Why Blood Tests Don't Save More Lives



# New Life-Saving Data on VITAMIN K

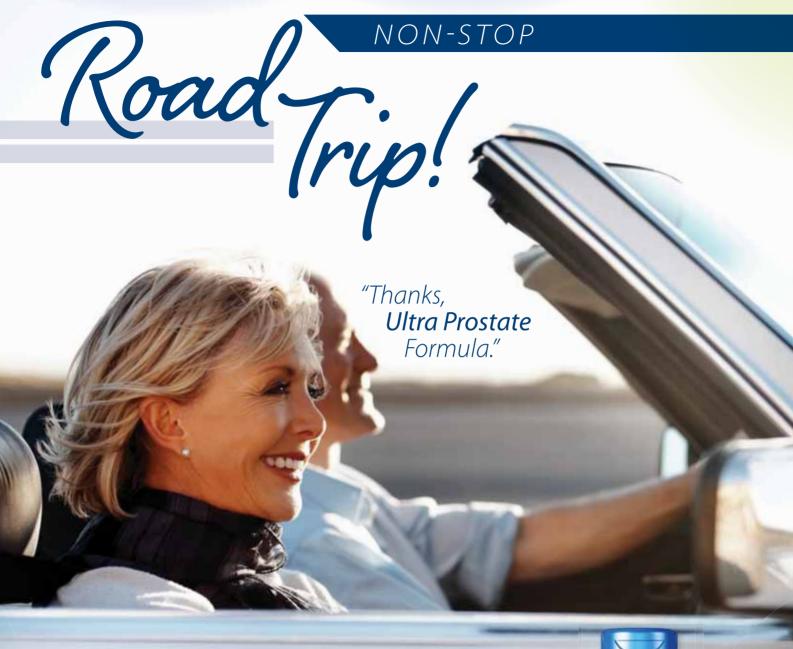
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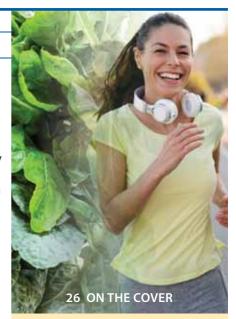
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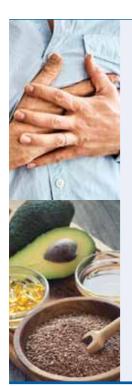
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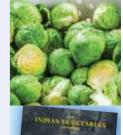
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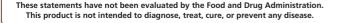
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