plaque** along the arterial walls and interact with a *dysfunctional* endothelium. In this scenario, platelets begin to clump together, causing a cascade that can lead to blood flow being cut off to vital tissue (such as a portion of the heart muscle).

In the venous system, normal blood flow can slow, and if left to **stagnate** too long, the blood within these veins begins to coagulate (clot). The problem of deep vein thrombosis, however, extends beyond mere stagnating pools of blood in the lower legs.

Symptoms of Deep Vein Thrombosis

A blood clot in one of the deep veins can include the following symptoms:⁸³

- Mild to severe pain in the affected arm or leg
- Swelling of an arm or leg
- Redness or color change
- · Warmth of the skin

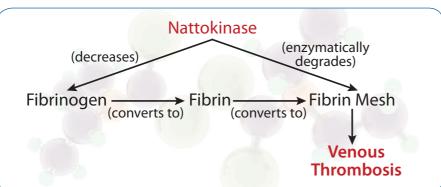
A venous blood clot that has traveled to the lung (called a pulmonary embolism) has symptoms that include:⁸³

- Chest pain
- Shortness of breath



- Green tea^{41,42}
- Fish oil⁴³⁻⁴⁶
- Olive polyphenols⁴⁷⁻⁵¹
- Ouercetin⁵²
- Resveratrol⁵³⁻⁵⁶
- Grape seed extract⁵⁷⁻⁶¹
- French maritime pine bark extract⁶²
- Lycopene^{63,64}
- Pomegranate⁶⁵
- Garlic⁶⁶⁻⁷²
- Flax seed oil^{73,74}
- Ginger^{66,75-80}
- Curcumin^{81,82}





Formation of Deep Venous Thrombosis

Fibrinogen converts to **fibrin** that creates a **fibrin mesh** inside a vein. Platelets and red blood cells adhere to fibrin mesh to cause deep vein

Nattokinase reduces fibrinogen and has fibrinolytic effects, which means it can enzymatically break down the fibrin mesh component of blood clots.

Beyond "Antiplatelet" Strategies

Inflammation and some other factors that contribute to arterial thrombosis (such as excess homocysteine) also are linked with an increased risk of deep vein thrombosis.84,85 Several of these dynamics, however, play a *larger* role in the venous system than in arteries.

What happens with deep vein thrombosis is that **fibrinogen**, also a proinflammatory regulator, excessively converts to a fibrin **mesh** that traps red blood cells. In deep vein thrombosis, formation of **fibrin** is also linked with excess inflammation in veins.86 This helps explain the limited efficacy of antiplatelet drugs (aspirin and Plavix®) in venous thrombosis, i.e., they don't stop the initiating phase of proinflammatory fibrinogen converting to red blood cell-trapping fibrin.

There are multiple underlying coagulation factors that can initiate a deep venous thrombosis. Some require preventive treatment using anticoagulant drugs (warfarin, Pradaxa[®], Eliquis[®], Xarelto[®]). There is a common venous thrombotic mechanism, however, that can be impeded with a low-cost nutrient.

Nattokinase has been shown to decrease fibrinogen levels and help dissolve **fibrin clots** that obstruct blood flow.32,87

The flow chart on this page shows a simplified version of the complex process of coagulation involved in thrombosis formation. The **fibrinogen**/ fibrin-dissolving effects of nattokinase can help stop this coagulation cascade at several checkpoints.

As We See It

Impressive Human Data!

A recent study highlighted the risk of deep vein thrombosis in response to conditions that predispose to stagnation of venous blood in the lower extremities (legs) specifically, air travel.96

Many people are not aware that air travel, and associated pooling of venous blood in the legs due to inactivity during flight, is a major risk factor for deep vein thrombosis and potentially life-threatening pulmonary embolism. 97,98

Using ultrasound imaging, the presence of venous thrombosis was detected in a startling 5%-7% of passengers of flights lasting 7-8 hours.96 These passengers were asymptomatic, meaning they did not know they had developed a venous blood clot!

When a nondrug approach for prevention of deep vein throm**bosis** was studied in this group of passengers, no lower leg venous clots were detected and lower leg edema (swelling) was drastically reduced.

Cancer-Related Thromboembolism

Cancer is associated with a 4.1-fold increased risk of venous thromboembolism.88

Poor mobility, venous obstruction, and ongoing chemotherapy further increase risk of recurrence.^{89,90} Venous thromboembolism is associated with advanced and more aggressive cancers. 91,92

Cancer patients with **venous thromboembolism** have worse survival than cancer patients free of this complication. 91 For example, after a diagnosis of venous thromboembolism, the mortality rate at 6 months for cancer patients on anticoagulant therapy is 40%.92

Cancer patients today are dying prematurely from venous thromboembolism. This is why *Life Extension*® long ago recommended aspirin and low-molecular weight heparin as adjuvant cancer therapies. Not only do overly-active platelets contribute to **thrombosis**, but they facilitate metastasis.93-95

What might surprise you is that **venous thromboembolisms** can be the first clinical manifestation of cancer somewhere in one's body. About 10% of patients with unprovoked **venous thromboembolism** are diagnosed with cancer. Of these, more than 75% are diagnosed within the first year after their thrombotic episode.2

As We See It

Whenever you are faced with long-haul air travel, you should stand up every few hours and take a walk through the plane cabin. Consider obtaining high quality compression stockings to wear whenever you fly to reduce stagnation of blood in lower leg veins.99

As you will read in this month's issue, ingestion of two nutrients taken two hours before the plane departs and again six hours into the flight drastically reduced detection of venous blood clots and lower leg edema at the end of the flight. These nutrients have a dual effect of inhibiting platelet aggregation and helping to thwart fibrininduced clots.

The fact that short-term dosing of these two nutrients demonstrated such a profound effect in protecting against deep vein thrombosis implies significant systemic benefits for those who supplement daily.

Surgeon General's Call to Action

In a report published 8 years ago, the Surgeon General stated:

"DVT [deep vein thrombosis] and PE [pulmonary embolism] are major public health problems in the United States. Much is known about how to reduce their burden, yet this knowledge is not being applied systematically today. Without a concerted effort to stem this public health crisis, the incidence and burden of these diseases will only grow larger as the population ages."8

Sadly, this medical neglect continues as hurried physicians are not doing enough to prevent thrombotic events that not only cause DVT/pulmonary embolism, but many strokes and heart attacks.

An intriguing article in this month's issue describes the robust benefits of these nutrients for reducing deep vein thrombosis risk.

The first article unveils a novel way of enhancing the efficacy of your **probiotic** by selectively killing off harmful intestinal bacteria.

Obtain Nutrient Formulas at Year's Lowest Prices

This is the time of year when we **discount** prices on every one of our advanced nutritional formulas.

Longtime supporters take advantage of the once-a-year **Super Sale** to stock up on their favorite nutrient formulas.

Those who have engaged in healthy lifestyle choices should find comfort knowing that nutrients they may have been using for decades confer considerable protection against **thrombosis**, which in persons over age 50.

Our Commitment

No organization is working harder to accelerate human age reversal research than Life Extension.

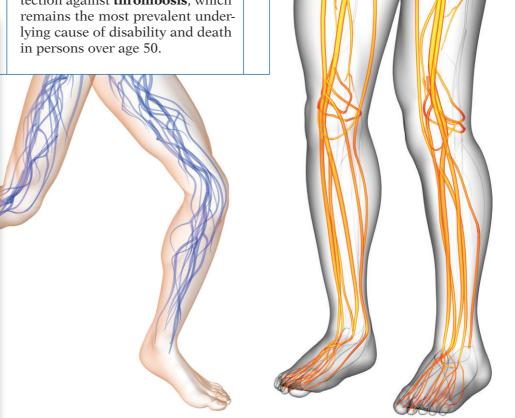
Your support enables scientists to engage in biomedical research that would have been inconceivable just a few years ago.

To order nutrients you need today at Super Sale prices, call 1-800-544-4440.

For longer life,

William Faloon

(References may be found on page 14.)



Identify Common THROMBOTIC RISK Blood Factors

Special Introductory Price

Numerous factors in **blood** can contribute to *abnormal* blood clots.

Thrombosis is the term used to describe **clots** that form inside blood vessels.

Rather than wait for an acute stroke, coronary blockage, or deep vein thrombus, one can evaluate common thrombotic risk factors that are measured in the blood.

A new low-cost panel has been designed to detect abnormalities in one's blood that can be corrected before a disabling or lethal blood vessel clot develops.

One of these clotting factors is **fibrinogen** which markedly increases risk of thrombosis, especially stroke.

The new **Thrombotic Risk Panel** provides the following tests at a special introductory price of only \$139 (if ordered before January 31, 2017):

Inflammatory/Clotting Markers Insulin Resistance Markers

- Fibrinogen
- C-reactive protein
- Homocysteine
- PT/PTT/INR

Blood Cells

- Platelet count
- Red Blood Cell count
- Anemia markers
- White Blood Cell count (includes differentiation)

- Glucose
- Insulin
- Hemoglobin A1C

Lipid Markers

- Cholesterol (total)
- I DI
- HDI
- **VLDL**
- **Triglycerides**

Liver/Kidney Function and more!

Don't Delay—Order today, visit www.LifeExtension.com or call one of our knowledgeable Wellness Specialists at 1-866-864-3027.

This offer is good through January 31, 2017.



Fasting Not Required

Conventional protocol says that one should fast for 8-12 hours prior to a blood draw. The fast means no eating or drinking anything except water and taking one's prescription medica-

Life Extension® will soon make an announcement based on new evidence indicating that more valuable data may be obtained if one is NOT in a fasting state when their blood is drawn.

If you are used to fasting prior to a blood test you may continue this practice, but you now have the **option** of eating/drinking what you **normally** do in a typical day.

Just write down the time of your last meal and what it contained. This provides a snapshot of what your blood **normally** looks like on a typical day as opposed to what may be artificially different levels that occur during a fasting state.

Thrombotic Risk Panel • Item # LC100055 Retail: \$259 • Introductory price: \$139

Additional Blood Tests for Thrombosis

Unlike arterial thrombosis risk due to elevated LDL-C cholesterol and hs-CRP, as well as elevated arterial and venous thrombosis risk due to fibrinogen and homocysteine, some individuals suffer from other coagulation disorders that require additional blood tests to detect.

Additional tests available from your doctor for coagulation factors to evaluate for venous thrombosis risk include:

- Factor V Leiden (mutation)
- Prothrombin gene G20210A (mutation)
- Antiphospholipid antibodies (e.g. lupus anticoagulant, cardiolipin antibody, beta-2 glycoprotein antibody)
- Protein Cantigen and activity
- Protein S antigen and activity



When Anticoagulant Drugs Are Needed

People with artificial heart valves or atrial fibrillation are at high risk for developing a thrombus that breaks loose and travels up a carotid artery, where it can cause an acute ischemia stroke. 100,101

There are also inherited conditions in which blood clotting proteins improperly react, either causing blood to overcoagulate or preventing expression of normal clot dissolving factors. Some of these coagulation disorders that result in too much clotting include:

- Factor V Leiden
- · Antithrombin III (ATIII) deficiency
- Protein C or protein S deficiency
- Prothrombin (PT) gene mutation
- · Antiphospholipid antibody syndrome

Those in a hypercoagulable state are usually prescribed one of the four following anticoagulant drugs:

- Pradaxa®
- Eliquis®
- Xarelto®
- Coumadin® (warfarin)

The major side effect risk of these drugs is unwanted bleeding. These drugs also don't always prevent a thrombotic event. Those who choose the oldest of these drugs (warfarin) are subjected to severe vitamin K deficiency that rapidly calcifies tissues. This can lead to future degenerative illnesses (such as accelerated atherosclerosis and aortic valve stenosis).

Despite these side effect risks, those at high thrombotic risk should work closely with their physician to use the anticoagulant drug that best meets their individual needs. To review our detailed report on the pros and cons of each of these drugs, log on to:

LifeExtension.com/thrombosis

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Vitamin D3	1,000 IU	2,000 IU
Vitamin E	50 IU (synthetic)	100 IU (natural)
Thiamine (vitamin B1)	1.5 mg	75 mg
Riboflavin (vitamin B2)	1.7 mg	50 mg (with riboflavin- 5-phosphate)
Niacin	20 mg	50 mg
Vitamin B6	3 mg	75 mg (with pyridoxal- 5-phosphate)
Folate	400 mcg (synthetic)	400 mcg (5-MTHF)
Vitamin B12	25 mcg	300 mcg
Biotin	30 mcg	300 mcg
Pantothenic acid	10 mg	100 mg
Calcium	220 mg	10 mg
lodine	150 mcg	150 mcg
Magnesium	50 mg	100 mg
Zinc	11 mg	30 mg
Selenium	19 mcg (one form)	200 mcg (three forms)
Manganese	2.3 mg	2 mg
Chromium	50 mcg	200 mcg
Molybdenum	45 mcg	100 mcg
Potassium	80 mg	25 mg
Alpha lipoic acid	none	25 mg
Boron	none	3 mg
Choline (as bitartrate)	none	20 mg
Inositol	none	50 mg
Lutein	250 mcg	5,000 mcg (from marigold extract)
Zeaxanthin	none	155 mcg (from marigold extract)
Lycopene	300 mcg	1,000 mcg
Natural mixed tocopherols (providing gamma, delta, alpha, and beta tocopherols)*	none	20 mg
Apigenin	none	5 mg



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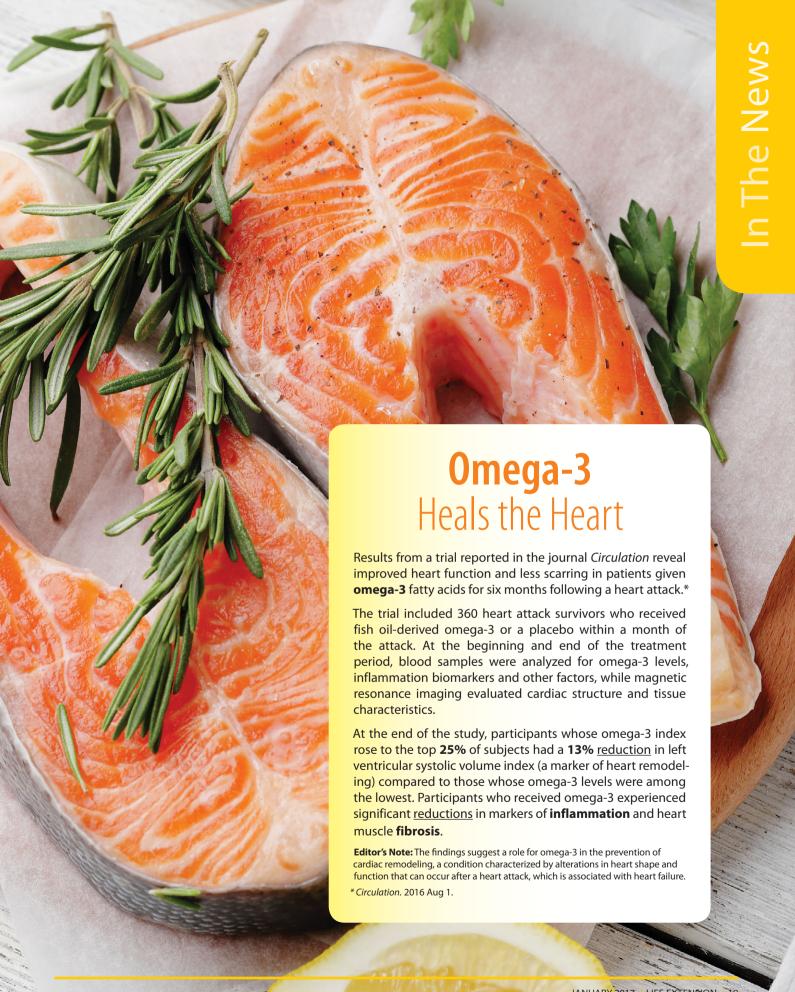
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Vitamin D

Associated with Shorter Hospital Stays in Ventilated Patients

A pilot trial reported in the *Journal of Clinical and Translational Endocrinology* found that critically ill patients receiving mechanical ventilation who were given a high-dose **vitamin D** supplement had a <u>shorter</u> hospital stay in comparison with those who received a placebo.*

Researchers randomized 31 ventilated intensive care unit patients to receive **50,000 IU** or **100,000 IU** of vitamin D3 daily for five days. Thirteen of the subjects had deficient plasma 25-hydroxyvitamin D levels of less than **20 ng/mL** at the beginning of the study. Blood samples were analyzed one and two weeks after the beginning of the treatment period to measure plasma vitamin D and other factors.

Average length of hospital stay was 25 days among those in the **50,000 IU** group and 18 days in the **100,000 IU** group, in comparison with 36 days in the placebo group. The high-dose **vitamin D** group thus spent <u>half</u> as much time hospitalized.

High doses were needed because these ICU patients were in acute need to rapidly build up their vitamin D levels.

Editor's Note: "High-dose vitamin D may have multifactorial effects that could contribute directly or indirectly to hospital length of stay, including salutary effects via improved 25-hydroxyvitamin D levels on respiratory or other skeletal muscle function, by modulation of the pro-inflammatory milieu, and by regulation of immune functions, among other contributors," authors Jenny E. Han and colleagues conclude.

* J Clin Transl Endocrinol. 2016 Jun;4:59-65.







Drug More Than Doubles Chances of Five-Year Survival Rate

According to a study presented at the annual meeting of the *American Association for Cancer Research* in New Orleans, patients with advanced **melanoma** who were treated with the immune-oncology drug **Opdivo®** had far higher survival rates. After 5 years, **34%** of patients receiving Opdivo were alive compared to **16%** of melanoma patients receiving conventional treatment methods.*

Significantly, oncologists say the results indicate that patients who survive for about four years are highly unlikely to relapse because their immune systems have eradicated or controlled their tumors.

Opdivo® is an immune checkpoint inhibitor. This type of drug blocks certain proteins made by some types of immune system cells, such as T cells, and some cancer cells. These proteins suppress immune responses and can keep T cells from killing cancer cells. When these proteins are blocked, the "brakes" on the immune system are released and T cells are able to better kill cancer cells.

Editor's Note: The first checkpoint inhibitor to reach the market was Yervoy®. It targets a brake known as CTLA-4 and in a previous analysis was shown to result in long-term survival in about 17% of melanoma patients. A report on check-point inhibitor drugs was published in the June 2016 issue of *Life Extension Magazine®*. Oncology experts working with Life Extension® report favorable outcomes when checkpoint inhibitors are used against other malignancies.

* American Association for Cancer Research. 2016 April 16-20.



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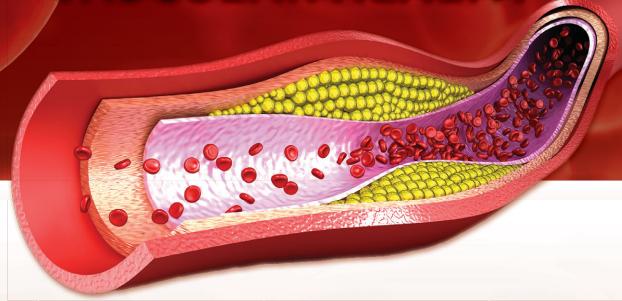
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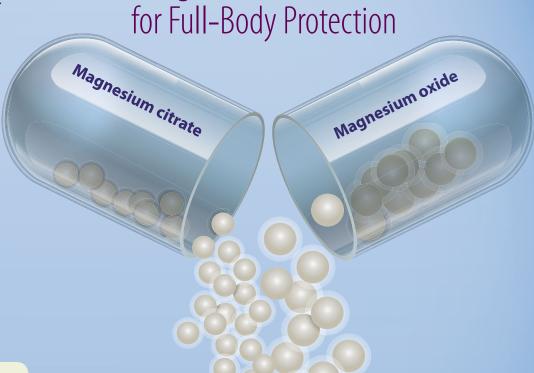
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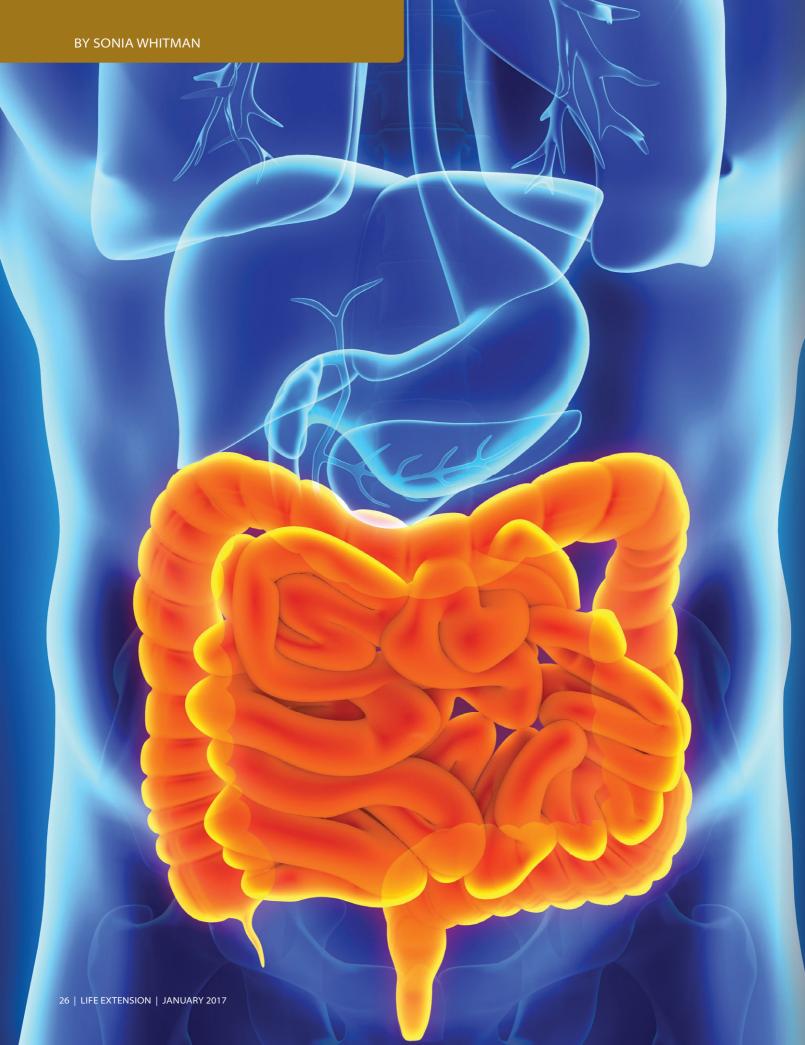
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Major Advance in INTESTINAL HEALTH

Probiotics are emerging as enormously popular health supplements.

When one ingests a **probiotic**, they add bacteria to their digestive tract that produce benefits ranging from alleviating **intestinal distress** to strengthening **immunity**.

Another virtue of **probiotics** is that they can slowly crowd out harmful **bacteria** strains in the intestines.

In a significant advance, a natural method of selectively killing <u>undesirable</u> **bacteria** has been developed.

When combined with a probiotic in the animal model, there was an exponential <u>increase</u> in beneficial bacteria with a parallel <u>decrease</u> in unfriendly flora like *E. coli*.¹⁻³

This article describes a novel way of *optimizing* **probiotic** efficacy.

Most people in the United States have likely never heard of **phage therapy**. It was discovered in pre-World War I Eastern Europe, and for much of the pre-World War II era, it was thought to be a promising approach to controlling bacterial illnesses.4-6

In the 1940s, industrial giants like Eli Lily and L'Oréal developed bacteriophage "cocktails" aimed at treating a variety of bacterial infections. But the advent of antibiotics quickly cost phages the spotlight, though their effectiveness has never been in question.⁷

Phage therapy uses *bacteriophages*, which are submicroscopic packages of DNA or RNA enclosed in a protein envelope that **selectively** target harmful bacteria.

The name bacteriophage literally means "bacteria eating." A bacteriophage attaches itself to a bacterial cell wall and then destroys the host bacteria.

Bacteriophages are ubiquitous in nature, meaning they can be found almost everywhere—from soil, hot springs and the ocean depths to the animal and human body.8

The name **bacteriophage** literally means "bacteria eating."

Phages are a common and important component of gut flora and are found in various other parts of the human body such as the mouth and skin.^{8,9} Phages are currently used in the food industry to control diseasecausing organisms. 10-12

Numerous phages are classed as GRAS—or "generally recognized as safe"—and are commonly used for a variety of different applications, from controlling Listeria in cheese and E. coli in meat and on food-contact surfaces, to Salmonella in food.

With the targeted use of **bacteriophages**, it is possible to seek out and effectively reduce specific populations of unhealthy organisms that have taken over the intestinal microbiome.

The use of bacteriophages allows an exceptionally specific approach to eliminating detrimental bacteria. This is in direct contrast to **antibiotics**, which employ a mass-killing technique that eliminates healthy and detrimental bacteria, leaving us vulnerable to attack by other organisms.13

By removing common pathogenic bacteria in one's gut, **bacteriophages** enable beneficial *probiotic* bacteria to thrive, allowing them to more effectively rebalance the microbiome.

When **phages** are combined with **probiotics** in animal models there are huge reductions in the targeted harmful bacteria with a simultaneous increase in beneficial bacteria.

The Importance of a Healthy Gut

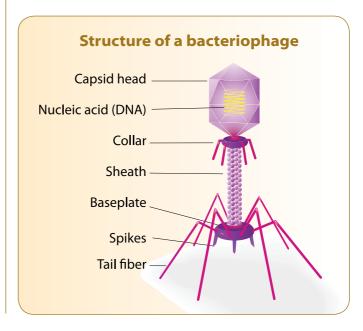
When scientists speak of the intestinal *microbiome*, they refer to the community of trillions of microorganisms that populate the human gut.14 We're just now beginning to understand that this collection of microorganisms can determine our state of health and illness.

It is impossible to avoid all threats to our intestinal microbiome. The increase in our consumption of meats, fats, processed carbohydrates, preservatives, and other additives can alter our bacterial populations. 15-17 And, of course, the overuse of antibiotics kills both healthy and bad bacteria indiscriminately.^{7,18,19} As a result, our delicate **microbiome** is becoming disrupted and imbalanced, which results in a condition called *dysbiosis*.

Dysbiosis is associated with almost every known disease process, from obesity and cancer to cardiovascular disease, diabetes, allergic reactions, neurodegenerative disorders such as Alzheimer's, and even mental health issues such as depression.²⁰⁻²²

Imbalances in the intestinal microbiome steadily worsen with age,23-26 and can be blamed for such common complaints as feeling generally unwell, reduced sleep quality, and feeling "foggy." 27-31

The impact of an unbalanced microbiome can also lead to another disorder related to dysbiosis called *small intestinal bacterial overgrowth.* People with this condition often have vague but persistent abdominal symptoms, such as pain, bloating, indigestion, gas, or cramping. 32,33 And in older adults, small intestinal bacterial overgrowth can contribute to malnutrition and eventual frailty, making the condition a major threat to the elderly.^{32,34}



Giving Probiotics a Boost

Many people live in a state of **dysbiosis** or imbalance that not only threatens their long-term health, but may also contribute to sleep disorders, to a sense of malaise or "fogginess," and to a range of stomach distress issues that cannot be explained by a specific disease. Fortunately, many studies have shown that positively changing the **microbiome**—shifting it toward a healthy profile and away from *dysbiosis*—can also change symptoms and disease risk.14,35,36

One important way of improving the intestinal microbiome is through the use of **probiotic bacteria**. Probiotics are a great *additive therapy* that increase the abundance of organisms that can rebalance an ailing microbiome. In some cases, however, probiotics by themselves have difficulty competing with the more aggressive microbes that contribute to dysbiosis in the first place.

A more comprehensive approach is to use **probi**otics in combination with another *therapy*, one that selectively targets and reduces the troublesome bacteria that are taking over the microbiome. This approach of targeting harmful bacteria while replenishing beneficial bacteria can make way for **probiotics** to better help restore the microbiome to a healthy, balanced state. That is how **phage therapy** works.

Phage therapy reduces potentially troublesome organisms that are overabundant in the microbiome, allowing beneficial organisms to flourish. This helps restore the **microbiome** to a healthy, balanced state.³⁷

Combating *E. coli* Bacteria

Escherichia coli (*E. coli*) is a bacterium that normally lives in our intestines. Most types of E. coli are harmless, but some can cause disease.

A troublesome strain of *E. coli* called **H10407** causes abdominal cramps, diarrhea, and gas. 38,39 This pathologic E. coli strain also suppresses growth of beneficial bacteria and produces a state of *dysbiosis*. 40

In an effort to help restore a balanced and healthy **microbiome**, researchers have developed a cocktail of **bacteriophages** that target dangerous strains of *E. coli*.

Unlike typical pre- or probiotics, this **phage cock**tail is effective within hours, not days, and in very small doses.⁴⁰ It functions not only in the colon (large intestine), where **dysbiosis** is a problem, but also in the small intestine, where undesirable bacterial over**growth** occurs. It does not cause flatulence, a constant problem with fiber-containing prebiotics.⁴⁰

Laboratory studies show that this form of **phage therapy** removes harmful bacteria, making way for beneficial **probiotic bacteria** to establish and form a healthy microbiome. This has been demonstrated—with impressive results.



What You Need to Know

The Revival of Phage Therapy

- An imbalance in the intestinal microbiome (dysbiosis) is now associated with many serious age-related disorders. Studies show that it is prevalent in older people, particularly those with sleep disorders, vague abdominal complaints, and malaise.
- Changing the intestinal microbiome towards a state of improved health and function is a desirable solution, but until now options have been limited.
- Bacteriophages selectively target and destroy specific harmful bacteria.
- In combination with a mixture of probiotic bacteria, a cocktail of bacteriophages has been shown to reduce the abundance of undesirable bacteria, while freeing beneficial organisms to thrive and increase in numbers.
- Using this probiotic/bacteriophage mixture shows high promise in relieving the functional changes and tissue damage wrought by dysbiosis, and should be added to the regimen of anyone interested in rebalancing their microbiomes to feel better.

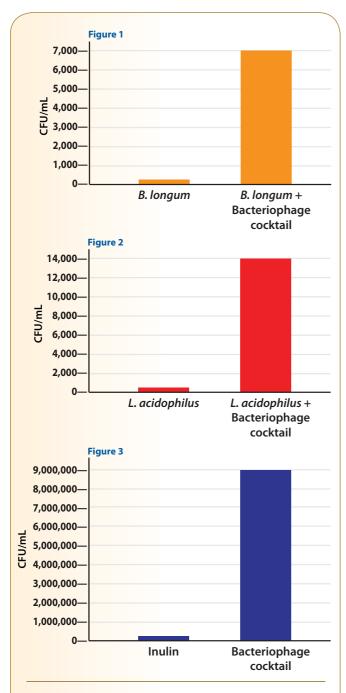


Figure 1: Beneficial Bifidobacterium longum grew nearly 7000 times better despite competition from E. coli, when the culture dish was treated with bacteriophage cocktail.40

Figure 2: Beneficial Lactobacillus acidophilus grew nearly 20 times better, despite competition from E. coli, when the culture dish was treated with bacteriophage cocktail.40

Figure 3: Beneficial Lactobacillus paracasei grew poorly in culture against competing bacteria when treated with prebiotic inulin (left). After treatment with bacteriophage cocktail, the desirable organism grew nine million times more vigorously than with inulin alone.40

Phage Cocktail Promotes Survival of Probiotic Organisms

In one experiment, culture dishes were prepared with a **beneficial bacteria** (*Bifidobacterium longum*), along with a competitive *E. coli* bacteria.⁴⁰

A second set of dishes was prepared to which the **bacteriophage** mixture was added. After just 5 hours, there was a visible difference.

In the dishes without **bacteriophage**, there was little growth of the desirable *B. longum* organisms, indicating their inability to compete with *E. coli*.

In the dishes with the **bacteriophages**, colonies of B. longum skyrocketed to more than 7,000 times their numbers compared to dishes without the bacteriophage. This study demonstrates that the competitive *E. coli* had been greatly removed from the picture by the bacteriophage (Figure 1).

In a related experiment, a probiotic organism (Lactobacillus acidophilus) was grown in culture, again in competition with E. coli.⁴⁰ In a very similar fashion, without the **phage cocktail**, the *E. coli* suppressed growth of the beneficial L. acidophilus. However, in the presence of the phage mixture, L. acidophilus thrived, reaching colony counts that were 20-fold higher compared to the culture not receiving the **phages**. (**Figure** 2). In similar experiments, growth of the probiotic organism *L. rhamnosus* was enhanced **15-fold** and the beneficial B. bifidum was enhanced nearly 10-fold when grown in culture with the **phage** cocktail.

These experiments demonstrate the value of the **phage** mixture in promoting the growth and survival of probiotic organisms.

In another Petri dish experiment, the common prebiotic inulin was used in an attempt to stimulate the growth of the beneficial probiotic *Lactobacillus* paracasei. When the prebiotic was used alone, it failed to ensure the survival of the desired organisms. But when the **phage** mixture was added, it produced an astonishing *nine million-fold* increase in growth of the probiotic Lactobacillus paracasei (Figure 3).

Phage Cocktail Decreases E. coli and Improves Normal Gastrointestinal Function

Moving out of the Petri dish and into the animal model, one group of laboratory mice was given the probiotic B. longum in combination with the diseasecausing E. coli strain H10407.41 A second group of mice was given the same mixture, but with the addition of the **phage cocktail** that targets *E. coli*.

After 24 hours, compared with the control group, animals in the **phage-treated** group gave powerful evidence of a *reduction* in the disease-causing bacteria as can be seen on the following page:

- About a **10-fold decrease** of *E. coli* in the small intestine
- About a **100-fold decrease** of *E. coli* in the colon (large intestine)
- About a **100-fold decrease** of *E. coli* in fecal matter

At the same time, the **phage-treated** animals showed remarkable increases in the beneficial (B. longum) bacteria as follows:

- About **100-fold** <u>increase</u> of *B. longum* in the small intestine
- About **100-fold** increase of *B. longum* in the colon
- About **40-fold** increase of *B. longum* in fecal matter

This study also revealed marked differences in the appearance of tissue samples and digestive function across the different groups of mice.

Mice treated with E. coli and B. longum alone were constipated, and intestinal segments showed swelling, redness, and leaks compared with healthy animals. The mice given **B.** longum and **E.** coli plus the **phage** mixture had normal bowel movements, while their intestinal tissues showed no detrimental changes.

Safety of Phage Therapy

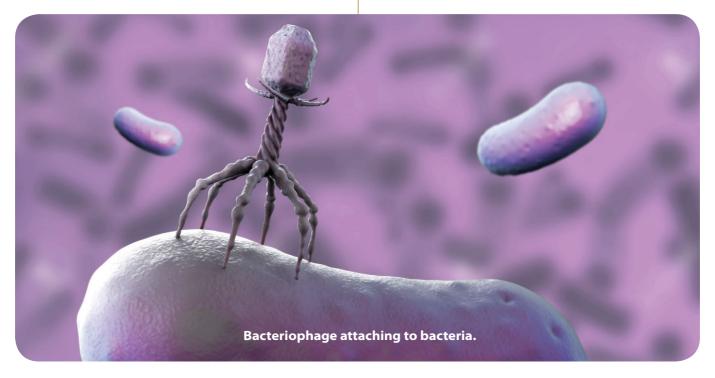
In 1981, Russian researchers reported the successful use of phage therapy to treat antibiotic-associated dysbiosis—displacement of normal intestinal bacteria with pathogenic bacteria—in 500 low-birth-weight infants.

The subjects had all been given antibiotics for at least two to three weeks to treat sepsis and pneumonia, killing off their gut bacteria. They were then administered both a specific bacteriophage and a probiotic (Bifidobacteria) strain. The infants experienced both depletion of their pathogenic bacteria and replenishment of healthy, intestinal bacteria.⁴²

Phage therapy has been used for a wide variety of infections involving numerous pathogens. Therapies have included infections of the gastrointestinal tract, skin, head and neck, bone, chest, and abdomenand have involved an array of pathogens such as Staphylococcus, Streptococcus, E. coli, Salmonella, and Pseudomonas. Success rates range up to 80%-95% for phage therapy with either no, or only mild reversible, side effects. 5,42,43

Bacteriophages were successfully used in numerous human clinical and therapeutic settings and demonstrated an extremely strong safety profile. 43-46

The reason for the very safe interaction between phages and human tissue likely results from human exposure to vast numbers of phages over the entire course of evolution. This naturally high human tolerance to phages contrasts sharply with the risks inherent in compounds that are relatively novel in human evolution—such as drugs.5





end of the "antibiotic era."

In severe cases, physicians desperately search for alternatives to save their patients' lives.

This has prompted excitement about an innovative approach developed more than 100 years ago, but overlooked with the advent of antibiotics.

In fact, long before antibiotics were discovered, Eastern European doctors were successfully neutralizing bacterial infections with **phage therapy**, a treatment aimed at selectively targeting and destroying harmful bacteria.

With the rise in antibiotic-resistant bacteria, **phage therapy** is experiencing a revival among the scientific community because of its effectiveness and safety profile. The potential for phage therapy is so great that just last year the National Institutes of Health sponsored a symposium titled:

"Bacteriophage Therapy: An Alternative Strategy to Combat Drug Resistance"

Phage cocktails have been shown to effectively treat common bacterial invaders, including staph, strep, and E. coli. 1-3

Phage therapy also beneficially encourages growth of healthy bacteria in the gut microbiome, which can also reduce harmful bacteria.

With so much scientific investigation into the multiple health benefits of a balanced microbiome, **phage therapy** is rapidly emerging as a new venue to enhance the benefits of probiotics.

Summary

In addition to being associated with major agerelated disorders, an unbalanced intestinal microbi**ome** (dysbiosis) can also be responsible for general "blah" feelings as well as vague but troubling intestinal symptoms that plague aging individuals.

A novel approach to bringing the gut **microbi**ome back into a healthy balance uses safe bacteriophages to selectively reduce harmful bacteria while encouraging beneficial **probiotic** organisms to flourish.

Called **phage therapy**, this technique is harmless to humans, but deadly for specific, troublesome bacteria.

Studies show that when **probiotic** organisms are accompanied by targeted **phage** therapy, the beneficial bacteria grow up to thousands of times their baseline rate, thanks to the removal of the more aggressive microbes.

Phage therapy shows real promise in relieving the functional changes and tissue damage wrought by *dysbiosis* and could be especially valuable for aging individuals experiencing intestinal discomfort, sleep disturbances, or general malaise.

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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Natural Approach to Guard Against DEEP VEIN Thrombosis

Long periods spent sitting—behind a desk, in a car, on a long flight, or on a couch—increase the <u>risk</u> of **deep vein thrombosis (DVT)**, a blood clot that forms mainly in a deep vein in the leg, which can lead to a *pulmonary embolism*, a condition that is often fatal.^{1,2}

Practically anyone can be at risk, and the statistics are frightening. According to the American College of Cardiology, those who **sit** more than **four hours** a day have a **48%** increased risk of mortality from blood clots that originate in the veins.³

Through a series of well-designed studies, scientists investigating certain natural extracts found that they can <u>block</u> these **blood clots** from forming and can help break down small clots before they grow.⁴⁻⁷ They also found that these extracts safely restore the natural anti-clotting and clot-reversal processes, making a dangerous clot less likely to form and quicker to resolve if it does.⁷⁻¹⁰

When tested on **humans** in a placebo-controlled study, those taking a dual **plant extract** experienced **zero** cases of deep vein thrombosis compared to **5.4%** afflicted in the control group.⁶

Veins have valves, which work with the natural pumping action of the leg muscles to prevent backflow of blood into the lower limbs. With age those valves tend to leak. In people who don't move around much, blood tends to pool in their legs.

According to the Centers for Disease Control and Prevention (CDC), as many as **900,000** Americans could be affected by **venous thromboembolisms** (blood clots) every year.¹¹ Compare that to epidemics like all forms of **cancer**, which kill about **570,000** Americans annually.

As sedentary lifestyles become common, the threat of **deep vein thrombosis** continues to grow and claim an ever larger number of lives.

Research conducted by the American College of Cardiology found that individuals that spent four hours or more a day sitting compared to those who spent less than 2 hours had an approximate **125%** increase in risk of **cardiovascular** events. This association was independent of traditional risk factors like smoking, hypertension, physical activity, and body mass index.³ Some even consider sitting to be the new smoking!¹²

How Deadly Venous Thromboembolism Occurs

A deep **vein thrombosis** can occur quickly, often at first with no warning symptoms. But when this clot breaks apart, symptoms occur with a deadly vengeance. Large pieces of the clot travel silently through the circulatory system and eventually block blood flow into the **lungs**. This sudden event produces a **pulmonary embolism**, a blockage that can severely reduce, and even entirely stop, critical blood flow to the lungs—an event that is commonly fatal *within minutes*.

About **30%** of those with a venous thromboembolism will die within one month, and about **25%** of those fatalities will occur as sudden death. And about **33%** of survivors have a recurrence within 10 years.¹¹ Venous thromboembolisms are estimated to kill up to **100,000** Americans annually.¹¹

Lifestyle changes can reduce the risk, such as quitting smoking, regularly exercising, and eating a healthy diet, but these are often insufficient to prevent a deadly catastrophe. Anticlotting drugs involve a risk of undesired bleeding, ^{13,14} and compression stockings have shown limited effectiveness. ^{15,16} The reason that these approaches provide limited protection is that with physical pooling of blood, natural clot-breakdown systems slowly lose pace with the body's clot-forming systems. ^{17,18}

Clearly, a new defensive strategy is needed.



Possible Symptoms of Deep Vein Thrombosis

Deep vein thrombosis often occurs without warning symptoms. But if you do notice any of the following indications, call your doctor, especially if they appear suddenly:²⁷

- Swelling in one or both legs
- Tenderness or pain in one or both legs, even if it's only when standing or walking
- · Warm skin on your leg
- Red or discolored skin on your leg
- · Veins you can suddenly see
- Tired legs

A natural, dual plant extract formula has been studied based on its ability to simultaneously inhibit venous clot formation *and* promote venous wall elasticity.

Combating Deep Vein Thrombosis

After searching for natural, deep vein thrombosisinhibiting interventions, scientists identified <u>two</u> ingredients that demonstrated powerful preventive effects:

- 1. **Nattokinase**, an enzyme extracted from soybeans fermented with the bacterium *Bacillus natto*, and
- 2. French maritime pine bark, a natural extract rich in polyphenols.

Nattokinase was shown in studies to break down *fibrin*—long, strand-like molecules that make up the main protein found in clots—and its precursor, *fibrinogen*, both of which are involved in **red blood cell-induced** clot formation.^{8,19,20} Nattokinase decreased levels of other factors in the blood-clotting cascade, while raising levels of factors that prevent clotting. Specifically, nattokinase reduces the activity of two proteins (factor VIII and factor VII) that can produce unwanted clotting when elevated. No adverse effects or undesirable bleeding were reported.⁴

French maritime pine bark extract was demonstrated to reduce platelet aggregation, while increasing the activity of a blood flow-boosting *enzyme* that generates *nitric oxide* in blood vessels. Nitric oxide plays a critical role in regulating vascular function, which reduces thrombotic risks.

Nattokinase and **pine bark extract** were shown to work together to prevent clots as well as to improve the microcirculation of the legs.^{9,21}

French maritime pine bark extract was also found to inhibit the action of "protein-melting" matrix metal*loproteinase* enzymes. These enzymes would otherwise degrade elastic proteins in the blood vessel walls, making them stiff and reducing blood flow. 10,22

Given these observations, scientists recognized that these two extracts could result in significant prevention of deep vein thrombosis by:4,7-10,19-22

- 1. Inhibiting unwanted clot formation within
- 2. Improving microcirculation in the veins of the legs, and
- 3. Promoting elasticity of vessel walls.
- 4. Inducing breakdown of fibrin clots.

Now let's look at a more detailed evaluation of these ingredients.

Nattokinase Breaks Down Blood Clots

Before designing human trials, scientists conducted animal studies that clearly demonstrated the beneficial effects of nattokinase.

Studies in dogs showed that nattokinase produces a mild—but steady—increase in the rate of fibrin degradation in the blood. This effect works to prevent clots and to reduce the size and toughness of any existing ones.8

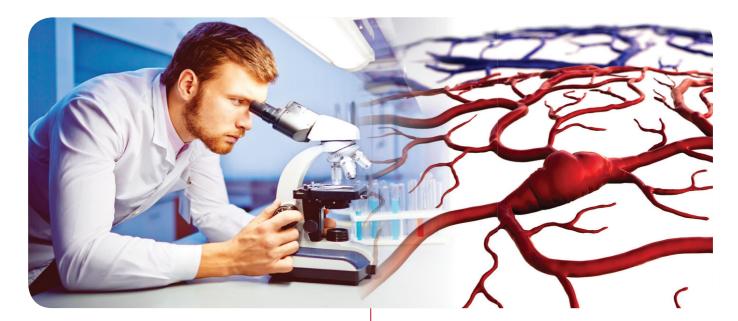
When **nattokinase** was given to dogs with experimentally-induced blood clots, researchers were literally able to watch the clots break down in real time using a type of X-ray technology called angiography.8

What You Need to Know

Preventing **Deep Vein Thrombosis**

- Spending long periods sitting can lead to deep vein thrombosis, which can suddenly, and without warning, trigger a pulmonary embolism.
- The medical establishment provides no safe or practical solutions to prevent this deadly risk.





Then, in spontaneously hypertensive rats—a standard model for studies of high blood pressure in humans—nattokinase was demonstrated to break down fibrinogen (the precursor to fibrin) in the blood. And by a related mechanism, nattokinase reduced blood pressure, potentially by preventing conversion of the hormone angiotensin into its active, blood pressure-boosting form.²³

A human study evaluating the effects of nattokinase found that it improves markers of coagulation. For this study, researchers recruited volunteers comprised of healthy individuals, cardiovascular disease patients, and dialysis patients. On a daily basis for two months, subjects took two capsules of **nattokinase**, each containing **2,000 fibrinolytic units**.⁴

The researchers found that all three groups demonstrated significant <u>decreases</u> in procoagulation factors VII and VIII, and fibrinogen compared to baseline, suggesting that nattokinase works equally well in individuals with normal and impaired endothelial and coagulation function.⁴

No adverse effects were seen in this study, 4 and safety tests completed on nattokinase confirmed its low-risk status. 20

Preventing Venous Thromboses with French Maritime Pine Bark Extract

Scientists also conducted human studies on French maritime pine bark extract's effects on the risks of venous thrombosis.

First, they enlisted 198 people at risk for deep or superficial venous thromboses during flights ranging from seven to 12 hours, with an average flight time of 8.25 hours.⁵ Test subjects were randomly assigned to either the control or test group. The test group took two capsules, each containing **100 mg** of French maritime pine bark extract, two to three hours preflight, two more capsules six hours into the flight, and one capsule the day after. The controls took placebos at the same intervals. All subjects underwent ultrasound scans of their leg veins within 90 minutes before and after their flights to detect clots.

The French maritime pine bark extract-treated group showed **no** blood clots for an event rate of **0.0%**. But the placebo group showed **four** superficial venous thromboses and one deep venous thrombosis, equivalent to an event rate of **5.15%**. No adverse events were observed.⁵ This study presented evidence to support that French maritime pine bark extract can prevent dangerous blood clots during prolonged sitting.⁵

Guarding against Lower Leg Fluid Accumulation

The same scientific team conducted a similar study to evaluate ankle swelling during long-haul air flights.²⁴ Aside from being uncomfortable, **ankle swelling** is an excellent indicator of poor blood return up the veins of the legs, making it a great way to assess the risk of deep vein thrombosis.

The team enlisted 169 volunteers at risk of deep vein thrombosis due to remaining seated during a long flight. The same dose of **200 mg** of French maritime pine bark extract was given at the same intervals as in the earlier study. Using standard measurements, edema (swelling) was measured before and after flights, as well as the rate of swelling.

Compared to preflight levels, the edema score in **placebo** subjects was <u>increased</u> by **58.3**%. The edema score in the French maritime pine bark extract-supplemented volunteers increased only **17.9**%. This dramatic decrease in edema score represented a significantly reduced thrombotic risk.²⁴ Similarly, the ankle swelling rate was increased during the flight by a mean of **91**% in controls, while the French maritime pine bark extract recipients showed only a **36**% increased ankle swelling rate, a much safer rate.²⁴

These studies showed the capacity of French maritime pine bark extract to reduce the risk of deep vein thrombosis without any of the side effects of anticlotting drugs. However, the question remained as to whether this extract was superior to compression stockings, which are known to be safe but not necessarily effective in reducing post-thrombotic syndrome. Post-thrombotic syndrome is a common complication of an otherwise localized deep vein thrombosis, in which blood pools in the affected leg because it cannot return to the heart—causing skin swelling, thinning, and discoloration and sometimes, painful, infection-prone leg ulcers.

To settle this issue, scientists conducted a study comparing French maritime pine bark extract to compression stockings in their ability to prevent post-thrombotic syndrome.²⁵ In this study, 156 patients who had experienced a single, major episode of deep

Risk Factors for Deep Vein Thrombosis

Deep vein thrombosis (DVT) and the deadly pulmonary embolism it can trigger (venous thromboembolism) is often the result of spending long periods sitting or standing. But other factors can put some individuals at an especially high risk. These risk factors include:²⁷

- · use of oral contraceptives,
- · advanced age,
- smoking,
- pregnancy,
- · severe obesity,
- · limited mobility,
- recent surgery or trauma,
- · sitting on long-haul flights, and
- cancer.

vein thrombosis were divided into three groups. For 12 months, one group used the compression stockings, the second group took **50 mg** of French maritime pine bark extract three times daily, and the third group used both the stockings and the same daily French maritime pine bark extract regimen.²⁸ The researchers measured edema score, ankle circumference, and volume of the previously deep-vein thrombosis-afflicted leg compared with the other leg.

These confirmatory findings indicate that this novel dual-extract formula helps prevent deep vein thrombosis.

Their findings soundly confirmed the superior effectiveness of French maritime pine bark extract:²⁵

- 1. In the **compression stocking-only** group, two new deep vein thrombosis cases occurred in the first six months, compared with <u>no</u> new deep vein thrombosis cases in either of the **pine bark** extract groups.
- 2. After the first six months, French maritime pine bark extract alone proved significantly more effective than compression stockings alone for relieving symptoms of edema (while the combination of both was better still).
- 3. Leg volume and ankle circumference measurements showed French maritime pine bark extract-plus-stockings to be superior to stockings alone.
- 4. In the microcirculation (blood flow in the tiniest vessels), French maritime pine bark extract—but not compression stockings—enhanced blood flow, raised oxygen levels in circulating blood, and decreased carbon dioxide levels, demonstrating improved function.

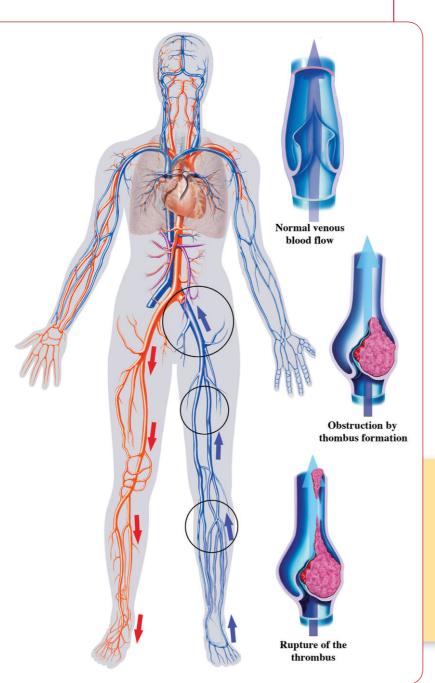
Human Clinical Trial Combining Nattokinase and French Maritime Pine Bark Extract

Scientists set out to test a formulation combining **nattokinase** and **French maritime pine bark** extract in a randomized, placebo-controlled human trial.⁶ All 204 passengers on a New York-to-London flight were instructed in deep vein thrombosis-prevention techniques: isometric exercises, standing and moving for

five to ten minutes, and keeping hydrated. Passengers were randomly assigned to receive either capsules of placebo or capsules of the proprietary blend of nattokinase and French maritime pine bark extract. All subjects took the **blend** two hours predeparture and again six hours later. Ultrasound scans were done before and after the flight to detect clots.

Passengers taking the supplement had **zero** deep vein thrombosis cases. However, five of the control passengers developed a deep vein thrombosis, and two others developed superficial clots, for a total of **seven** events—a 5.4% DVT rate among controls, compared to a **0.0**% rate among the test subjects. The scientists also measured leg swelling, which was equal between the two groups preflight. Edema increased by 12% in the controls. But edema decreased by 15% in the supplemented passengers.6

These findings confirm that this novel dual-extract formula helps prevent deep vein thrombosis in individuals who spend long periods sitting and reduces the risk of sudden death from a resultant pulmonary embolism. No adverse side effects were reported.



Summary

Deep vein thrombosis is a serious risk for anyone who spends long periods sitting and can lead to a deadly pulmonary embolism.

The two novel ingredients described in this article were shown to protect against venous thrombosis.

These extracts inhibit unwanted venous clot formation, improve leg microcirculation, and promote vessel-wall elasticity.

In a placebo-controlled human trial, these two nutrients prevented deep vein thrombosis in <u>all</u> volunteers who supplemented with it and decreased leg swelling.

While the medical establishment provides no safe or practical solutions, these two agents are available to augment the effects of taking frequent breaks from any kind of prolonged sitting.

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

Thromboemmbolism

This diagram illustrates normal venous blood flow, venous obstruction by a thrombus formation, and the rupture of the thrombus. A blood clot such as this forming in the leg of an individual with deep vein thrombosis (DVT) can break off without warning and lodge in the lungs or other organs, often resulting in death.

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Too Much TV Can Lead to **Pulmonary Embolism**

Our parents always used to say that watching too much TV wasn't good for us. It turns out they may have been more right than they realized. A recent study published in the journal Circulation found that a sedentary, couch-potato lifestyle can lead to fatal blood clots.28

Starting in 1988, researchers from Japan's Osaka University recruited more than 86,000 subjects and had them keep track of how much television they watched over a two-year period. After 19 years, researchers found that 59 study participants had died of a pulmonary embolism—a blood clot in their lungs. Lack of activity causes a person's blood flow to slow down, which can lead to a blood clot, usually in the pelvis or leg. The blood clot can then travel to the lungs and become wedged in a small blood vessel in the lungs with fatal results.

The researchers calculated that the study subjects had a 40% increased risk of incurring a pulmonary embolism for every two hours of TV they watched each day beyond a threshold of 2.5 hours. The risk for participants who watched five hours or more was **2.5 times** greater than the risk for those who watched under 2.5 hours.

To avoid an embolism, study co-author Dr. Hiroyasu Iso recommends that, while watching TV, you get up "after an hour or so" and "stand up, stretch, walk around. Or while you're watching TV, tense and relax your leg muscles for five minutes."

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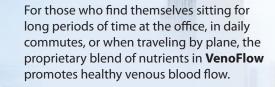
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MacuGuard® Ocular Support • 60 softgels, Item #01992 Offers triple eye protection with <i>meso</i> -zeaxanthin, lutein, and <i>trans</i> -zeaxanthin. This product is <u>not</u> needed by those already taking Health Booster , which contains these same ingredients.	\$25	\$15.75 (four-bottle purchase)

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Mitochondrial Energy Optimizer with BioPQQ® • 120 capsules, Item #01868 This glycation-protection formula helps maintain cellular function, protein structural integriand mitochondrial biogenesis. It contains high potency carnosine along with R-lipoic acid, benfotiamine, luteolin, and taurine. These ingredients cost far more if taken separately.		\$43.20 (four-bottle purchase)
Mega Green Tea Extract • 725 mg, 100 lightly caffeinated vegetarian capsules, Item #00953 A highly concentrated 98% polyphenol extract delivering 45% of the health-promoting cated	\$30 hin EGCG.	\$16.20 (four-bottle purchase)
Life Extension Mix™ • 315 tablets, Item #02155 This high-potency multinutrient now includes apigenin , which has been shown to ease inflammation and support healthy cell growth and differentiation, and SelenoExcell® , a natural selenium source. (Also available in capsule and powder form.)	\$80	\$46.80 (four-bottle purchase) \$39.38 (ten-bottle purchase)
AMPK Activator • 90 vegetarian capsules, Item #01907 Activating AMPK "turns off" many of the destructive factors of aging, enabling cells to return to their youthful vitality. Research shows that the <u>two</u> plant extracts contained in this formula promote AMPK activation.	\$48	\$29.70 (four-bottle purchase)
Cognitex® with Brain Shield® • 90 softgels, Item #01896 Optimal support for the brain. Includes gastrodin, alpha-glyceryl phosphoryl choline, vinpoc phosphatidylserine, uridine-5′-monophosphate, and more. Available with or without pregne	\$ 60 retine, nolone.	\$35.10 (four-bottle purchase)
Super K with Advanced K2 Complex • 90 softgels, Item #01834 Provides two forms of vitamin K2 (1,000 mcg of immediate-release MK-4 and 200 mcg of long MK-7), along with 1,500 mcg of K1.	acting \$30	\$18.23 (four-bottle purchase)
Triple Action Thyroid • 60 vegetarian capsules, Item #02003 A combination of ashwagandha, guggul, and Korean ginseng extract work in synergy to support healthy thyroid function.	\$36	\$21.60 (four-bottle purchase)
Memory Protect • 36-day supply, Item #02101 This new product contains microdose lithium and proline-rich polypeptide, which have be found to block and (in the case of proline-rich polypeptide) to reverse cognitive decline.	een \$24	\$14.40 (four-box purchase)
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Super R-Lipoic Acid • 300 mg, 60 vegetarian capsules, Item #01208 Superior efficacy compared to alpha-lipoic acid—supplies 240 mg of stabilized R-lipoic acid	\$49	\$30.38 (four-bottle purchase)
Advanced Bio-Curcumin® with Ginger and Tumerones • 30 softgels, Item #01924 Absorbs up to 7-times better, with added benefits of ginger and turmeric extracts.	\$30	\$18.23 (four-bottle purchase)
Extend-Release Magnesium • 60 vegetarian capsules, Item #02107 Provides immediate-release magnesium citrate along with a 6-hour extended-release magnesium for <i>optimal</i> benefits.	\$13	\$7.88 (four-bottle purchase)



How Vegetable Extracts Protect Against Cancer

Apigenin is a polyphenol found in vegetables such as parsley and celery. It is receiving increased attention as a low-cost nutrient to protect against common cancers.

What makes **apigenin** so fascinating is how it functions to starve cancer cells, promote cancer cell destruction, and protect cellular DNA against environmental toxins (that can result in future malignancies).

Compounds such as indole-3-carbinol (I3C) are found in cruciferous vegetables. These cruciferous compounds have been shown to work in complementary ways with apigenin (non-cruciferous) to combat cancer and other age-related diseases.

According to a study published in the **International Journal** of Oncology:

"Cancer prevention through diet may be largely achievable by increased consumption of fruits and vegetables. Considerable attention has been devoted to identifying plant-derived dietary agents which could be developed as promising chemopreventives. One such agent is apigenin."1



Apigenin Protects Cells from Cancer

Apigenin fights *oxidative stress* and *inflammation*—two factors that play a role in cancer development.²⁻⁹ Oxidative stress and inflammation generate DNA damage that can lead to uncontrolled proliferation of nonfunctioning cells, i.e., cancer.^{10,11} But that's just one aspect of how apigenin functions.

Unique mechanisms of apigenin have led researchers to intensely evaluate it.¹² What they've discovered is that *apigenin* has a host of other anticancer properties:

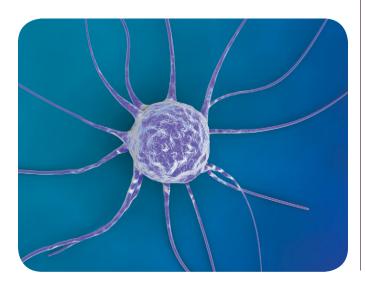
1. Apigenin Stops Cancer Cells from Replicating

Apigenin attacks cancer at a variety of stages and in different ways. At every step, apigenin seems to aggressively stop cancer's various pathways. Apigenin has the ability to stop cancer cells from replicating, to reduce their invasiveness, and to slow their growth. Scientists believe this is largely related to its ability to shut down **nuclear factor-kappa B** (NF-kB). When activated, NF-kB leads to a flood of proinflammatory molecules that can promote tumor growth and the spread of cancer. 15,16

In an animal study of nonmelanoma skin cancer, apigenin inhibited the production of inflammatory signaling molecules known to promote tumor proliferation.¹⁷

2. Apigenin Causes Cancer Cells to Die Off

In a cell study of chronic lymphocytic leukemia (a common malignancy in older adults), apigenin prevented the DNA mutations tied to cancer, while promoting the naturally-occurring cell death (*apoptosis*) that cancer cells otherwise evade.¹⁸



Apigenin may promote apoptosis by **reactivating** an important cancer-suppressor gene called **p53**. Inactivation of p53 is a common feature of cancer cells, which results in the cell's loss of control over when to replicate and when to die naturally. By restoring p53 activity, apigenin essentially allows cancer cells to die a natural death. 12

3. Apigenin Cuts off Cancer Cells' Ability to Grow

In additional to causing cancer cells to die off naturally, studies have also shown that apigenin modulates a host of factors that can give cancer a promotional boost once it gets started.

Apigenin has repeatedly been shown to regulate the *insulin-like growth factor* signaling pathway known to promote the growth of prostate cancer cells when deregulated. Under the influence of apigenin, those cells quit their explosive growth and proceed to kill themselves off by apoptosis.¹²

4. Apigenin Starves Cancer Cells

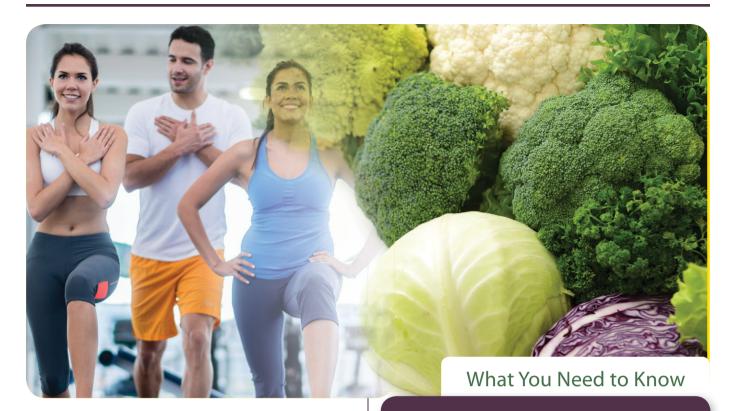
Finally, apigenin appears capable of literally *starving cancer cells into submission* through several unrelated, but complementary, mechanisms.

First, apigenin suppresses the expression of a protein essential for transporting glucose into cancer cells.^{22,23}

Apigenin further promotes this "energy crisis" by going after cancer cells' mitochondria, the tiny intracellular power plants that generate energy. When mitochondria from human liver cancer cells were treated with apigenin, their membranes became leaky to the extent that it destroyed affected cancer cells.²⁴

Based on the mechanisms by which apigenin thwarts cancer at every turn, new human studies are needed to further explore this impressive cancerdestroying polyphenol. The **underlying mechanisms** discussed here have been found to be effective in animal models of **leukemia** and the following solid tumor malignancies:

- Prostate^{14,25}
- Larynx (voice box)²²
- Leukemia¹⁸
- Liver (hepatocellular carcinoma)²⁶
- Pancreatic⁹
- Skin¹⁷
- Breast²⁷



Apigenin Protects Brain Cells

Alzheimer's and Parkinson's are two common neurodegenerative diseases.²⁸⁻³⁰ Together, they cause untold misery for patients, their families, and their other caregivers. Numerous studies show apigenin's ability to reduce some of the known contributors to neurodegeneration.

For example, apigenin fights *excitotoxicity*, the neuronal damage that occurs over a lifetime of intense brain cell stimulation.^{3,6} This is critical since excitotoxicity promotes brain cell death and dysfunction in both Alzheimer's and Parkinson's diseases.^{31,32}

Apigenin also was shown to protect dopamine-producing cells of deep brain centers affected by Parkinson's disease. This important action reduces neuroinflammation and the activation of inflammatory cells in the brain.^{33,34}

When applied to brain cells in culture, apigenin protected those cells from toxicity induced by **beta amyloid**, the toxic "junk" protein found in abundance in the Alzheimer's brain.³⁵

Finally, given the importance of elevated blood sugar in the development of Alzheimer's (it has been called "Type III diabetes" by some researchers), it is interesting to note that apigenin attenuates the cognitive decline seen in adult diabetic rats.⁴ Animals treated with apigenin have demonstrated improved learning and memory retention in a mouse model of Alzheimer's disease.³⁶

Health Benefits of Vegetable Extracts

- Vegetable extracts present in broccoli, celery, parsley, kale, Brussels sprouts, and many others—have long been revered for their health-promoting benefits.
- It is now recognized that four compounds, apigenin, I3C, DIM, and BITC, are responsible for the majority of that protection, and in very specific and complementary ways.
- All four have powerful cancer-protective effects, attacking and preventing malignancies through a multitude of overlapping mechanisms.
- Apigenin also has exceptional cardiovascular, metabolic, and neuroprotective properties, while I3C/DIM promote cardiovascular and liver health.
- Consumption of these plant compounds provides broad-spectrum protection against many of the most common symptoms of aging.



Apigenin, I3C, DIM, and BITC are compounds found in vegetables that offer wide-ranging protection against the factors that damage our DNA.

Apigenin Promotes Cardiometabolic Health

Oxidative stress and inflammation are deadly to heart and blood vessel cells. Inflammation/oxidation can also induce the kind of damage in liver and fat tissues that can promote weight gain, diabetes, and other metabolic changes that raise cardiovascular disease risk. Apigenin can contribute substantially to protecting against all of these effects.

At the most fundamental level, apigenin has been shown to prevent new cholesterol molecules from being synthesized in liver cells, which can reduce the amount of cholesterol in circulation.³⁷ In a similar fashion, animal studies show that apigenin lowers blood sugar levels by decreasing insulin resistance, decreasing elevated insulin levels, and decreasing the formation of new glucose in the liver.^{38,39}

Studies in diabetic rats reveal that apigenin can improve the function of the endothelial cells that line arteries and directly control blood flow and pressure.³⁹

Preclinical studies show that apigenin helps protect the heart muscle against ischemia/reperfusion injury, the serious damage that occurs in the minutes to hours following a heart attack or stroke, when oxygen-starved (ischemic) tissue is suddenly flooded with oxygen-rich blood as circulation is restored (reperfusion).⁴⁰⁻⁴²

I3C: Complementary Tissue Protection from Cruciferous Vegetables

I3C (indole-3-carbinol) is a major component found in cruciferous vegetables. While I3C has many overlapping activities with apigenin, it also adds a substantial number of unique functions.⁴³ Many of these benefits are also produced by *3,3'-diindolylmethane* (**DIM**), a condensation product of I3C molecules.^{44,45}

Through these unique functions, I3C and DIM provide complementary protection against cancer, heart disease, and more. Let's take a look.

I3C/DIM: Unique Protection against Cancer

The most promising of I3C's cancer-fighting properties has to do with its impact on enzymes that metabolize the sex hormones *estrogen* and *testosterone*.⁴⁶

For example, I3C and DIM have been found to reduce a carcinogenic form of estrogen called **16-alpha-hydroxyestrone** while boosting beneficial **2-hydroxyestrone**. A large study of over 10,000 women showed that those with highest amounts of **2-hydroxyestrone** compared to 16-alpha-hydroxyestrone had a remarkable **42**% lower risk of breast cancer. Since hormone-dependent cancers such as breast and ovarian tumors thrive on unhealthy hormone balances, this is an important step in cancer prevention. ^{43,48-50}

There's also considerable evidence that I3C modifies the function and expression of estrogen receptors on cells. Studies show that I3C downregulates the expression of **estrogen receptor alpha**, which

is known to cause cancer-promoting cellular changes when it is overexpressed.⁵⁰⁻⁵³ Turning down estrogen receptor alpha allows greater influence for the *protective* **estrogen receptor beta**, which further reduces estrogen-dependent cancer risk.⁵⁰

I3C/DIM Promote Cardiometabolic Health

I3C and DIM have multiple antiobesity effects.

Obese mice treated with I3C showed decreased body weight, fat accumulation, fat-mediated release of inflammatory cytokines, and fat infiltration by inflammatory cells—all of which produce obesity-associated health risks.⁵⁴⁻⁵⁷

I3C and DIM also reduce blood sugar, insulin, and markers of sugar-induced protein damage in mice on high-fat diets. These are effects that contribute to reduced atherosclerosis and cardiovascular disease risks. 58,59

I3C/DIM Provide Wide-Ranging Liver Protection

I3C and DIM also have impressive liver-protective properties. These are partly due to their ability to suppress oxidative stress, in part through their activation of AMPK, and in part through their anti-inflammatory properties. ⁶⁰⁻⁶²

These mechanisms of action protect the liver against fat accumulation, inflammatory changes, malignancies, and even *fibrosis*, the scarring and toughening that precedes liver failure.⁶³

In fact, animal studies show that I3C and DIM can prevent fatty liver disease caused both by alcohol and a high-fat diet.^{60,61}

BITC Offers Complementary Cancer Protection

Benzyl isothiocyanate (**BITC**) is another compound found in cruciferous vegetables that provides cancer protection that complements those of other cruciferous compounds.

One of its most startling properties is its ability to inhibit **cancer stem cells**. Many malignancies recur or fail to respond to treatment because of just a few cancer stem cells that can "hide out" and emerge after treatment is complete.

Both lab and animal studies show that BITC inhibits breast cancer stem cells, while also suppressing cell signaling molecules that contribute to the "stemness" of such cells.⁶⁴⁻⁶⁶ As a result, BITC has been found to be effective against numerous types of cancer.

For example, an animal study demonstrated that BITC reduced the number of prostate cancers in cancer-prone mice.⁶⁷ BITC also prevents the stimulation of breast cancer growth induced by high-fat diets, prevents invasion and new blood vessel formation in brain and head and neck cancers, and induces cell death by apoptosis in lung cancers.⁶⁸⁻⁷¹

Summary

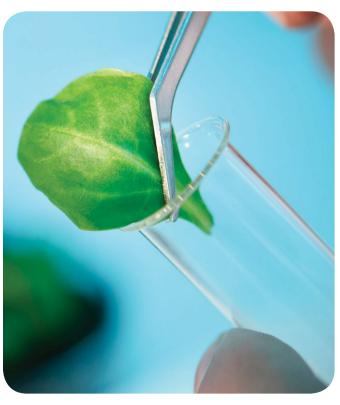
Apigenin, I3C, DIM, and BITC are compounds found in **vegetables** that offer wide-ranging protection against the factors that damage our DNA.

These compounds work in a complementary way to protect our bodies against cancer, cardiovascular and metabolic diseases, neurodegeneration, and even liver disease.

Together, these **botanical compounds** represent the virtues of consuming more healthy **vegetables** in ones' everyday diet.

For those who cannot consistently eat large amounts of celery, parsley and other vegetables, **apigenin** is available in low-cost multinutrient formulas that provide a wide variety of beneficial plant extracts.

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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Selenium's Impact on Cancer Reduction

Selenium is a trace mineral critical for human life.

While there are many different <u>forms</u> of selenium, not all of them provide the same health benefits. This article presents data on the unique benefits of **selenium-enriched yeast**.

While selenium can be found in foods such as Brazil nuts, pinto beans, and beef, the amount of selenium that is obtained from our diet is highly uneven. This is because the amount of natural selenium in the soil fluctuates from region to region.

Many areas of the United States such as Texas, the Southwest, lower Southeast and Northwestern mountain states have selenium <u>deficient</u> soil. Therefore foods grown in these regions will not be rich in selenium.¹

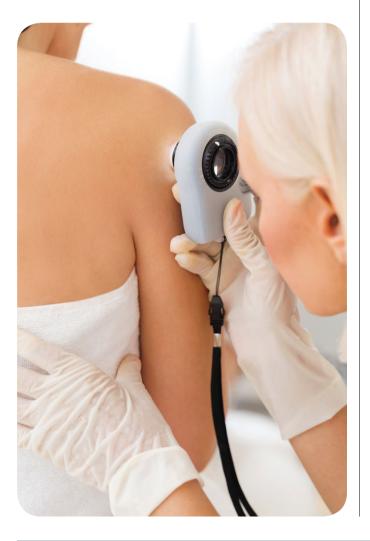
In the past year, two new studies reveal that selenium status is suboptimal in many people in the industrialized world, and that <u>low</u> selenium status raises the risk for colorectal cancers and cancers of the liver and gallbladder.^{2,3}

Ample selenium has been shown to be a relevant factor in protecting against a uniquely male malignancy, prostate cancer.⁴⁻⁹

But men aren't the only ones at increased cancer risk when their selenium levels drop, and prostate cancer is far from the only cancer affected by poor selenium status. Back in 1983, a group of researchers began a human study to examine the effects of **selenium-enriched yeast** on skin cancer.

The researchers found that subjects supplementing with this yeast-based selenium had an approximately 46%-63% reduction in the risk of colorectal, lung, and prostate cancers, with a 50% reduction in the risk of mortality for all cancers. The results were later published in the *Journal of the American Medical Association*⁷ and made **headline news** stories around the world. This generated intense interest in the scientific community in the protective power of selenium.

Recent studies amplify the importance of maintaining satisfactory selenium levels and their impact on cancer risk reduction.



Selenium Lowers Cancer Risk

The association between inadequate selenium status and the risk for many different cancers naturally leads to the question, "Can supplemental selenium reduce cancer risk?"

The answer is that optimal selenium levels through supplementation provide a protective **function** against the risk of cancer.

Studies going back to the 1970s have generally supported supplemental selenium as a cancer risk-reducing intervention.

As mentioned near the beginning of this article, a randomized, controlled trial published in **1996** demonstrated that **200 mcg** of selenium daily from selenium-rich yeast was associated with a **50%** reduction in the risk of dying from cancer, a **37%** reduction in the risk of developing cancer, and reductions of **58%**, **63%**, and **46%** in the risks for developing colorectal, prostate, and lung cancers, respectively.⁷

Deeper analysis showed that, among men with baseline normal levels of prostate-specific antigen, or **PSA**, a marker of cancer risk, those in the treated group showed an overall **74%** reduction in their risks of prostate cancer.⁸

Colon cancer is a selenium-responsive malignancy, as shown in a placebo-controlled **2013** study that supplemented active subjects with **200 mcg** of selenium, along with zinc and vitamins A, C, and E over a 5-year period. ¹⁰

These study subjects consisted of 411 people with *polyps* removed during screening colonoscopy, indicating their higher risk for colorectal cancers.

In the supplemented group, **38** had a recurrence of polyps, while **62** recurred in the placebo group.

This worked out to a **39**% reduction in the risk of recurrence in those who supplemented with selenium and vitamins, compared with <u>un</u>-supplemented placebo recipients.

A **2015** study shows that cervical cancer may also be preventable with selenium supplementation.¹¹

In this study, 56 women diagnosed at biopsy with cervical intraepithelial neoplasia (precancerous lesions) were randomly assigned to receive **200** mcg of selenium from yeast or a placebo, daily for 6 months.

The findings showed that **88%** of supplemented women had regression of their precancerous lesions, compared with only **56%** in the placebo group. This significant difference was accompanied by decreases in fasting blood sugar, insulin, and insulin resistance, all factors associated with increased cancer and metabolic risks.



Some insight into how selenium supplementation works to reduce cancer comes from a study of blood cells from hemodialysis patients, who are at known increased risk for DNA damage, the precursor of new cancers.12

Forty-two dialysis patients randomly received either selenium-rich yeast or a placebo of standard yeast for 3 months.

The dialysis patients had significantly lower selenium levels than did healthy controls, as expected, but these rose significantly with supplementation.

Markers of DNA damage in circulating white blood cells (a good indicator of damage throughout the body) were three times higher in dialysis patients than in controls at the outset. After 3 months of selenium supplementation, DNA damage markers fell in the dialysis patients, to levels 16% lower than those in healthy controls.

No similar changes were seen in the placebo recipients.

- for more than 25 key enzymes that can't
- Most of those enzymes are involved in systems that protect cells and their DNA from oxidative stress that, unopposed, leads to cancer.
- Studies show that those with lower selenium blood levels are at substantially increased risk for cancers in a variety of organs.
- Selenium supplementation, on the other hand, has been shown effective at reducing cancer incidence and producing protective biochemical shifts that shield cells and DNA from damage.
- No single selenium form, however, provides all of the cancer-preventing benefits available from selenium.
- Instead, a thoughtful regimen that includes multiple forms of selenium is most likely to offer comprehensive cancer- prevention properties.

Not All Selenium is the Same

There are several different forms of selenium and not all of them offer the same health benefits. For example, in a very large and well-publicized 2009 study, researchers from the Selenium and Vitamin E Cancer Prevention Trial (SELECT) reported that, after treating more than 35,000 men with one form of selenium (L-selenomethionine) alone (200 mcg), selenium plus vitamin E, or placebo, they found no significant differences in participants' risk for prostate cancer in any of the groups compared with placebo.¹³

Because this was such a large study, it garnered an outsized share of attention, and many scientists and other readers came away with the impression that selenium has no role in cancer prevention.

It should be noted that multiple scientific studies dating back to the 1970s show that different forms of selenium provide a spectrum of protection against cancer. Life Extension® long ago discussed the importance of including more than one form of selenium in one's daily supplement program.

Selenium in a variety of forms has been shown to produce significant reductions in cancer risks.

One form of selenium that has received relatively little attention is selenium-enriched veast, obtained from high-selenium brewer's yeast.

The study discussed near the beginning of this article, which involved 1,312 patients over a total of 8,271 person-years, found that daily supplementation with selenium-enriched yeast (200 mcg/day), had a 50% reduced risk of dying from cancer, and a 37% decreased risk of developing cancer, compared with placebo recipients.7

Further, with regard to specific cancers, the study found reductions of 46%, 58%, and 63% in the risks of lung, colorectal, and prostate cancers, respectively.⁷

> Selenium in its different forms has demonstrated cancer-preventive effects.



As of this writing, selenium status has been significantly associated with lower cancer incidence, as shown in the following table:

Type of Cancer	Impact of Selenium Status
Bladder ¹⁸	39 % reduction with higher selenium levels
Lung ¹⁹	90 % reduction with higher selenium levels
Larynx (throat) ¹⁹	77% lower with higher selenium levels
Prostate ⁴	71% reduction with higher selenium levels
Head-and-Neck ²¹	45% reduction with higher selenium levels
Lung ²⁰	140% higher with <u>low</u> selenoprotein levels



Different Forms of Selenium

The different forms of selenium have been shown to provide various protective properties against cancer, oxidative stress, DNA damage and even shielding against toxic metal poisoning. For this reason, it is recommended that it is best to ingest a **200 mcg** "cocktail" daily of various selenium forms to provide broad-spectrum coverage against the diseases of aging.^{14,15}

Here are brief descriptions of well-studied forms, highlighting how each can contribute to seleniumbased protection against cancer.

Sodium selenite is a simple chemical salt of sodium and selenium. This form of selenium has the ability to ramp up our natural immune system to find and destroy tumor cells. 16,17

Selenium-methyl L-selenocysteine triggers cancer cell suicide (apoptosis) and also acts on more advanced cancer cells that have lost the fundamental "suicide gene."¹⁴

Selenium from yeast provides advanced protection against oxidative stress and resulting DNA damage, to reduce the risks that a cell will undergo transformation into a malignancy.⁶

Because optimum cancer prevention requires protection from DNA damage, enhanced self-destruction of malignant cells, and a boosted immune system, the benefits of using multiple <u>forms</u> of **selenium** are obvious.

Summary

Preventing cancer requires a multifactorial approach.

One mechanism involved in cancer development is oxidative stress and the DNA damage that it causes.

Selenium is a natural element capable of fighting oxidative stress and has been shown to be effective in preventing cancer and the biochemical changes that precede it.

But because one large study found no significant effects on prostate cancer risks using just one form of selenium (L-selenomethionine), many have turned away from this valuable mineral.

The fact is that selenium in its different forms has demonstrated strong cancer-preventive effects.

Recent studies amplify the importance of maintaining satisfactory selenium levels.

Don't be fooled by spurious science, often based on flawed studies. Selenium-enriched yeast supplements have demonstrated strong potential for reducing cancer risk. Other forms of selenium contribute to enhanced immunological destruction of early cancers, trigger cancer cell "suicide," and protect tissues from oxidative stress.

Overall, the evidence points to a daily **selenium** dose of **200 mcg** to reduce cancer risk, which should ideally utilize multiple forms to obtain comprehensive protection.

No single nutrient should be relied on to protect against cancer. This magazine consistently reminds readers of dietary and lifestyle factors that play huge roles in one's risk of developing malignant disease. ●

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

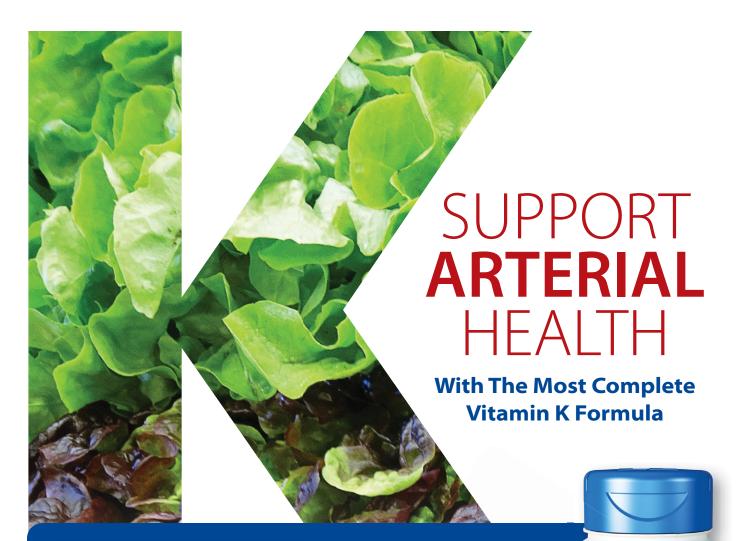
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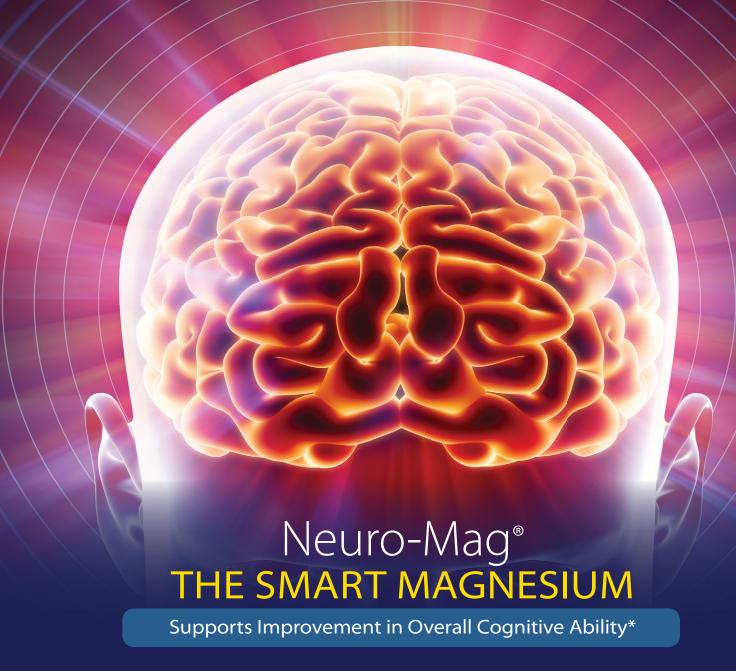
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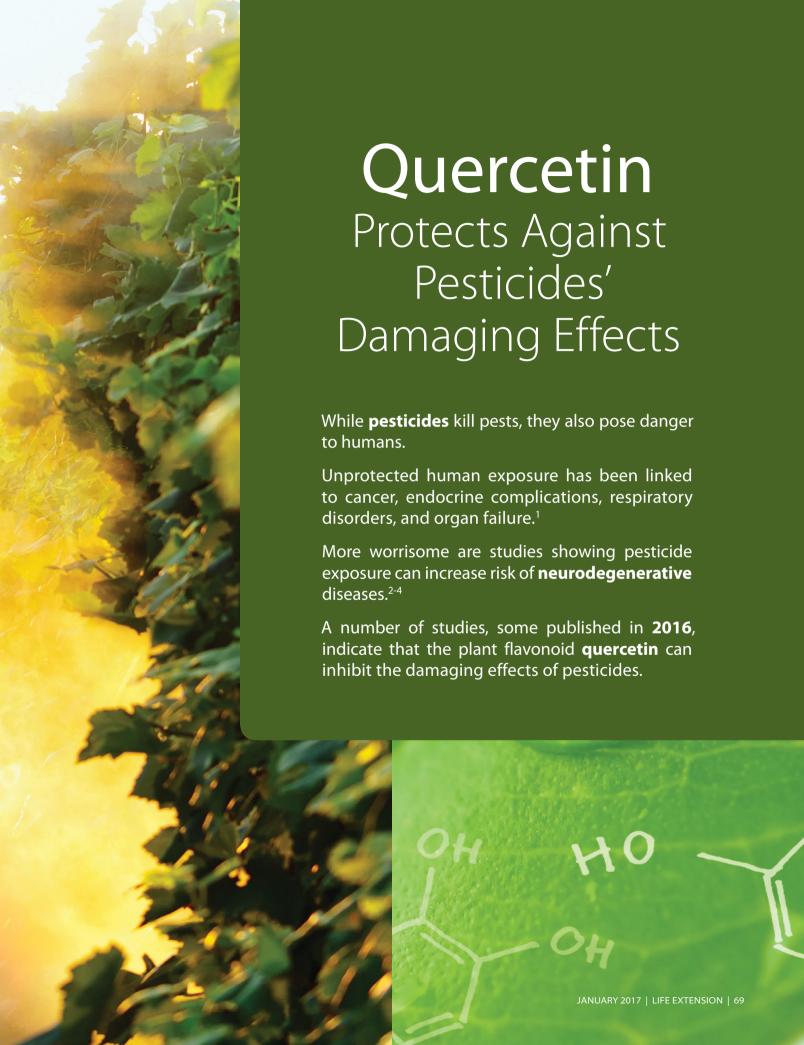
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* Alzheimers Dis. 2015;49(4):971-90.



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Pesticides are found in our food, air and water. Harmful chemicals are so abundant in our environment that we often absorb them without eating food.

With advances in manufacturing, many pesticides are no longer sprayed on crops. Instead, *systemic pesticides* are mixed in with fertilizer and absorbed by plants through their vascular system. This makes it *impossible* for consumers to wash off pesticide residue because it's within the fruits and vegetables that you eat.

While everyone is vulnerable to the dangers of pesticides, those especially at risk include children, pregnant women, sick persons, and the elderly. Pesticide exposure can result in conditions ranging from learning disabilities to Parkinson's disease and cancer.³⁻⁶ Because many different types of pesticides are used by farmers, exposure can overwhelm our efforts to stay healthy. Even everyday lawn chemicals pose risks and can remain in our body for decades.^{7,8}

Quercetin, a flavonol occurring in certain fruits and vegetables, 9-11 shows promise in animal studies to protect against some of the dangers of pesticides. 12,13 Until farmers completely stop their use, we are always at risk. But initial studies in the laboratory show that quercetin can provide some measure of cellular protection.

Quercetin Protects Against Pesticides

In various studies conducted on rats, **quercetin** was shown to provide substantial protection against a number of pesticides.

A June **2016** study in the journal *Human & Experimental Toxicology* found that quercetin had protective effects against toxicity induced by a mixture of *organophosphate* pesticides, which were originally developed as biological warfare agents.

The United States Environmental Protection Agency has indicated that organophosphate pesticides are dangerous not only to humans but also to bees and wildlife. ¹⁴ The study showed that quercetin can protect against organophosphate pesticide toxicity through the following mechanisms: ¹³

- Preserving energy, fatty acid, and sex hormone metabolism
- Inhibiting oxidative stress
- Protecting against DNA damage
- Preserving kidney and liver function

Also, although there is no specific treatment for poisoning by the pesticide *paraquat*, researchers are investigating quercetin's ability to minimize oxidative stress through its free-radical scavenging properties as a possible therapy.¹⁵

In a **2016** study, scientists demonstrated that quercetin protects cells against oxidative stress and cellular death caused by *dichlorvos*, another organophosphate pesticide. In lab cells exposed to this pesticide, pretreatment with quercetin significantly reduced dichlorvos-induced cell death, inhibited ROS (reactive oxygen species) generation, reduced levels of malondialdehyde (a marker of oxidation), and modulated the activities of two primary antioxidants (catalase and superoxide dismutase).¹⁶

In the same year, another rat study produced very similar findings. Various parameters representing the negative health effects of a mixture of pesticides were ameliorated when quercetin was simultaneously administered.¹⁷

As the population ages, researchers have increasingly focused on the potentially devastating brain effects of pesticides.





Pesticides Linked to Alzheimer's Disease

About 5.4 million Americans live with Alzheimer's disease,18 the sixth leading cause of death in the US.19 After age 65, the risk of Alzheimer's disease or vascular dementia doubles every five years.²⁰

Early-stage Alzheimer's disease patients are usually anxious, well aware that something is wrong. Shortterm memory is poor and patients have difficulty finding ordinary words. Crippling depression sets in with the awareness that something is being lost that will never be regained.

A great deal of Alzheimer's disease research has gone into finding the genes that increase susceptibility. However, multiple other factors are implicated, including chronic infections, declining hormone levels, inflammation, mitochondrial dysfunction, oxidative stress—and toxic chemicals.²¹⁻²⁶ Chief among this latter group are the over 1,000 chemicals that comprise pesticides, use of which has quintupled since 1945.27

Recent studies link chronic pesticide exposure to increased prevalence of dementia, including Alzheimer's disease.^{2,28} **Organophosphate** pesticides have been shown to lead to microtubule derangements and tau hyperphosphorylation—a hallmark of Alzheimer's disease. (Phosphorylation is the addition of a phosphate group to a molecule, which turns a protein enzyme on or off, altering its activity.) This mechanism of action suggests that, at the cellular and molecular level, these pesticides may at least partly account for the neurodegeneration of Alzheimer's disease.28

Ouercetin vs. Pesticides

- Although designed to kill pests, pesticide compounds can also trigger negative health effects in humans.
- Excessive or prolonged exposure to pesticides has been linked to cancer, endocrine complications, infertility, respiratory disorders, organ failure, birth defects, mood changes, and Alzheimer's disease.
- Prominent among the flavonoids produced by plants to protect themselves from destructive forces is the potent compound quercetin, which conveys similar protection to humans.
- Accumulating evidence demonstrates that quercetin protects against many of the biological effects of pesticides.

Scientists evaluating 86 Alzheimer's disease patients and 79 controls found that serum levels of dichloro*diphenyldichloroethylene* (DDE)—a metabolite of the pesticide dichlorodiphenyltrichloroethane (DDT) were **3.8-fold higher** in Alzheimer's disease patients.²⁹

Although banned in the US in 1972, people still come into contact with DDT through imported foods or by living near farmlands where DDT was formerly sprayed or near industrial sites where manufacturers dumped DDT-containing products.³⁰⁻³³

Similarly, exposure to the pesticide *beta-hexa-chlorocyclohexane* (beta-HCH) was found to be detectable in **76**% of Parkinson's disease patients, compared to **40**% of those <u>without</u> the disease. Based on serum levels of this pesticide, researchers could predict a Parkinson's diagnosis with a high degree of confidence.³⁴

Quercetin's Broad Brain-Protective Effects

Exposure to pesticides can result in a range of very subtle neurological symptoms that are not commonly recognized by the medical community, but that can be devastating to the individual, especially over time. These include loss of memory, poor coordination, reduced stimuli-response speed, decreased vision, altered or uncontrollable moods or other behavior, and impaired motor skills.³⁵

Rapidly accumulating evidence has identified quercetin as a potent neuroprotective nutrient.

Quercetin protects brain cells from excitotoxicity, the damage done by the repeated excitatory electrical impulses observed in Alzheimer's and other neurodegenerative diseases. ³⁶⁻³⁹ Its potent antioxidant mechanisms <u>reduce</u> toxicity of *beta amyloid* proteins that accumulate in the brain, eventually producing memory loss and dementia. ^{40,41} Quercetin has also been found to prevent brain-cell death in animal models of Parkinson's disease. ⁴²

The Alzheimer's Disease Risk in Your Home and How to Protect Yourself

When thinking of **pesticides**, most people focus on the over 1,000 chemicals used in agriculture today. But these 1,000 chemicals have been formulated into over 20,000 products—a fifth of which are designed for nonagricultural uses—in homes, gardens, schools, playgrounds, offices, golf courses, and hospitals. So microscopic traces of pesticides easily drift into homes. Studies show that detectable levels of these compounds can be found in many people, even newborns.²⁷

Just as they do in pests, pesticides can affect the human nervous system. These common pesticides include:⁵¹

- Organophosphates, a synaptic poison that damages the junction between two nerve cells,
- · Carbamates, a synaptic poison,
- Pyrethroids, an axonic poison that damages axons that conduct impulses away from the cell,
- · Avermectins, an axonic poison,
- · Imidacloprid, a synaptic poison, and
- Fipronil, an axonic poison.

Quercetin has been shown to inhibit pesticide damage by preserving energy, fatty acid, and sex hormone metabolism, inhibiting oxidative stress, protecting against DNA damage, and preserving kidney and liver function.¹³

In addition, research findings show that quercetin is associated with improved cardiovascular health, reduced cancer risk, milder allergic responses, and improved resistance to infection after extensive exercise.⁵² And, a recently published study demonstrated that quercetin enhanced memory recall in a mouse model of early-stage Alzheimer's disease—as well as in *human*, early-stage Alzheimer's disease patients.⁵³



Oxidative stress and chronic brain inflammation eventually produce changes that can lead to neurodegeneration.

The brain has a powerful free-radical defense system that upgrades cellular defenses such as **glutathione** and prevents brain-cell death. This natural defensive shield has been found to be activated by **quercetin.**⁴³ The quercetin flavonol also increases brain expression of protective **paraoxonase 2** (PON2), which scavenges free radicals that damage mitochondrial membranes and cause them to lose their electrical potential.^{44,45}

Oxidative stress and chronic brain inflammation eventually produce changes that can lead to neuro-degeneration.⁴⁶ Quercetin can preserve vital brain-cell function in the face of those changes, limiting the cell death that produces neurodegenerative diseases.⁴⁷

Not all free radical scavengers deliver the same defense as quercetin against the type of oxidative stress damage associated with Alzheimer's disease. In one study, isolated rat brain cells were exposed to either quercetin or vitamin C and then to hydrogen peroxide, which replicates the oxidative cell damage that occurs in Alzheimer's disease. Quercetin-treated brain cells showed significantly less damage to cellular membranes.⁴⁸

Quercetin's Potent Neuroprotection

Additional to stimulating cellular defenses against oxidative stress, quercetin activates sirtuins (SIRT1) and induces autophagy (removal of cellular waste debris)—both possible mechanisms for its neuroprotection.⁴⁹

This same review found that the neurotoxicity of several toxic agents, including *polychlorinated biphenyls*, *MPTP* (1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine)—and the insecticide **endosulfan**—was <u>decreased</u> by quercetin *in vivo*. Findings showed that "...quercetin ameliorates Alzheimer's disease pathology and related cognitive deficits in an aged...Alzheimer's disease mouse model."⁴⁹

Quercetin was shown to protect brain mitochondria against **endosulfan**. This pesticide normally induces oxidative stress in brain mitochondria by significantly lowering levels of catalase, superoxide dismutase (SOD), and glutathione. In rats, the pesticide resulted in swelling of mitochondria and higher levels of the oxidative stress marker malondialdehyde.⁵⁰

Pretreatment with **quercetin** was demonstrated to protect the brain mitochondria from oxidative stress, lipid peroxidation, and mitochondria swelling normally induced by **endosulfan**. The activities of the natural enzymes systems and the mitochondrial content of glutathione and malondialdehyde were all returned to healthy levels. The study author concluded:

"Thus, although endosulfan can have neurotoxic effects in brain[s of] rats, this toxicity can be prevented by quercetin." ⁵⁰

Summary

Pesticide compounds may increase the risk of diseases most commonly associated with aging, including **neurodegenerative** disorders.

Plants naturally produce molecules known as **flavonoids** to allow them to withstand a host of destructive forces, including chemical toxins.

New and accumulating evidence demonstrates that the plant flavonoid **quercetin** delivers protection—for the body and the mind—against many pathological effects associated with **pesticide** exposure.

The typical daily supplemental dose of quercetin is **150-400 mg**. It is often included in **resveratrol** formulas because of evidence showing that **quercetin** and **resveratrol** provide complementary health benefits when taken together. •

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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BY CHANCELLOR FALOON

PQQ Reduces Arthritis Inflammation

Pyrroloquinoline quinone, or PQQ, is a vitamin-like compound that has demonstrated impressive biological effects. It is found in tiny quantities in plant foods.¹

PQQ is responsible for creating <u>new</u> mitochondria as well as maintaining existing mitochondria within the cell.² PQQ's unique abilities have led many researchers to believe it can slow down the progression of aging.

Past studies have shown that PQQ can improve cardiovascular and brain health.³⁻⁸ More recent research has shown that PQQ can inhibit breakdown of healthy bone.

The latest findings indicate that PQQ has the potential to decelerate the deterioration of joints in **rheumatoid arthritis** and **osteoarthritis**.^{10,11}

Arthritis is a leading cause of disability in the US.¹² An urgent need exists to find treatments to prevent or delay its onset. Recent studies published in the journal *Inflammation* indicate PQQ may very well exert a protective effect in the **joints**.^{10,11}

Research Update

Rheumatoid Arthritis

Rheumatoid arthritis occurs when the body's immune system mistakenly attacks the cells in its joints. This leads to the release of inflammatory cytokines and enzymes that damage not just the cartilage, but also the bone. The disease does not just manifest in the joints. It produces a dangerous amount of inflammation that affects the rest of the body. Rheumatoid arthritis increases the risk of cardiovascular disease. Those with rheumatoid arthritis suffer an approximately 40% increased risk of overall mortality. 13,14

In response to the immune system's attack on joints, cells called **fibroblast-like synoviocytes**, which are found in the joints' synovial fluid, release inflammatory molecules. ¹⁵ Fibroblast-like synoviocytes can circumvent healthy cell turnover and instead release inflammatory molecules via a protein complex called NF-kappaB. ¹⁰

In one of the new studies from the journal *Inflammation*, researchers tested human fibroblast-like synoviocytes in vitro, or outside of the body. The scientists used an inflammatory agent to activate the release of cytokines from the fibroblast-like synoviocytes cells. In one of the groups of cells, POO was also added.10 The group of cells that did not receive POO had increased levels of proinflammatory cytokines. The cells that received the PQQ had a decreased production of the proinflammatory cytokines. In addition, PQQ was also able to halt the activation of NF-kappaB. The researchers also noted that POO may be able to attenuate certain enzymes (such as matrix metalloproteinases) that degrade a protein called type-II collagen present in our joints.



In a second part of this study, the researchers tested the effects of POO on two groups of mice with an animal model of rheumatoid arthritis. The scientists gave intraperitoneal injections of PQQ to one group of mice but not the other. After 45 days there was a dramatic difference with the POO-administered mice showing remarkable protection against inflammatory degeneration. The researchers observed narrowing of space between joints, increased inflammatory cell infiltration, and cartilage damage in the group that did not receive POO. Due to these impressive results, the researchers hypothesized that POO may be helpful in the treatment of other inflammatory conditions as well.

Osteoarthritis

Approximately **33**% of Americans 65 and older are affected by osteoarthritis, making it the most common form of arthritis. ¹⁶ It was believed that osteoarthritis was simply the result of age-related "wear and tear" of the joints, as well as the body failing to produce enough cartilage. ^{17,18}

Scientists now understand that the underlying causes of osteoarthritis are similar to that of rheumatoid arthritis.^{19,20}

A group of researchers conducted a study to test the effects of PQQ for osteoarthritis treatment.11 In this study, human chondrocytes, cells that produce and maintain cartilage, were tested in vitro. The researchers manipulated the environment of the cells to create inflammation and mimic the effects of osteoarthritis. One group of cells received PQQ before the researchers manipulated the environment while the other group did not. The researchers observed increased levels of collagen-degrading enzymes (matrix metalloproteinase) in the group that did not receive PQQ.

One of the pivotal biomarkers for inflammation is nitric oxide. Under normal physiological conditions nitric oxide acts as an anti-inflammatory as well as a vasodilator. But in certain circumstances, overproduction can lead to inflammation.²¹ In the group of cells that received PQQ, there was a significant reduction in joint-degrading enzymes and nitric oxide.

In addition, the researchers tested the effects of PQQ on mice who underwent a surgical procedure to induce osteoarthritis. Prior to treatment, one group received injections of PQQ with additional injections daily. The other group did not receive any. The group that received PQQ had significantly less severe cartilage damage compared to the group that did not receive PQQ.

Practical Approaches for Arthritis

The studies thus far are laboratory models and not directly relevant to human arthritis patients. PQQ's most impressive results have been seen in studies showing its support for mitochondrial function, cardiovascular health, and brain health.^{1,22,23}

For those seeking natural approaches to combat arthritis, there are natural anti-inflammatory compounds in widespread use today including: undenatured type II collagen, boswellia, and curcumin.

Derived from chicken cartilage, **undenatured type II collagen** has robust data showing reduction in pain and increase in function in dogs, horses, and humans.²⁴⁻²⁷ Results from one study showed, in just 90 days, a **40**% reduction on a pain scale for osteoarthritis patients supplementing daily with undenatured type II collagen.²⁷

Boswellia is an Indian plant that has demonstrated potent anti-inflammatory properties. It has been shown to inhibit the pro-inflammatory enzyme *5-lipo-oxygenase* or 5-LOX.^{28,29} This enzyme is responsible for facilitating the production of leukotriene, a pro-inflammatory compound that damages joints and cartilage.

Boswellia has also been shown to inhibit a type of *matrix metal-loproteinase* cartilage-degrading enzyme.³⁰ An impressive study on osteoarthritis patients showed that daily boswellia supplementation caused a **40.1%** reduction on a pain scale compared to placebo in just 30 days.³¹

Curcumin is the most active constituent of the turmeric root. A study tested the effects of curcumin or a nonsteroidal antiinflammatory drug (NSAID) on 45 humans during a flare-up of rheumatoid arthritis. The researchers tested the patients' blood for a systemic inflammatory marker called **C-reactive protein** and their pain score at the beginning of the study and then eight weeks afterwards. The results showed a 52% drop in C-reactive protein in the curcumin group. In the group receiving the NSAID, C-reactive protein increased by 1.5%. The NSAID group had a 42.1% improvement in pain scores, which was similar to the curcumin group, which had a 44.5% improvement (relief) in pain scores.³²

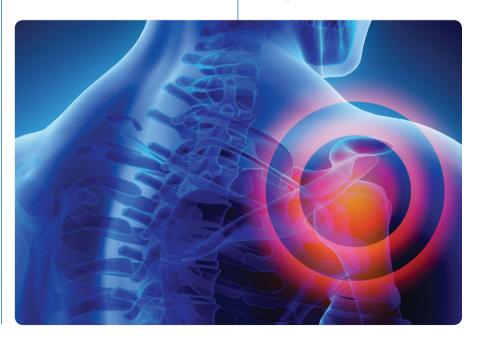
Summary

PQQ has been shown to promote mitochondrial biogenesis (creation of new mitochondria), which is essential for older cells to retain youthful energy output. PQQ also protects against mitochondrial damage.²

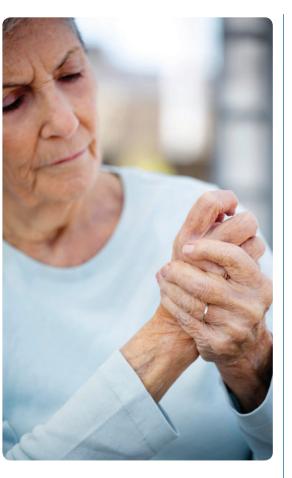
The two new studies described in this Research Update provide preliminary data to pave the way for human trials in arthritis patients. PQQ was effective at suppressing cartilage-degrading enzymes and inflammatory markers in human cells and in the laboratory mouse model.

These findings open a pathway for clinical studies to evaluate whether PQQ can benefit osteoarthritis and rheumatoid arthritis patients. Those seeking to relieve arthritis symptoms today, without resorting to drugs, have access to a variety of low-cost nutrients with clinically validated effects. •

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



Research Update



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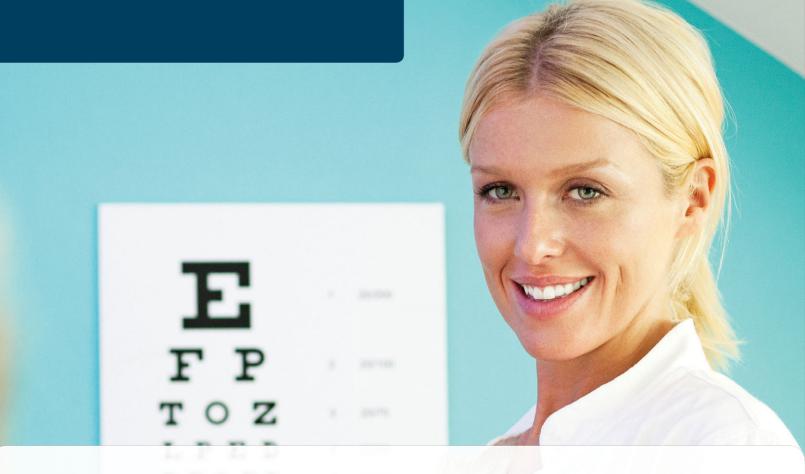
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Metformin and Glaucoma

Glaucoma is the second leading cause of blindness in the world.¹

A medical group submitted a report to *Life Extension Magazine*® that provides persuasive data that the **AMPK-activating** drug **metformin** may be of significant benefit in protecting the eyes against the threat of blindness from **open angle glaucoma**.

This report is written with some technical language that may make it challenging for some of our readers to understand.

We choose to publish it with the caveat that a succinct practical suggestion on how to use **metformin** to potentially <u>reduce</u> **glaucoma risk** be made in the introduction.

So here is what the medical group that authored this report recommends:

"Those with elevated intraocular pressure (IOP) and/or glaucoma should ask their doctor about prescribing a modest

250 mg-500 mg dose of metformin

twice a day after meals as it may have unique beneficial mechanisms in protecting against this blinding disorder."

We welcome you to read the report beginning on the next page that describes underlying pathologies of **open angle glaucoma** and how **metformin** can help to counteract them.



Aqueous Humor, Its Functions and Its Relation to Glaucoma Production

Glaucoma is a disease characterized by the increase of *intraocular pressure* due to various pathologies related to *aqueous humor* production, circulation, and drainage. In addition, the disease produces subsequent damage to the retina and atrophy of the **optic nerve** resulting in reduced visual acuity and ultimately leading to blindness.¹¹

Aqueous humor is a transparent, watery fluid that provides nutrition to the front part of the eye. It also transports the metabolic debris produced there to the bloodstream, thus maintaining transparency of the lens and cornea so light rays can pass through cleanly and provide clear vision. Most importantly, it keeps the cornea inflated with hydrostatic pressure, like water in a balloon.

There are many varieties of glaucoma, the most common being **open angle glaucoma**, in which the angle where the cornea and the iris meet is as wide and open as it should be, but the aqueous humor drainage channels become blocked over time and aqueous humor builds up. This raises intraocular pressure.^{12,13}

As pressure is exerted on the sensitive retina over time, it results in damage to nerve cells and their projection, the optic nerve. 11 Once the **optic nerve** is damaged, it can't be repaired, even if the raised *intraocular pressure* is corrected (figure 1). 14,15 Abnormally high pressure inside the eye usually causes this retinal and optic nerve damage.

Because open angle glaucoma occurs due to

the effects of aging, it may be that the disease is treatable with **metformin**, because of the drug's general antiaging properties.

How Metformin Functions in the Body

Metformin works to reduce blood sugar in several ways. It decreases the amount of glucose made by the liver, decreases the amount of sugar absorbed into the body, and makes insulin receptors more sensitive. Metformin does not increase insulin levels as many antidiabetic medications do, which makes it unlikely to cause dangerously low drops in blood sugar. ^{16,17} It's therefore considered safe for nondiabetics to take.

Let's examine how metformin works as an antiaging therapeutic agent and extrapolate the findings in terms of its ability to fight **glaucoma**.

AMPK activation helps to mimic the beneficial effects of calorie restriction.

Metformin enhances the activity of an enzyme found within all our cells called *adenosine monophosphate-activated protein kinase*, or AMPK for short. **AMPK activation** helps to mimic the beneficial effects of **calorie restriction** and exercise, the best documented method of slowing and reversing degenerative aging processes and biomarkers of human aging.¹⁸

The biological effects of increased **AMPK activity** include <u>inhibition</u> of **fat storage**, <u>reduced</u> **triglyceride synthesis**, and <u>increased</u> **glucose uptake** into muscle for metabolism.¹⁹⁻²⁷ AMPK activation also enhances destruction of diseased or dying cells as well as removal of intracellular metabolic debris – a method to slow and reverse degenerative aging processes of various organs.⁹

Further, experiments have shown that metformin, through **AMPK activation**, promotes the functional activity of the **sirtuin** family of **genes**, which is associated with **longevity**. Scientists have identified several signaling pathways involved in the regulation of aging processes that promote longevity. One of these signals, named **p53**, controls cell proliferation and is known as a **tumor-suppressor gene**. Loss of p53 predisposes normal cells to cancer. Metformin helps protect **functional p53** so cells are <u>less</u> likely to become **cancerous** ^{9,28-31}

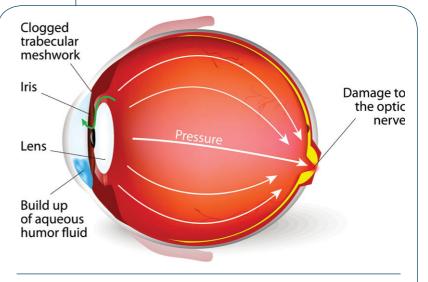


Figure 1 shows the production of aqueous humor by ciliary processes and obstruction to its drainage at trabecular meshwork exit channels. This results in raised intraocular pressure (arrows), pressing on the sensitive retina and optic nerve, resulting in glaucoma and ultimately loss of vision.



Metformin's Inflammation-Reducing Property

Nuclear factor-kappa B (NF-kB) is an internal cell signal that induces **chronic inflammation** responsible for many diseases, from cancer to heart attack, neuro-degenerative diseases and even glaucoma.³²⁻³⁵

NF-kB activation is blamed for many chronic diseases that ravage us as we age. Metformin produces higher **AMPK activity** which <u>decreases</u> expression of **NF-kB**.³⁶

By blocking NF-kB, metformin is thought to promote longevity by <u>inhibiting</u> systemic inflammatory processes in the body, which play havoc in all our vital organs including the brain and heart, as well as the eyes.

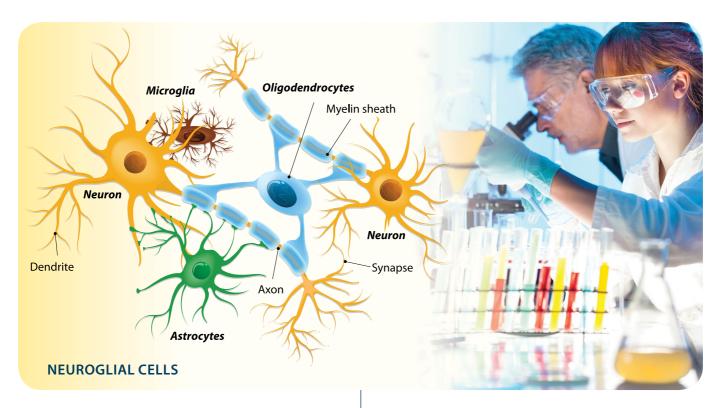
A recent study has found that metformin relieves neuropathic and other pain by decreasing the activation of microglial cells in the spinal cord that are an integral part of the central nervous system and its proper functioning as discussed below.⁶

Metformin's Effects on Nerve Cells (Neurons) and Their Physical Supporter-Glial Cells

Common characteristics for many neurodegenerative diseases include changes in glial cells, progressive neuronal loss, increased inflammation and oxidative stress.³⁷ Thus decreasing the activation of glial cells in the brain is one promising approach to reducing the inflammation in the brain responsible for various neurodegenerative diseases including Parkinson's and Alzheimer's disease. This is exactly what a group of researchers found in an animal model of neuropathic

The Benefits of Metformin

- Metformin has been prescribed for decades as an effective treatment against type II diabetes. But studies have shown metformin to have a number of other beneficial effects as well. These include promoting longevity, weight loss, and reduced cancer risk, as well as reducing chronic pain.²⁻⁷ The drug also has antiaging effects that mimic calorie restriction, and it favorably modulates genes thought to be involved in aging.^{8,9}
- Now, new research reveals metformin also reduces the development of open angle glaucoma, a progressive optic neuropathy and a leading cause of blindness.
- A University of Michigan study has found metformin to be linked with a 25% reduction in the risk of developing open angle glaucoma. Other medications used to treat type II diabetes did not have a similar benefit. Metformin is the <u>only</u> drug that has an intraocular pressure-reduction therapeutic effect.
- Everyone over age 50 would be welladvised to get tested for glaucoma and to ask their physician about possibly taking metformin, which could be preferable to typical antiglaucoma drugs, considering their common side-effects and lack of antiaging properties.



pain treated with metformin—glial cell activation was decreased and chronic pain was reduced.⁶ We believe these findings may have implications to decrease neuropathic pain in thousands of patients treated with the expensive drug gabapentin (Neurontin[®]).

Glial cells have multiple functions:^{38,39}

- 1. They surround neurons and hold them in place.
- 2. They supply nutrients and oxygen to neurons.
- 3. They insulate one neuron from another.
- 4. They destroy pathogens and remove dead neurons with various processes, including production of inflammatory cytokines that, besides attacking the invading microorganism, can promote neurodegenerative diseases.
- 5. They transport the brain's interstitial fluids between neurons to the cerebrospinal fluid, through drainage channels they create called glymphatics.
- 6. They play a role in synapse formation and the transmission of electrical signals from one nerve cell to another, especially in memory centers of the brain. Basically, glial cells are caregivers to nerve cells and facilitate nerve cell activity in the central nervous system and maintain a homeostatic milieu for nerve cells to function properly.

Now let us examine how metformin can help fight glaucoma and age-related neurodegenerative diseases related to glial cell pathology.

Originally marketed as an agent for type II diabetes. metformin has been found to have a number of other uses in clinical practice, including, in one study, the ability to decrease the activation of glial cells in the spinal cord.⁶ Researchers reported complete resolution of suffering in some rats with induced neuropathic pain. This study reveals the impact of metformin on the nervous system glial cells, which are believed to be associated with chronic pain. If that is the case, is it possible that metformin can protect other parts of the nervous system, such as the retinal ganglion cells, by inhibiting the activity of glial cells that produce inflammatory cytokines that are toxic to neurons? This would also explain our finding and the findings of other scientists that those on metformin have better cognition with reduction in dementia. 40,41 In our practice we routinely prescribe metformin for people over the age of 50 to be taken twice daily after meals to prevent future neurodegenerative diseases and aging.

How Does Metformin Reduce Glaucoma?

A recent study found that metformin reduces the intraocular pressure of primary open angle glaucoma. ¹⁰ Open angle glaucoma is a progressive optic neuropathy characterized by loss of retinal ganglion cells and optic nerve atrophy. ⁴² It's the most common

form of glaucoma and is often asymptomatic and may even go undetected for a while.⁴³ By the time vision is noticeably impaired, the loss is irreversible, because once the nerve cells are dead in the retina with degeneration of the connected nerve fibers, nothing can restore them.

Open angle glaucoma is a manifestation of aging along with other neurodegenerative diseases. Normally, through autophagy, our cells purge themselves of accumulated debris, often called "cellular metabolic junk." Autophagy is a natural mechanism that disassembles cells' unnecessary or dysfunctional components as they age and lose their function. But over time, our cells lose this housekeeping ability. 44,45 Metformin has been shown to promote this process.⁴⁶

According to a study at the University of Michigan, metformin was associated with a 25% reduction in the risk of developing **open angle glaucoma**. They also found that other oral antidiabetic medications used to treat type II diabetes did not confer a similar risk reduction. **Metformin** is the only drug endowed with this **intraocular pressure-reduction** therapeutic

This retrospective cohort study was based on longitudinal data from more than 150,000 patients with type II diabetes and no preexisting record of open angle glaucoma. Forty percent filled at least one met-

Autophagy: Cellular House Cleaning

Christian de Duve, 1974 Nobel Laureate in physiology or medicine, coined the term autophagy (meaning "self-eating") in 1963. This year, biologist Yoshinori Ohsumi, of the Tokyo Institute of Technology, has been awarded the Nobel Prize in physiology or medicine for his discoveries in autophagy, the process whereby a cell recycles part of its own cellular debris (cellular house cleaning).

Scientists had been aware of autophagy for decades, but knew little about how it worked until Ohsumi's pioneering experiments in the 1990s. It's important because autophagy can eliminate invading intracellular bacteria. Disrupted autophagy has been linked to Parkinson's and Alzheimer's disease, type II diabetes and other disorders that particularly affect the elderly.

We know that metformin enhances autophagy, which is how it reduces diseases of aging such as Parkinson's and Alzheimer's, and may reduce the incidence of glaucoma. This is one more reason to prescribe metformin.

formin prescription. During the 10-year study period, 5,893 (3.9%) of the patients of a large health care network developed the disease. The researchers compared users of metformin with nonusers, analyzing the data by means of regression modeling. Each model demonstrated substantial reductions in open angle glaucoma risk among those using metformin. In two years, a diabetic patient taking a daily **2,000 mg** dose

Metformin was associated with a **25%** reduction in the risk of developing open angle glaucoma.

of metformin would have a 20.8% reduction in open angle glaucoma risk, compared with a diabetic patient who had no metformin exposure. 10

Glaucoma is the second leading causes of blindness in the world.1 It is estimated that in excess of 2.5 million people have glaucoma in the United States, and that more than 120,000 people are legally blind from the disease.⁴² Many people who have it aren't aware of it. Blindness from glaucoma is six to eight times more common in African Americans than Caucasians, and, after cataracts, is the leading cause of blindness among them.⁴³ We advise all those over the age of 50, especially African Americans and women, to get their eyes tested for glaucoma and ask their physician if it is appropriate to start taking metformin, not as an antidiabetic, but for its antiaging, antiglaucoma properties. Women have longer life expectancy and are more likely than men to develop age-related eye diseases like glaucoma.47



There are dozens of drugs available to treat glaucoma.^{48,49} Many of them have systemic complications and lack the antiaging effect of metformin on the rest of the body. Wouldn't it make sense to prescribe metformin to prevent glaucoma and at the same time delay and/or reverse the rayages of aging?

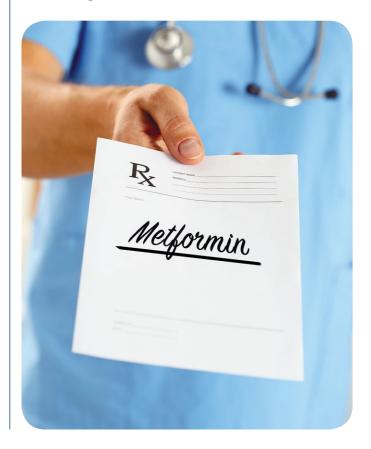
Researchers at the University of Michigan Kellogg Eye Center have suggested a clinical trial protocol in which newly diagnosed glaucoma patients would be randomized to receive either an IOP (intraocular pressure)-lowering drug plus metformin or a glaucoma drug plus a placebo. ⁵⁰ From our point of view, a randomized clinical trial which might take decades may not be needed. Metformin is inexpensive, widely used to treat type II diabetics, and has hardly any adverse effects.

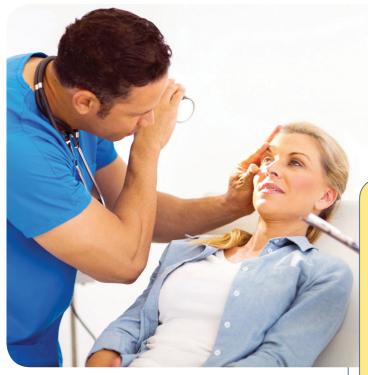
Summary of How Metformin Wards Off Glaucoma

Given how aqueous humor is formed, how it circulates and exits the eye, there are numerous possible explanations for how metformin works. It may act to reduce open angle glaucoma risk at multiple levels, which need to be further examined. The possible mechanisms are:

- 1. Metformin, by inhibiting an inflammatory reaction and its related cytokines, may reduce aqueous humor production by ciliary processes, and bring it to stability.
- 2. By promoting autophagy, it may prevent exfoliated cells from blocking the aqueous humor drainage channels of the meshwork and the Schlemm's canal.
- 3. By AMPK activation, it may reverse the biomarkers of human aging in the uveal aqueous humor production structures and transportation channels of aqueous humor.
- 4. Due to increased AMPK activation, as the aqueous humor circulates, it comes in contact with trabecular meshwork and may cleanse the glycation around the endothelial cells of the trabecular meshwork, thus allowing the aqueous humor to pass to exits without resistance.
- 5. Metformin in the aqueous humor may cleanse and open the pores in the Schlemm's canal and uvea-scleral pathways by activation of AMPK, resulting in autophagy within the disease-afflicted lining cells of trabecular meshwork.

- 6. By autophagy, it may effectively cleanse the platelet clumps and lipid deposits in the trabecular meshwork and the Schlemm's canal that facilitates the easy drainage of the aqueous humor without increasing intraocular pressure.
- 7. Metformin protects the functional p53 gene while repressing and/or blocking the proinflammatory NF-kB by reversing or inhibiting inflammatory process in the body, including the eyes. ^{29,30,36} By reduction of inflammatory cytokines, it may protect the retina and prevent the degeneration of ganglion cells and optic nerve fibers, thus reducing the chances of blindness.
- 8. Metformin reduces resistance to insulin, thus helping uptake and metabolism of circulating sugar, and preventing the adverse effects of hyperglycemia such as glycation—the bonding of a protein or lipid molecule with a sugar molecule. 16,17,51
- 9. Loss of ganglion cells in the retina is a leading cause of blindness in open angle glaucoma.⁴² This could be prevented with metformin by decreasing activation of glial cells in the retina and optic nerve.





FDA Approves Human Trials on Metformin Antiaging Effects

Further studies will point out the multiple ways metformin reduces the incidence of open angle glaucoma in older people as it provides antiaging protection in other organs and tissues and possibly even prevents or reduces the incidence of age-related macular degeneration.

Interestingly, the FDA's approval of the first human trials to see if metformin can protect against diseases of aging was headlined in news media reports. We hope this study includes the drug's effect on the eyes of the aging population.

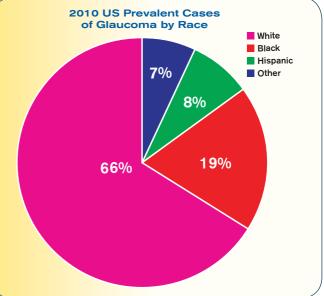
For decades, Life Extension has discussed the antiaging effects of metformin. Finally, the FDA has heard their call. This study may take decades to reveal its findings, hence our practice has started advocating for metformin use for people over the age of 50 to promote good health and reverse, inhibit, or stop the ravages of aging.

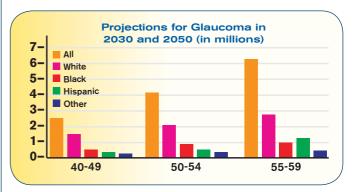
Although it can cause lactic acidosis if taken in doses that are much larger than required for treatment, metformin is essentially very safe. The public should demand that the FDA approve metformin for use without prescription as an over-the-counter medication, both in oral form and as ophthalmic drops.

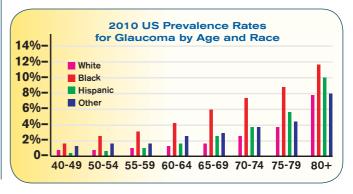
This will reduce medical cost and improve the health of many, with reduction in age-related diseases (which cost billions to care for). It will also bestow longevity, with probable reductions of neurodegenerative



Glaucoma Statistics and Data⁴⁷







FELOPZD

diseases such as Parkinson's and Alzheimer's, and at the same time provide good eyesight. Until that happens, the best alternative is for patients to ask their physicians to prescribe metformin for them and put it in writing that they, the patients, will not hold their doctors responsible for any untoward effects. Those with elevated intraocular pressure and/or glaucoma should ask their doctor about prescribing a modest 250 mg-500 mg dose of metformin twice a day after meals.

When developed, metformin ophthalmic drops, besides preventing open angle glaucoma, may also prevent or delay the development of age-related macular degeneration and diabetic retinopathy, and restore good vision to the aging population inexpensively. We appeal to the pharmaceutical industry to develop metformin ophthalmic drops with other adjuvant therapeutic agents to treat various eye diseases such as open angle glaucoma, age-related macular degeneration, retinitis pigmentosa, diabetic retinopathy and uveitis. •

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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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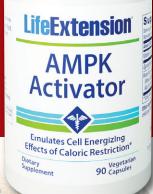
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Recipes for a Modern Vegetarian Lifestyle

BY GARRY MESSICK

Solla Eiríksdóttir is a celebrity chef in her native Iceland, famed for her TV shows and four restaurants which cater to fans of vegan and vegetarian dishes and raw food. She has published five cookbooks in Iceland, but *Raw: Recipes for a Modern Vegetarian Lifestyle* is her first cookbook in English.

Solla, 55, collaborated with her 36-year-old daughter, Hildur, on the new book, and the duo had a specific aim in mind. "What we're doing is transforming fruit and vegetables into real dishes instead of just making salad all the time," explains Solla. "We love the freshest raw materials so your taste buds are screaming for more."

Solla began investigating raw foods just after Hildur was born. She was suffering with a number of allergies at the time, and her doctor wanted to treat her ailments with drugs, but that would have required her to stop breast-feeding her infant daughter. She refused, and instead went to a nutritionist who put her on a vegan diet. Amazingly, Solla's allergies vanished within six months. From there, she got a job in a vegetarian restaurant, where she began to develop her cooking skills. She subsequently went on to study at the Living Light Culinary Institute in Fort Bragg, California.

Why raw? In the introduction to her new cookbook, the Icelandic chef explains that while most vegetables and fruits are rich in fiber, minerals, vitamins, antioxidants, phytonutrients, and enzymes, "Some of these compounds are sensitive to heat and a significant amount can be lost in the process of cooking." Enzymes, in particular, which catalyze digestion, denature with high heat, leading proponents of raw food to usually restrict application of heat in their cooking.

Raw features 75 recipes, all equally healthy and delicious. Solla and Hildur helpfully include symbols that denote when a particular dish is gluten-free, dairy-free, nut-free, raw or vegan. The following are four sample recipes from the book. Bon appetit!

Healthy Eating

Chia and Millet Flake Porridge

Serves 2

1 cup almond milk

3 tablespoons chia seeds

1 cup rolled millet flakes

1 teaspoon vanilla powder

1 teaspoon ground cinnamon

1/2 teaspoon lemon juice

A pinch of salt

1/2 banana, thinly sliced for topping

For the raspberry compote:

1 pear, peeled, cored, and chopped into small pieces or grated

1 cup raspberries (fresh or frozen)

1 tablespoon shredded fresh ginger root



Put the almond milk and chia seeds in a clean glass jar, put the lid on, and shake for 2-3 minutes, or until combined. Stir in the millet flakes, vanilla, cinnamon, lemon juice, and salt, then put the lid back on and set aside to rest for 15-30 minutes, or overnight.

For the compote, put the pear, raspberries, and ginger into a medium bowl and mash with a fork until it is the consistency you like. We like it slightly chunky. Alternatively, place the ingredients into a food processor and process using the pulse button.

When ready to serve, pour half the raspberry compote into a bowl or a glass jar, add the chia porridge, and top with a layer of thinly sliced banana. Spoon the remaining raspberry compote on top and eat.

Tofu Scramble with Kale and Avocado

Serves 2-3

2 tablespoons olive oil
½ onion, finely chopped
2 cloves of garlic, chopped
4 kale leaves, stems (stalks) removed

For the tofu marinade:

2-3 tablespoons almond milk
2 tablespoons nutritional yeast flakes
1 tablespoon tamari
1 tablespoon mustard
½ teaspoon ground turmeric
¼ teaspoon red pepper (chili) flakes
A pinch of salt

To garnish:

1 cup tofu

1/2 avocado sliced Sprouts of your choice (optional) Start by marinating the tofu. Stir the almond milk, nutritional yeast flakes, tamari, mustard, turmeric, red pepper (chili) flakes, and salt together in a bowl. Before adding the tofu, squeeze out all the liquid. A good way to do this is to wrap the tofu in a clean dish cloth and squeeze it gently so the water comes out through the cloth. Be gentle so the tofu doesn't become

a paste. You may need to use 2 cloths because a lot of liquid is likely to come out. When all the water has been squeezed out, crumble the tofu into a bowl with all the remaining marinade ingredients, and mix to combine.

Heat the olive oil in a saucepan over medium heat. Add the onion and garlic and cook for 3-4 minutes, or until golden brown. Add the kale leaves, stir for 1 minute, then add the tofu and cook for another 4-5 minutes. Serve in a bowl and garnish with sliced avocado and sprouts, if using.



Rainbow Pasta with Pesto

Serves 3-4

1 rutabaga (swede)

1-2 carrots

1 beet (beetroot)

1 small zucchini (courgette)

2 tablespoons lemon juice

1 tablespoon olive oil

For the green pesto:

1/2 cup cashew nuts

1 handful of basil

2-3 kale leaves, stems (stalks) removed

1-2 tablespoons nutritional yeast flakes

1 large clove garlic

1/4-1/2 teaspoon sea salt flakes

1/4-1/2 teaspoon cold-pressed olive oil

For the pesto, put the cashew nuts in a bowl, pour in enough water to cover, and soak for about 2 hours. Drain and discard the soaking water.

Put the cashew nuts into a food processor with the remaining ingredients, except the olive oil, and blend. The texture of the pesto should be chunky. Transfer the pesto to a bowl and add the olive oil. Stir gently to mix together. Spoon into a clean glass jar and set aside.

Peel the rutabaga (swede), carrots, and beet (beetroot) and use a julienne peeler or spiralizer to shred the vegetables into spaghetti-like strips.

Put your vegetable spaghetti into a bowl and add the lemon juice and olive oil. Stir together, then cover with plastic wrap (Clingfilm) and leave for 15-25 minutes to let the "spaghetti" soften. Serve with green pesto.



Healthy Eating

Ouinoa Pizza Crust

Makes 1 pizza

 3/4 cup quinoa
 1/2 teaspoon sea salt flakes
 1 clove garlic
 1/2 teaspoon freshly ground black pepper
 2 teaspoons dried oregano
 1/4 cup grated vegan cheese
 1 tablespoon olive oil

For topping:

Scant ½ cup vegan cream cheese ½ zucchini (courgette), very thinly sliced
2-3 tablespoons pine nuts
3-4 sprigs rosemary
1 tablespoon truffle oil

Put the quinoa into a bowl, pour in enough water to cover, and let soak overnight.

The next day, preheat the oven to 375 degrees and line a baking sheet with parchment (baking) paper.

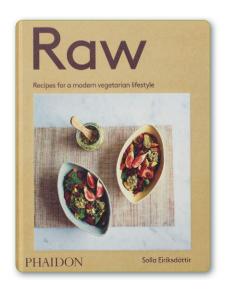
Drain and rinse the quinoa, then put it into a blender together with ¼ cup water, the salt, garlic, black pepper, and oregano and blend until smooth. Pour the batter into a bowl and mix the cheese and olive oil.



Put a 9 inch tart ring on the prepared baking sheet and pour in the quinoa batter. Bake for 20 minutes, then remove from the oven. Wearing oven mitts, flip the crust over by covering it with another baking sheet, grasping both sides of the 2 baking sheets, and flipping the sheets with the crust between them. Bake on the second sheet for another 5-10 minutes.

Remove the crust from the oven and lower the temperature to 345 degrees. Spread the crust with the cream cheese, top with the zucchini (courgette) slices, and sprinkle with the pine nuts. Bake for another 8 minutes.

Meanwhile, in a small skillet, briefly cook the rosemary sprigs in the truffle oil over medium heat. When the pizza is ready, sprinkle with the fried rosemary and serve.



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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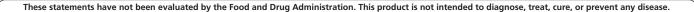
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References

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Multi Stem Cell Skin Tightening Complex Neck Rejuvenating Anti-Oxidant Cream Pigment Correcting Cream Rejuvenating Serum Rejuvenex® Body Lotion RejuveneX® Factor Firming Serum Renewing Eye Cream Resveratrol Anti-Oxidant Serum

Skin Lightening Serum Skin Restoring Phytoceramides with Lipowheat® Skin Stem Cell Serum Stem Cell Cream with Alpine Rose Tightening & Firming Neck Cream Triple-Action Vitamin C Cream Ultimate MicroDermabrasion Ultra Eyelash Booster

Ultra Lip Plumper Ultra Rejuvenex® Ultra RejuveNight® Ultra Wrinkle Relaxer Under Eye Refining Serum Under Eye Rescue Cream Vitamin C Serum

Shade Factor

Vitamin D Lotion Vitamin E-ssential Cream Youth Serum

Sleep

Bioactive Milk Peptides Enhanced Natural Sleep® with Melatonin Enhanced Natural Sleep® without Melatonin Fast-Acting Liquid Melatonin Glycine L-Tryptophan Melatonin Optimized Tryptophan Plus

Sports Performance

Creatine Capsules Creatine Whey Glutamine Powder (Vanilla Flavor) New Zealand Whey Protein Concentrate (Natural Chocolate and Vanilla Flavor) Tart Cherry with CherryPure® Whey Protein Isolate (Chocolate and Vanilla Flavor)

Vitamins Ascorbyl Palmitate Benfotiamine with Thiamine Beta-Carotene BioActive Complete B-Complex Buffered Vitamin C Powder Fast-C® with Dihydroquercetin Gamma E Mixed Tocopherol Enhanced with Sesame Lignans Gamma E Mixed Tocopherol/Tocotrienols High Potency Optimized Folate Inositol Caps Liquid Emulsified Vitamin D3 Liquid Vitamin D3 Low-Dose Vitamin K2 Methylcobalamin MK-7 Natural Vitamin E No Flush Niacin Optimized Folate (L-Methylfolate)
Pantothenic Acid (Vitamin B-5) Pyridoxal 5'-Phosphate Caps Super Absorbable Tocotrienols Super Ascorbate C Capsules Super Ascorbate C Powder Super K with Advanced K2 Complex Vitamin B12 Vitamin B6 Vitamin C with Dihydroquercetin Vitamin D3 with Sea-Iodine™ Vitamin D3 Vitamins D and K with Sea-Iodine™

Weight Management

7-Keto® DHEA Metabolite Advanced Anti-Adipocyte Formula Advanced Natural Appetite Suppress CalReduce Selective Fat Binder DHEA Complete Garcinia HCA HCActive™ Garnicia Cambogia Extract Integra-Lean® Mediterranean Trim with Sinetrol™-XPur Optimized Irvingia with Phase 3™ Calorie Control Complex Optimized Saffron with Satiereal® Super Ctrimax®
Super CLA Blend with Guarana and Sesame Lignans Super CLA Blend with Sesame Lignans Waist-Line Control™

Women's Health

Advanced Natural Sex for Women® 50+ Breast Health Formula Femmenessence MacaPause® Natural Estrogen Progesta-Care® Super-Absorbable Soy Isoflavones Ultra Soy Extract

SUPER SALE SAVINGS ON ALL PRODUCTS

			YO	UR PRIC	E				YO	UR PRIC	E
ITEM N	o. PRODUCT	Retail Each	1 Unit	4 Unit	10 Unit	ITEM I	No. PRODUCT	Retail Each	1 Unit	4 Unit	10 Unit
	A	\$	Each	Each	Each QTY Total	70000	BLOOD PRESSURE MONITOR (ACCUFIT™) • med/lg cuff	\$ 79.99	Each	Each	Each QTY To
01524	ACETYL-L-CARNITINE • 500 mg, 100 veg. caps	3/1 00	25.50	22.50			BLOOD PRESSURE MONITOR • Digital wrist cuff	69.95			
01324	σ, σ ,			35.00	_		BLOOD PRESSURE (Triple Action AM/PM) • 60 veg. tabs			28.00	
01628	ů ,			16.50			. , ,				
							BLUEBERRY EXTRACT • 60 veg. caps			15.00	
01630	ADVANCED LINID CONTROL + 120 veg. caps			31.50			BLUEBERRY EXTRACT W/ POMEGRANATE • 60 veg. caps			20.25	
01828			22.50	20.25	_		BONE FORMULA (DR. STRUM'S INTENSIVE) • 300 caps			37.50	
00681	3, 111, 11		44.99		-		BONE RESTORE • 120 caps			14.25	
29727	AHCC® (KINOKO® GOLD) • 500 mg, 60 veg. caps		52.47	0400			BONE RESTORE W/VITAMIN K2 • 120 caps	24.00		16.50	
00457			27.75				BONE STRENGTH FORMULA W/KOACT® • 120 caps			30.00	
	AMPK ACTIVATOR • 90 veg. caps			33.00		00313	BONE-UP® • 240 caps	28.95		20.41	
01509	ANTI-ADIPOCYTE FORMULA W/MERATRIM® & INTEGRA LEAN® (Advanced) • 60 veg. caps	39.00	29.25	27.00			BORON • 3 mg, 100 veg. caps	5.95		3.94	
02140	ANTI-ALCOHOL w/HEPATOPRO COMPLEX • 60 caps	22.00	16.50	15.00			BOSWELLA • 100 caps			22.50	
01625	APPLEWISE POLYPHENOL EXTRACT	21.00	15.75	14.25			BRAIN SHIELD® GASTRODIN • 300 mg, 60 veg. caps			22.50	
	600 mg, 30 veg. caps				_	01253	BRANCHED CHAIN AMINO ACIDS • 90 caps			12.75	_
01039	ARGININE/ORNITHINE • 500/250, 100 caps	17.99	13.49			01942	BREAST HEALTH FORMULA • 60 caps	34.00	25.50	22.50	_
00038	ARGININE/ORNITHINE POWDER • 150 grams	22.95	17.21	14.25		00893	BRITE EYES III • 2 vials, 5 ml each	34.00	25.50	24.00	
01624	(L)-ARGININE CAPS • 700 mg, 200 veg. caps	26.50	19.88	17.44		26576	BROCCO MAX® • 60 veg. caps	26.95	20.21		_
02004	ARTERIAL PROTECT • 30 veg. caps	48.00	36.00	33.00		01203	BROMELAIN (Specially-coated) 500 mg, 60 enteric coated tablets	21.00	15.75	14.25	
01617		44.00	33.00	30.00			C				
01610	120 veg. caps	26.00	07.00	04.00		01653	CALCIUM CITRATE W/VITAMIN D • 300 caps	24.00	18.00	15.94	
01618	ARTHROMAX® ADVANCED W/UC-II® & APRÉSFLEX® 60 caps	30.00	27.00	24.00			CALCIUM D-GLUCARATE • 200 mg, 60 veg. caps			11.25	
02108	ARTHROMAX® HERBAL JOINT FORMULA • 60 veg. caps	40.00	30.00	27.00		†01823	CALREDUCE SELECTIVE FAT BINDER			28.50	
01404	ARTHRO-IMMUNE JOINT SUPPORT • 60 veg. caps	32.00	24.00	21.00			120 mint chewable tablets				
00919	ARTICHOKE LEAF EXTRACT • 500 mg, 180 veg. caps	30.00	22.50	21.00		01700	CARDIO PEAK™ w/STANDARDIZED HAWTHORN & ARJUNA 120 veg. caps	36.00	27.00	24.00	
01533	ASCORBYL PALMITATE • 500 mg, 100 veg. caps	22.50	16.88	15.00		00016	CARNITINE W/GLYCOCARN® (Optimized) • 60 veg. caps	26.00	27.00	24.00	
00888	ASHWAGANDHA EXTRACT (Optimized) • 60 veg. caps	10.00	7.50	6.75			. (1)		11.25	24.00	
01805	ASIAN ENERGY BOOST • 90 veg. caps	24.00	18.00	16.50			L-CARNITINE • 500 mg, 30 veg. caps			9.90	
01066	ASPIRIN • 81 mg, 300 enteric coated tablets	6.00	4.50	4.00			CARNOSINE • 500 mg, 60 veg. caps			24.00	
01923	ASTAXANTHIN WITH PHOSPHOLIPIDS • 4 mg, 30 softgels	16.00	12.00	10.50			CARNOSINE (Super) • 500 mg, 90 veg. caps		46.50 10.50	42.00	
	В						CAT MIX • 100 grams powder			8.25	
00920	BENFOTIAMINE W/ THIAMINE • 100 mg, 120 veg. caps	19.95	14.96	13.95		01899	CHILDREN'S FORMULA LIFE EXTENSION MIX™ 100 chewable tablets	20.00	15.00	13.50	
00925	BENFOTIAMINE (Mega) • 250 mg, 120 veg. caps	30.00	22.50	20.25		00550	CHLORELLA • 500 mg, 200 tablets	23.98	17.99		
01206	BERRY COMPLETE • 30 veg. caps	21.00	15.75	14.00		01571	CHLOROPHYLLIN • 100 mg, 100 veg. caps	24.00	18.00	15.00	
01496	BERRY COMPLETE W/ACAI (Enhanced) • 60 veg. caps	29.00	21.75	19.50		01359	CHO-LESS™ • 90 capsules	35.00	26.25		
00664	BETA-CAROTENE • 25,000 IU, 100 softgels	11.50	8.63			01910	CHOL-SUPPORT™ • 60 liquid veg. caps	48.00	36.00	32.00	
01622	BIFIDO GI BALANCE • 60 veg. caps	20.00	15.00	13.50		01504	CHROMIUM W/CROMINEX® 3+ (Optimized)	9.00	6.75	6.00	
01873	BILBERRY EXTRACT • 100 mg, 90 veg. caps	36.00	27.00	24.00			500 mcg, 60 veg. caps				
01512	BIOACTIVE MILK PEPTIDES • 30 caps	18.00	13.50	12.00		01503	CINSULIN® W/INSEA2® AND CROMINEX® 3+ • 90 veg. caps	38.00	28.50	25.50	
01631	BIO-COLLAGEN W/PATENTED UC-II® • 40 mg, 60 small caps	36.00	27.00	24.00		01906	CISTANCHE (Standardized) • 30 veg. caps	20.00	15.00	12.00	
*01006	BIOSIL™ • 5 mg, 30 veg. caps		15.99			01818	CITRIMAX® (Super) • 180 veg. caps	40.00	30.00	28.50	
	BIOSIL™ • 1 fl oz		25.59			00818	CLA BLEND W/SESAME LIGNANS (Super) 120 softgels	36.00	27.00	24.75	19.75
	BIOTIN • 600 mcg, 100 caps		5.63	4.88		00210	CLA BLEND W/GUARANA & SESAME LIGNANS (Super)	42.00	31.50	28 75	
01709			12.00	10.50		00019	120 softgels	72.00	01.00	20.70	
	BLACK CUMIN SEED OIL W/BIO-CURCUMIN® • 60 softgels		24.00			01896	COGNITEX® W/BRAIN SHIELD® • 90 softgels	60.00	45.00	39.00	36.00
	BLAST™ • 600 grams of powder	26.95				01897	COGNITEX® W/PREGNENOLONE & BRAIN SHIELD®	62.00	46.50	39.75	37.50
02025			33.00	28.00		01.404	90 softgels	20.00	00.50	00.05	04.00
	, , ,					01421	COGNITEX® BASICS • 60 softgels	38.00	28.50	26.25	24.00
	SUBTOTAL OF COLUMN 1						SUBTOTAL OF COLUMN 2				

OFFER ENDS JANUARY 31, 2017

ITC:	J. PRODUCT	D : "		OUR PRI			4 -
ITEM I	No. PRODUCT	Retail Each \$	1 Unit Each	4 Unit Each	10 Unit	ITEN QTY Total	1 N
1659	COGNIZIN® CDP CHOLINE CAPS • 250 mg, 60 veg. caps		27.00	25.50	Lauii	8016	33
1945	COMPLETE B-COMPLEX (BioActive) • 60 veg. caps	12.00	9.00	8.00		8012	23
2098	COMPREHENSIVE NUTRIENT PACKS ADVANCED • 30 packs	90.00	67.50	61.50		8010)7
1949	COQ10 w/d-LIMONENE (Super-Absorbable) 50 mg, 60 softgels	25.00	18.75	16.50	15.00	8013	
1948	COQ10 w/d-LIMONENE (Super-Absorbable) 100 mg, 100 softgels	46.00	34.50	28.00	26.25	8013	
1929	COQ10 (Super Ubiquinol) • 100 mg, 60 softgels	56.00	42.00	36.00	33.00	8010)2
1733	COQ10 w/BIOPQQ® (Super Ubiquinol) • 100 mg, 30 softgels	54.00	40.50	33.00	30.00	8010)9
1426	COQ10 w/ENH MITOCHONDRIAL SUPPORT™ (Super Ubiquinol) • 100 mg, 60 softgels	62.00	46.50	39.00	36.00	8011 8013	
1425	COQ10 w/ENH MITOCHONDRIAL SUPPORT™ (Super Ubiquinol) • 50 mg, 100 softgels	58.00	43.50	34.50	31.50	8010	
1427	COQ10 w/ENH MITOCHONDRIAL SUPPORT™ (Super Ubiquinol) • 50 mg, 30 softgels	20.00	15.00	12.00		8014 8013	
)1431	COQ10 w/ENH MITOCHONDRIAL SUPPORT™ (Super Ubiquinol) • 200 mg, 30 softgels	62.00	46.50	39.00	36.00	8011	
00862	CRAN-MAX® • 500 mg, 60 veg. caps	17.50	13.13	11.25		8015	
1424	CRAN-MAX® WITH ELLIROSE™ (Optimized) • 60 veg. caps	18.00	13.50	12.00		8012	
1529	CREATINE CAPSULES • 120 veg. caps	10.95	8.21	6.94		8011	
1746	CREATINE WHEY GLUTAMINE POWDER • 454 grams (vanilla)	30.00	22.50	19.50		8010	
1429	CR MIMETIC LONGEVITY FORMULA • 60 veg. caps	39.00	29.25	27.00		8015	
0407	CURCUMIN® (Super Bio) • 400 mg, 60 veg. caps	38.00	28.50	26.25		8014	12
1924	CURCUMIN® W/GINGER & TURMERONES (Advanced Bio)	30.00	22.50	20.25		8011	2
	30 softgels					8013	30
1804	CYTOKINE SUPPRESS™ W/EGCG • 30 veg. caps	30.00	22.50	20.25		8014	13
	COSMESIS					8014	18
0157	ADVANCED ANTI-GLYCATION PEPTIDE SERUM • 1 oz	53.00	39.75	34.50		8016	51
0154	ADVANCED LIGHTENING CREAM • 1 oz	65.00	48.75	42.75		8016	52
0155	ADVANCED PEPTIDE HAND THERAPY • 4 0z	46.00	34.50	29.25		8016	60
0152	ADVANCED TRIPLE PEPTIDE SERUM • 1 oz	65.00	48.75	42.75		8011	16
0140	ADVANCED UNDER EYE SERUM W/STEM CELLS • .33 oz	49.00	36.75	31.50		8010)1
0139	AMBER SELF MICRODERMABRASION • 2 oz	49.00	36.75	31.50		8011	13
0158	ANTI-AGING FACE OIL • 1 oz	59.00	44.25	39.00		8010)4
0118	ANTI-AGING MASK • 2 oz	72.00	54.00	47.52		8012	29
80151	ANTI-AGING REJUVENATING FACE CREAM • 2 oz	65.00	48.75	42.75		8013	36
30153	ANTI-AGING REJUVENATING SCALP SERUM • 2 oz	46.00	34.50	29.25		8014	15
30134	ANTI-GLYCATION SERUM W/BLUEBERRY & POMEGRANATE EXTRACTS • 1 oz	33.00	24.75	23.51		8014	19
80133	ANTIOXIDANT FACIAL MIST • 2 oz	32.00	24.00	22.80		0065	58
30127	ANTIOXIDANT REJUVENATING FOOT CREAM • 2 oz	45.00	33.75	32.10		0147	
80128	ANTIOXIDANT REJUVENATING FOOT SCRUB • 2 oz	59.00	44.25	38.94		0164	
0117	ANTIOXIDANT REJUVENATING HAND CREAM • 2 oz	64.00	48.00	43.12		0060	
0105	ANTI-REDNESS & ADULT BLEMISH LOTION • 1 oz	74.50	55.88	49.17		0147	
0147	BIOFLAVONOID CREAM • 1 oz	46.00	34.50	29.25		0033	
0144	BROCCOLI SPROUT CREAM • 1 oz	46.00	34.50	29.25			
80156	COLLAGEN BOOSTING PEPTIDE SERUM • 1 oz	59.00	44.25	39.00		0045	
80120	CORRECTIVE CLEARING MASK • 2 oz	64.50	48.38	42.57		8800	
	DNA REPAIR CREAM • 1 oz		36.75	31.50		0168	
	ESSENTIAL PLANT LIPIDS REPARATIVE SERUM • 1 oz		56.21	49.46		0135	8
	SUBTOTAL OF COLUMN 3						

			YO	UR PRIC	E		
ITEM No	o. PRODUCT	Retail Each \$	1 Unit Each	4 Unit Each	10 Unit Each	QTY	Total
80163	EYE LIFT CREAM • 0.5 fl oz	59.00	44.25	39.00			
80123	FACE REJUVENATING ANTIOXIDANT CREAM • 2 oz	69.50	52.13	45.87			
80107	FINE LINE-LESS • 1 oz	74.50	55.88	49.17			
80131	HAIR SUPPRESS FORMULA • 4 oz	59.00	44.25	38.94			
80137	HEALING FORMULA ALL-IN-ONE CREAM • 1 oz	53.00	39.75	34.07			
80115	HEALING MASK • 2 oz	64.50	48.38	42.57			
80102	HEALING VITAMIN K CREAM • 1 oz	79.50	59.63	52.47			
80109	HYALURONIC FACIAL MOISTURIZER • 1 oz	58.00	43.50	38.28			
80110	HYALURONIC OIL-FREE FACIAL MOISTURIZER • 1 oz	58.00	43.50	38.28			
80138	HYDRATING ANTIOXIDANT FACE MIST • 4 oz	39.95	29.96	28.50			
80103	LIFTING & TIGHTENING COMPLEX • 1 oz	74.50	55.88	49.17			
80146	LYCOPENE CREAM • 1 oz	28.00	21.00	19.05			
80135	MELATONIN CREAM • 1 oz	33.00	24.75	20.33			
80114	MILD FACIAL CLEANSER • 8 fl. oz	59.00	44.25	38.94			
80159	MULTI STEM CELL SKIN TIGHTENING COMPLEX • 1 oz	59.00	44.25	39.00			
80122	NECK REJUVENATING ANTIOXIDANT CREAM • 2 oz	64.00	48.00	42.24			
80111	PIGMENT CORRECTING CREAM • 1/2 oz	74.00	55.50	48.84			
80106	REJUVENATING SERUM • 1 oz	74.50	55.88	49.17			
80150	RENEWING EYE CREAM • 1/2 oz	65.00	48.75	42.75			
80142	RESVERATROL ANTI-OXIDANT SERUM • 1 oz	46.00	34.50	29.25			
80112	SKIN LIGHTENING SERUM • 1/2 oz	85.00	63.75	56.10			
80130	SKIN STEM CELL SERUM • 1 0Z	74.00	55.50	51.75			
80143	STEM CELL CREAM W/ALPINE ROSE • 1 oz	66.00	49.50	43.50			
80148	TIGHTENING & FIRMING NECK CREAM • 2 oz	39.00	29.25	26.25			
80161	TRIPLE ACTION VITAMIN C CREAM • 1 oz jar	59.00	44.25	39.00			
80162	ULTIMATE MICRODERMABRASION • 8 fl. oz	39.00	29.25	26.25			
80160	ULTRA EYELASH BOOSTER • 0.25 oz (2 units \$39)	59.00	44.25				
80116	ULTRA LIP PLUMPER • 1/3 oz	64.00	48.00	42.24			
80101	ULTRA WRINKLE RELAXER • 1 oz	89.95	67.46	59.82			
80113	UNDER EYE REFINING SERUM • 1/2 oz	74.50	55.88	49.17			
80104	UNDER EYE RESCUE CREAM • 1/2 oz	74.50	55.88	49.17			
80129	VITAMIN C SERUM • 1 oz	85.00	63.75	56.10			
80136	VITAMIN D LOTION • 4 oz	36.00	27.00	25.25			
80145	VITAMIN E-ESSENTIAL CREAM • 1 oz	28.00	21.00	19.50			
80149	YOUTH SERUM • 1 oz	65.00	48.75	42.75			
	D						
00658	7-KETO® DHEA METABOLITE • 25 mg, 100 caps	28.00	21.00	18.00			
01479	7-KETO® DHEA METABOLITE • 100 mg, 60 veg. caps	40.00	30.00	27.00			
01640	DHA (Vegetarian) • 30 veg. softgels	20.00	15.00	13.50			
00607	DHEA • 25 mg, 100 tablets (Dissolve in mouth)	14.00	10.50	8.81			
01478	DHEA COMPLETE • 60 veg. caps	48.00	36.00	32.40			
00335	DHEA • 25 mg, 100 caps	16.00	12.00	11.00			
00454	DHEA • 15 mg, 100 caps	14.00	10.50	9.00			
00882	DHEA • 50 mg, 60 caps	19.00	14.25	12.75			
01689	DHEA • 100 mg, 60 veg. caps	24.00	18.00	16.50			
01358	DIGEST RC® • 30 tablets	19.95	14.96	12.75			
	SUBTOTAL OF COLUMN 4						

SUPER SALE SAVINGS ON ALL PRODUCTS

I FIVI IV	o. PRODUCT	Retail	1 Usa	4 Unit	10 I	ITEM N	lo. PRODUCT	Retail	1 Unit	4 Heit	10 Unit
		Each \$	Unit Each	Unit Each	Unit Each QTY		QUIATE TOPOTO CONT. 11	Each \$	Unit Each	Unit Each	Each
	DIGESTIVE ENZYMES (Enhanced Super) • 60 veg. caps	22.00		15.00			GINGER FORCE® • 60 liquid caps	34.95		04.50	
	, , , , ,	28.00		18.00		01658	GINKGO BILOBA CERTIFIED EXTRACT™ 120 mg, 365 veg. caps	46.00	34.50	31.50	
	D, L-PHENYLALANINE • 500 mg, 100 veg. caps		14.06	12.00		00756	GLA WITH SESAME LIGNANS (Mega) • 60 softgels	19.50	14.63	13.50	
	DMAE BITARTRATE • 150 mg, 200 veg. caps		13.50	11.25		00345	(L-) GLUTAMINE CAPSULES • 500 mg, 100 veg. caps	14.95	11.21	10.13	
	DNA PROTECTION FORMULA • 60 veg. caps			24.00		00141	(L-) GLUTAMINE POWDER • 100 grams	22.00	16.50	15.00	
	DOG MIX • 100 grams powder					00522	GLUCOSAMINE/CHONDROITIN CAPSULES • 100 caps	38.00	28.50	24.00	
2006	DOPA-MIND™ • 60 veg. tabs	48.00	36.00	32.00		01541	GLUTATHIONE, CYSTEINE & C • 100 veg. caps	20.00	15.00	13.50	
321	DR. PROCTOR'S ADVANCED HAIR FORMULA • 2 oz	39.95	29.96	24.00			GLYCINE • 1,000 mg, 100 veg. caps		9.00	8.10	
320	DR. PROCTOR'S HAIR SHAMPOO • 8 oz	24.95	18.71	16.50		01411				25.50	
	E					01111	100 mg, 60 veg. caps	00.00		20.00	
	ECHINACEA EXTRACT • 250 mg, 60 veg. caps	14.35	10.76	9.38		01620	GREEN COFFEE EXTRACT COFFEEGENIC®	32.00	24.00	21.00	
997	ENDOTHELIAL DEFENSE™ w/POMEGRANATE COMPLETE AND CORDIART™ • 60 softgels	68.00	51.00	46.50		20050	400 mg, 90 veg. caps	00.00	00.50	10.00	
997	ENDOTHELIAL DEFENSE™ w/GLISODIN® • 60 veg. caps	54.00	40 50	36.00			GREEN TEA EXTRACT (Mega) • lightly caffeinated,100 veg. caps			18.00	-
	EPA/DHA (Mega) • 120 softgels	20.00		13.50		00954	GREEN TEA EXTRACT (Mega) • decaffeinated, 100 veg. caps	30.00	22.50	18.00	
	ESOPHAGEAL GUARDIAN (Berry flavor) • 60 chewable tablets			24.00		01074		27.05	20.00		
	, , ,	20.00					5 HTP • 100 mg, 60 caps		20.96	22.22	
1042	EUROPEAN LEG SOLUTION DIOSMIN 95 600 mg, 30 veg. tabs	20.00	13.00	13.50		02002	HAIR, SKIN & NAIL REJUVENATION FORM W/VERISOL® 90 tabs	32.00	24.00	22.00	
706	EXTRAORDINARY ENZYMES • 60 caps	26.00	19.50	18.00		01738	HCA (Garnicia) • 90 veg. caps	17.00	12.75	11.25	
800	(CALIFORNIA ESTATE) EXTRA VIRGIN OLIVE OIL • 500 ml (16.9 fl. oz)	33.00	24.75	22.50			HCACTIVE™ GARCINIA CAMBOGIA EXTRACT • 90 caps	30.00	22.50		
514	EYE PRESSURE SUPPORT W/MIRTOGENOL® • 30 veg. caps	38.00	28.50	25.50			HEPATOPRO • 900 mg, 60 softgels		37.50	34.50	
	F						HUPERZINE A • 200 mcg, 60 veg. caps			27.00	
054	FACE MASTER® PLATINUM • Facial Toning System	199.00	199.00				HYDRODERM® • 1 oz		59.96		
965	FAST-ACTING JOINT FORMULA • 30 caps	39.00	29.25	27.00		00001	1 62	70.00	00.00	10.00	
717	FAST-C® W/DIHYDROQUERCETIN • 120 veg. tabs	26.00	19.50	18.00		01704	IMMUNE MODULATOR W/TINOFEND® • 60 veg. caps	17.00	12.75	11.25	
053	FEM DOPHILUS® • 30 caps	25.95	19.46			00955		29.50	22.13	19.91	
055	FEM DOPHILUS® • 60 caps	39.95	29.96			02005	IMMUNE SENESCENCE PROTECTION FORMULA™ • 60 veg. tabs			27.00	
064	FEMMENESSENCE MACAPAUSE® • 120 veg. caps	34.99	26.24				INNERPOWER™ • 530 grams powder		31.50	27.00	
007	FIBER-IMMUNE SUPPORT (Apple Cinnamon) • 235 grams	34.00	25.50	23.50		01674	,		46.50	43.50	
	FIBRINOGEN RESIST™ • 30 veg. caps		36.75				INTEGRA-LEAN® AFRICAN MANGO IRVINGIA			18.00	
	FLAX SEED (Organic golden) • 14 oz	11.67				01232	150 mg, 60 veg. caps	20.00	21.00	10.00	
	FLORASSIST® GI w/PHAGE TECHNOLOGY • 30 liquid veg. caps			22.50		01677	IRON PROTEIN PLUS • 300 mg, 100 caps	28.00	21.00	19.50	
	FLORASSIST® HEART HEALTH • 60 veg. caps			21.00		01492	IRVINGIA W/PHASE 3™ CALORIE CONTROL COMPLEX	56.00	42.00	36.00	
							(Optimized African Mango) • 120 veg. caps				
	FLORASSIST® ORAL HYGIENE • 30 lozenges			12.75			J, K, L				
	FLORASSIST® BALANCE • 30 liquid veg. caps			21.00			JARRO-DOPHILUS EPS® • 60 veg. caps		17.21		H
	FLORASSIST® MOOD • 60 caps			22.50			K W/ADVANCED K2 COMPLEX (Super) • 90 softgels		22.50		
	FLORASSIST® THROAT HEALTH • 30 lozenges			13.50			KRILL HEALTHY JOINT FORMULA • 30 softgels		24.00	21.75	H
	FOLATE HIGH POTENCY (Optimized) • 5,000 mcg, 30 veg. tablets						KRILL OIL • 60 softgels		25.46		
	FOLATE (Optimized) • 1,000 mcg, 100 veg. tablets		14.25			00316	KYOLIC® GARLIC FORMULA 102 • 200 veg. caps	27.45	20.59		
	FOLATE + VITAMIN B12 (BioActive) • 90 veg. caps		9.00	8.00		00214	KYOLIC® GARLIC FORMULA 105 • 200 caps	28.45	21.34		L
	FORSKOLIN • 10 mg, 60 veg. caps		12.00			00789	KYOLIC® RESERVE • 600 mg, 120 caps	28.95	21.71		
513	FUCOIDAN W/MARITECH® 926 (Optimized) • 60 veg. caps	36.00	27.00	24.75		01681	LACTOFERRIN • 60 caps	44.00	33.00	30.00	
25	G	15:	05.	0.5		00020	LECITHIN • 16 oz granules	18.00	13.50	12.00	
	GAMMA E MIXED TOCOPHEROL/TOCOTRIENOLS • 60 softgels			27.00		02155	LIFE EXTENSION MIX™ • 315 tablets	80.00	60.00	52.00	43.7
2075	GAMMA E MIXED TOCOPHEROL w/ENHANCED SESAME LIGNANS • 60 softgels			21.75		02157	LIFE EXTENSION MIX™ W/EXTRA NIACIN • 315 tablets	80.00	60.00	52.00	43.7
1394	GARLIC (Optimized) • 200 veg. caps	24.95	18.71	15.75		02154	LIFE EXTENSION MIX™ • 490 caps	90.00	67.50	58.00	47.5
100	GASTRO-EASE • 60 veg. caps	44.00	33.00	30.00		02156	LIFE EXTENSION MIX™ POWDER • 14.81 oz	80.00	60.00	52.00	43.7

OFFER ENDS JANUARY 31, 2017

	10 ORDER CALL. 1.954.700.04		_			
ITEM N	o. PRODUCT	Retail	1	UR PRI	10	
		Each \$	Unit Each	Unit Each	Unit Each QTY	/ Total
02165	LIFE EXTENSION MIX™ • 315 tablets w/o copper	80.00	60.00	52.00	43.75	
	LIFE EXTENSION MIX™ • 490 caps w/o copper	90.00	67.50	58.00		
02166	LIFE EXTENSION MIX™ POWDER • 14.81 oz w/o copper	80.00	60.00	52.00	43.75	
01608	LIVER EFFICIENCY FORMULA • 30 veg. caps		13.50	12.00		
	5-LOX INHIBITOR W/APRESFLEX® • 100 mg, 60 veg. caps	22.00	16.50	15.00		
	L-LYSINE • 620 mg, 100 veg. caps	9.00	6.75	6.00		
00455	LYCOPENE (Mega) • 15 mg, 90 softgels	35.00	26.25	22.50		
04.000	M	05.00	40.75	47.50		
	MACUGUARD® OCULAR SUPPORT • 60 softgels		18.75	17.50		
01993	MACUGUARD® OCULAR SUPPORT w/ASTAXANTHIN 60 softgels	44.00	33.00	30.00		
01459	MAGNESIUM CAPS • 500 mg, 100 veg. caps	12.00	9.00	7.50		
01682	MAGNESIUM (CITRATE) • 160 mg, 100 veg. caps	12.00	9.00	7.50		
02107	(EXTEND-RELEASE) MAGNESIUM • 60 veg. caps	13.00	9.75	8.75		
01908	MEDITERRANEAN TRIM WITH SINETROL™-XPUR 60 veg. caps	18.00	13.50	12.00		
01668	MELATONIN • 300 mcg, 100 veg. caps	5.75	4.31	3.75		
01083	MELATONIN • 500 mcg, 200 veg. caps	18.00	13.50	12.00		
00329	MELATONIN • 1 mg, 60 caps	5.00	3.75	3.47		
00330	MELATONIN • 3 mg, 60 veg. caps	8.00	6.00	5.16		
00331	MELATONIN • 10 mg, 60 veg. caps	28.00	21.00	18.00		
00332	MELATONIN • 3 mg, 60 veg. lozenges	8.00	6.00	5.16		
01734	MELATONIN (Fast-Acting Liquid) • 2 fl. oz (Citrus-Vanilla)	12.00	9.00	8.25		
01787	MELATONIN TIMED RELEASE • 300 mcg, 100 veg. tabs	12.00	9.00	8.25		
01788	MELATONIN TIMED RELEASE • 750 mcg, 60 veg. tablets	8.00	6.00	5.25		
01786	MELATONIN TIMED RELEASE • 3 mg, 60 veg. tabs	12.00	9.00	8.25		
02101	MEMORY PROTECT • 36 day supply	24.00	18.00	16.00		
01536	METHYLCOBALAMIN • 1 mg, 60 veg. lozenges (vanilla)	9.95	7.46	6.00		
01537	METHYLCOBALAMIN • 5 mg, 60 veg. lozenges (vanilla)	32.00	24.00	18.75	17.25	
00709	MIGRA-EEZE™ (Butterbur) • 60 softgels	33.00	24.75	22.00		
01522	MILK THISTLE (European) • 60 veg. caps	34.00	25.50	22.50		
01922	MILK THISTLE (European) • 60 softgels	28.00	21.00	18.75		
01925	MILK THISTLE (European) • 120 softgels	44.00	33.00	30.00		
01940	MIRAFORTE w/STANDARDIZED LIGNANS (Super) • 120 veg caps	62.00	46.50	42.00		
01869	MITOCHONDRIAL BASICS W/BIOPQQ® • 30 caps	44.00	33.00	30.00		
01868	MITOCHONDRIAL ENERGY OPTIMIZER w/BIOPQQ® • 120 caps	72.00	54.00	48.00		
00065	MK-7 • 90 mcg, 60 softgels	28.00	21.00	18.75		
00451	MSM (Methylsulfonylmethane) • 1,000 mg, 100 caps	14.00	10.50	8.96		
	N					
01534	N-ACETYL-L-CYSTEINE • 600 mg, 60 veg. caps	14.00	10.50	9.25		
01904	NAD+ CELL REGENERATOR™ • 100 mg, 30 veg. caps	34.00	25.50	19.50		
00066	NATTOKINASE • 60 softgels	25.50	19.13			
01807	NATURAL APPETITE SUPPRESS (Advanced) • 60 veg. caps	38.00	28.50	25.50		
00984	NATURAL BP MANAGEMENT • 60 tablets	44.00	33.00	30.00		
01892	NATURAL ESTROGEN • 60 veg. tabs	38.00	28.50	25.50		
01626	$\textbf{NATURAL SEX FOR WOMEN} \bullet \textbf{50+} \ (\texttt{Advanced}) \bullet \textbf{90 veg. caps}$	59.00	44.25	34.00		
01444	NATURAL SLEEP® • 60 veg. caps	13.00	9.75	7.50		
	SUBTOTAL OF COLUMN 7					

	ORDER ORLINE VISIT. WWW.ElieExt			UR PRIC	F	_	
ITEM N	o. PRODUCT	Retail Each	1 Unit	4 Unit	10 Unit	070/	Tatal
01551	NATURAL SLEEP® w/ MELATONIN (Enhanced) • 30 caps	\$ 22.00	Each 16.50	Each 15.00	Each	QIY	IOLAI
	NATURAL SLEEP® W/O MELATONIN (Enhanced) • 30 caps	20.00	15.00	13.50		ī	
	NATURAL SLEEP® MELATONIN • 5 mg, 60 veg. caps	18.00	13.50	12.00			
	NATURAL STRESS RELIEF • 30 veg. caps		21.00	18.00		ī	
	NERVIA® • 90 softgels		40.46				
	NEURO-MAG® MAGNESIUM L-THREONATE • 90 veg. caps		30.00	27.00			
	NEURO-MAG® MAGNESIUM L-THREONATE w/CALCIUM & VITAMIN D3 • 25 grams • Lemon flavor	40.00	30.00	27.00			
01990	NITROVASC w/CORDIART™ • 30 veg. caps	18.00	13.50	12.00			
01903	NK CELL ACTIVATOR™ • 30 veg. tablets	45.00	33.75	31.50			
00373	NO FLUSH NIACIN • 800 mg, 100 caps	19.00	14.25	12.75			
	0						
01824	OLIVE LEAF VASCULAR SUPPORT w/CELERY SEED EXTRACT (Advanced) • 60 veg. caps	36.00	27.00	24.00			
01988	OMEGA-3 PLUS EPA/DHA w/SESAME LIGNANS, OLIVE EXTRACT, KRILL & ASTAXANTHIN (SUPER) • 120 softgels	45.00	33.75	31.50	24.75		
01983	OMEGA-3 EPA/DHA w/SESAME LIGNANS & OLIVE EXTRACT (Super) • 60 softgels	18.00	13.50	12.00	9.38		
01982	OMEGA-3 EPA/DHA w/SESAME LIGNANS & OLIVE EXTRACT (Super) • 120 softgels	32.00	24.00	21.00	17.05		
01984	OMEGA 3 EPA/DHA w/SESAME LIGNANS & OLIVE EXTRACT (Super) • 120 enteric coated softgels	34.00	25.50	23.25	18.00		
01985	OMEGA 3 EPA/DHA w/SESAME LIGNANS & OLIVE EXTRACT (Super) • 60 enteric coated softgels	20.00	15.00	13.50	10.50		
01986	OMEGA 3 EPA/DHA w/SESAME LIGNANS & OLIVE EXTRACT (Super) • 240 small softgels	32.00	24.00	21.00	17.25		
01991	ONCE-DAILY HEALTH BOOSTER • 60 softgels	54.00	40.50	38.00			
02113	ONE-PER-DAY • 60 tablets	22.00	16.50	15.00			
01328	ONLY TRACE MINERALS • 90 veg. caps	15.00	11.25	9.38			
	P						
01789	PALMETTOGUARD® SAW PALMETTO W/BETA-SITOSTEROL 30 softgels	15.00	11.25	10.50	9.00		
01790	PALMETTOGUARD® SAW PALMETTO/ NETTLE ROOT W/BETA-SITOSTEROL • 60 softgels	28.00	21.00	19.50	18.00		
01323	PEAK ATP® WITH GLYCOCARN® • 60 veg. caps	54.00	40.50	37.50			
00342	PECTA SOL-C MODIFIED CITRUS PECTIN • 454 grams powder	109.95	93.46				
*01080	PECTA SOL-C® MODIFIED CITRUS PECTIN • 270 veg. caps	79.95	67.96				
01811	PEONY IMMUNE • 60 veg. caps	36.00	27.00	24.00			
00673	PGX® PLUS MULBERRY (WellBetX®) • 180 veg. caps	34.95	26.21				
01953	POMEGRANATE COMPLETE • 30 softgels	24.00	18.00	15.75			
00956	POMEGRANATE FRUIT EXTRACT • 30 veg. caps	19.50	14.63	13.16			
**01837	POMI-T [®] • 60 veg. caps	35.00	26.25	24.00			
01500	PQQ CAPS W/BIOPQQ® • 10 mg, 30 veg. caps	24.00	18.00	13.50	12.00		
01647	PQQ CAPS W/BIOPQQ® • 20 mg, 30 veg. caps	40.00	30.00	24.00	21.00		
00302	PREGNENOLONE • 50 mg, 100 caps	26.00	19.50	16.50			
00700	PREGNENOLONE • 100 mg, 100 caps	30.00	22.50	20.25			
**01373	PRELOX® NATURAL SEX FOR MEN® • 60 tablets	52.00	39.00	36.00		ĺ	
01576	PREVAGEN® • 30 caps	60.00	45.00				
*01577	PREVAGEN® ES • 30 caps	70.00	60.00			Î	
	PROBOOST™ THYMIC PROTEIN A • 30 packets	66.60	49.95			Î	
	SUBTOTAL OF COLUMN 8						

SUPER SALE SAVINGS ON ALL PRODUCTS

TO ORDER CALL: 1.954.766.8433 or 1.800.544.4440 ■ TO ORDER ONLINE VISIT: www.LifeExtension.com

				UR PRI			L PROPUST
ITEM N	lo. PRODUCT	Retail Each	1 Unit	4 Unit	10 Unit Each QTY Total	ITEM N	lo. PRODUCT
01441	PROGESTA-CARE® • 4 oz cream	\$ 36.39	Each 27.29	Each 25.72	Each QTY lotal	01934	SAMe (S-ADENOSYL-METHIONINE)
01928	PROSTATE FORMULA (Ultra NAT) • 60 softgels	38.00	28.50	26.25	24.00	01740	400 mg, 60 enteric coated tablets SEA-IODINE™ • 1,000 mcg, 60 veg. caps
01909	PROSTAPOLLEN™ (Triple strength) • 30 softgels	28.00	21.00	18.75			, 0, 0
01742	PROTEIN-ISOLATE (Whey) Vanilla • 403 grams	30.00	22.50	19.50			SELENIUM • 2 fl. oz dropper
01743	PROTEIN-ISOLATE (Whey) Chocolate • 437 grams	30.00	22.50	19.50			SE-METHYL L-SELENOCYSTEINE • 200 mcg, 90 veg. co
01770	(30.00	22.50	19.95			SHADE FACTOR • 120 veg. caps
01771	500 grams	00.00	00.50	10.05			SILYMARIN • 100 mg, 90 veg. caps
01771	PROTEIN CONCENTRATE (New Zealand Whey) Chocolate 640 grams	30.00	22.50	19.95		01249	
01812	PROVINAL® PURIFIED OMEGA-7 • 30 softgels	27.00	20.25	18.00			SKIN RESTORING PHYTOCERAMIDES w/LIPOWHEAT®
01676	PS CAPS (Phosphatidylserine) • 100 mg, 100 veg. caps	54.00	40.50	36.00		01000	30 liquid veg. caps
01508	PTEROPURE® Pterostilbene • 50 mg, 60 veg. caps	32.00	24.00	22.50		00961	SODZYME® w/GLISODIN® & WOLFBERRY • 90 veg. ca
01209	PUMPKIN SEED EXTRACT (Water-soluble) • 60 veg. caps	20.00	15.00	13.50		00657	SOLARSHIELD® SUNGLASSES • Smoke color
01637		64.00	48.00	45.00		00432	STEVIA™ (Better) • 100 packets, 1 gram each
	100 mg, 60 veg. caps					00438	STEVIA™ ORGANIC LIQUID SWEETENER (Better) • 2 oz
01217	PYRIDOXAL 5'-PHOSPHATE • 100 mg, 60 veg. caps Q, R	22.00	16.50	14.85		01476	STRONTIUM • 750 mg, 90 veg. caps
01309		22.00	16.50	15.00		01649	SUPER ABSORBABLE SOY ISOFLAVONES • 60 veg. ca
	RED YEAST RICE (Bluebonnet) • 600 mg, 60 veg. caps		13.20	13.00		01778	SUPER SELENIUM COMPLEX • 200 mcg, 100 veg. cap
	REGIMINT • 60 enteric-coated caps		14.96	14.00			Т
01708			22.50	20.25		02023	TART CHERRY W/CHERRYPURE® 60 veg. caps
	REJUVENEX® BODY LOTION • 6 oz		18.00	14.85	12 75	01827	TAURINE • 1,000 mg, 90 veg. caps
01621			48.75	37.50	12.70	01918	TEAR SUPPORT w/MAQUIBRIGHT® • 60 mg, 30 veg. c
01220			39.00	33.00	29 25	00133	L-TAURINE POWDER • 300 grams
	REJUVENIGHT® (Ultra) • 2 oz		29.96	27.00	20:20	*13685	TEN MUSHROOM FORMULA® • 120 veg. caps
	RESVERATROL W/PTEROSTILBENE • 100 mg, 60 veg. caps		27.00	24.00		01304	THEAFLAVIN STANDARDIZED EXTRACT • 30 veg. caps
02031			31.50	27.00		01683	(L) THEANINE • 100 mg, 60 veg. caps
	(Optimized) • 30 veg. caps					***01038	THERALAC® PROBIOTICS • 30 caps
02030	RESVERATROL (Optimized) • 60 veg. caps	46.00	34.50	31.00		00668	THYROID FORMULA (Metabolic Advantage™) • 100 ca
00889	RHODIOLA EXTRACT • 250 mg, 60 veg. caps	14.00	10.50	9.00		00349	TMG POWDER • 50 grams
01900	RIBOGEN™ FRENCH OAK WOOD EXTRACT 200 mg, 30 veg. caps	36.00	27.00	24.75		01859	TMG • 500 mg, 60 liquid veg. caps
00972	(D) RIBOSE POWDER • 150 grams	27 50	20.63	18.56		01400	TOCOTRIENOLS (Super-absorbable) • 60 softgels
	(D) RIBOSE TABLETS • 100 veg. tabs		24.00	21.00		01278	TOOTHPASTE • 4 oz (Mint) tube
	RICH REWARDS® BREAKFAST GROUND COFFEE • 12 oz. baq			21.00		01917	TRANQUIL TRACT™ • 60 veg. caps
01730	RICH REWARDS® BREAKFAST BLEND GROUND COFFEE		11.25	10.50		01468	TRIPLE ACTION CRUCIFEROUS VEGETABLE EXTRACT 60 veg. caps
01729	Natural Mocha • 12 oz. bag RICH REWARDS® BREAKFAST BLEND GROUND COFFEE	15.00	11.25	10.50		01469	TRIPLE ACTION CRUCIFEROUS VEGETABLE EXTRACT w/RESVERATROL • 60 veg. caps
01612	Natural Vanilla • 12 oz. bag RICH REWARDS® BREAKFAST BLEND WHOLE BEAN COFFEE	13.00	9.75				TRIPLE ACTION THYROID • 60 veg. caps
01610	12 oz. bag RICH REWARDS® DECAFFEINATED ROAST GROUND COFFEE	14.00	10.50				TRI SUGAR SHIELD® • 60 veg. caps TRUFIBER™ • 180 grams
	12 oz. bag					01389	TRUFLORA® PROBIOTICS • 32 veg. caps
	R-LIPOIC ACID (Super) • 240 mg, 60 veg. caps		36.75	33.75		01722	L-TRYPTOPHAN • 500 mg, 90 veg. caps
00070	RNA CAPSULES • 500 mg, 100 caps	17.95	13.46	12.12		01721	TRYPTOPHAN PLUS (Optimized) • 90 veg. caps
01.420	SAEEDAN W/SATIEDEAL® (Ontimized) a 60 year cans	26.00	27.00	24.00		02116	TWO-PER-DAY • 60 tablets
	SAFFRON W/SATIEREAL® (Optimized) • 60 veg. caps		27.00	24.00		02115	TWO-PER-DAY • 120 tablets
J 1935	SAMe (S-ADENOSYL-METHIONINE) 200 mg, 30 enteric coated tablets	25.00	18.75	16.50		02114	TWO-PER-DAY • 120 caps
01933	SAMe (S-ADENOSYL-METHIONINE) 400 mg, 30 enteric coated tablets	36.00	27.00	24.00		00326	L-TYROSINE • 500 mg, 100 tablets
	SUBTOTAL OF COLUMN 9						SUBTOTAL OF COLUMN 10

10 Unit Each QTY Total

4 Unit Each

45.00

5.40

7.50

12.00

9.50

8.63

9.00

12.00

12.66

12.00

8.25 9.00

6.50

16.50

7.13

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Retail

66.00 49.50

8.00 6.00

11.95 8.96

11.00 8.25 18.00 13.50

25.00 18.75

12.99 9.74

9.95 7.46

11.00 8.25 20.00 15.00 13.50

44.00 33.00 30.00 14.00 10.50

25.00 18.75 17.25

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39.95 33.96

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21.95 16.46 14.00 10.50

13.00 9.75 30.00 22.50 21.00 9.50 7.13

52.00 39.00 34.50 24.00 18.00

32.00 24.00 22.20

36.00 27.00 24.00 36.00 27.00 24.00 32.95 24.71 42.95 32.21 33.00 24.75 22.50 32.00 24.00 21.75 10.50 7.88

20.00 15.00 13.50 22.00 16.50

13.50 10.13

24.00 18.00 15.38 47.95 35.96

OFFER ENDS JANUARY 31, 2017

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	TO ORDER CALL: 1.954.766.8	433 ()	300.6)44.	1440
ITEM N	o. PRODUCT	Retail	Y0	UR PRIO	10	
II EIVI IV	U. FNUDUCI	Each \$	Unit Each	Unit Each	Unit	QTY Total
	U, V					
01921	URIC ACID CONTROL • 60 veg. caps	24.00	18.00	16.50		
00213	VANADYL SULFATE • 7.5 mg, 100 veg. tablets	15.00	11.25	9.38		
02102	VENOFLOW • 30 veg. caps	52.00	39.00	36.00		
00408	VENOTONE • 60 caps	18.95	14.21	12.00		
01327	VINPOCETINE • 10 mg, 100 veg. tablets	18.00	13.50	10.50		
00372	VITAMIN B3 NIACIN • 500 mg, 100 caps	7.65	5.74	4.99		
00098	VITAMIN B5 • 500 mg, 100 caps (Pantothenic Acid)	10.50	7.88	7.04		
01535	VITAMIN B6 • 250 mg, 100 veg. caps	12.50	9.38	8.25		
00361	VITAMIN B12 • 500 mcg, 100 lozenges	8.75	6.56	5.44		
01634	VITAMIN C w/DIHYDROQUERCETIN 1,000 mg, 60 veg. tablets	10.00	7.50	6.75		
00927	VITAMIN C w/DIHYDROQUERCETIN 1,000 mg, 250 veg. tablets	25.50	19.13	17.44		
00084	VITAMIN C POWDER (BUFFERED) • 454 grams	23.95	17.96	16.50		
01736	VITAMIN C-MAGNESIUM CRYSTALS (EFFERVESCENT) 180 grams	20.00	15.00	13.50		
01732	VITAMIN D3 • 2,000 IU, 1 fl. oz, Mint flavor	28.00	21.00	18.75		
01753	VITAMIN D3 • 1,000 IU, 90 softgels	7.00	5.25	4.50		
01751	VITAMIN D3 • 1,000 IU, 250 softgels	12.50	9.38	8.44		
01713	VITAMIN D3 • 5,000 IU, 60 softgels	10.00	7.50	6.50		
01718	VITAMIN D3 • 7,000 IU, 60 softgels	14.00	10.50	9.45		
01758	VITAMIN D3 W/SEA-IODINE™ • 5,000 IU, 60 caps	14.00	10.50	9.38		
00864	VITAMIN D3 LIQUID • 2,000 IU, 1 fl. oz	28.00	21.00	18.75		
01840	VITAMINS D AND K W/SEA-IODINE™ • 60 caps	24.00	18.00	16.50		
01863	VITAMIN E (Natural) • 400 IU, 90 softgels	28.00	21.00	19.50	18.00	
01936	VITAMIN K2 (Low dose) • 45 mcg, 90 softgels	18.00	13.50	12.00		
01902	WAIST-LINE CONTROL™ • 120 veg. caps	42.00	31.50	28.50		
	X, Y					
01919	X-R SHIELD • 90 veg. caps	15.00	11.25	9.75		
00409	XYLIWHITE™ MOUTHWASH • 16 oz	10.00	7.50			
	Z					
01813	ZINC HIGH POTENCY • 50 mg, 90 veg. caps	7.95	5.96	5.25		
01561	ZINC LOZENGES • 60 veg. lozenges	9.00	6.75	6.00		
01961	ZINC LOZENGES (Enhanced) • 30 veg. lozenges	12.00	9.00	6.00		
**01051	ZYFLAMEND® WHOLE BODY • 120 liquid veg. caps	72.95	54.71			
	BOOKS					
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- **PQQ:** This micronutrient has been shown to trigger the growth of new mitochondria in aging cells!7 PQQ also activates genes involved in protecting the delicate structures within the mitochondria.8-11
- **TAURINE:** Supports whole-body health and boosts new brain cell formation in the area of the brain connected to learning and memory.⁶
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Benfotiamine	150 mg
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WHAT'S INSIDE

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7 GREATEST THREAT TO LONGEVITY

The cause of most **disability** and **death** in persons over age 50 are abnormal **blood clots** that block **arteries** and **veins**. Fortunately, the healthy practices that readers of this magazine follow <u>reduce</u> risk of these occlusive **vascular** disorders.



86 METFORMIN REDUCES THE INCIDENCE OF OPEN-ANGLE GLAUCOMA

Scientists have discovered persuasive data that the **AMPK-activating** drug **metformin** may help protect against glaucoma. **Life Extension**° encourages those at risk to speak to their doctor about these findings.



58 ACHIEVING OPTIMAL SELENIUM STATUS

Volumes of research reveal how different **selenium** compounds exert their own unique effects in impeding malignant transformation.



26 KILL HARMFUL BACTERIA IN YOUR INTESTINES

When bacteria-killing **phages** were added to **probiotics**, there was a **40-100-fold** <u>increase</u> in beneficial **intestinal flora** with huge <u>reductions</u> of pathogenic *E.coli*. These findings represent a novel opportunity to achieve *optimal* intestinal flora balance.



48 CANCER PROTECTIVE VEGETABLE EXTRACT

Data show that **apigenin**, a polyphenol found in parsley and celery, can effectively starve cancer cells, guard DNA against toxins, and block malignant cell propagation.



36 PREVENTION OF DEEP VEIN THROMBOSIS

Sitting more than four hours a day increases the risk of potentially lethal blood clots, known as **deep vein thrombosis** (**DVT**), by **48%**. Researchers have developed two natural compounds that drastically *reduce* platelet aggregation- and fibrininduced clots.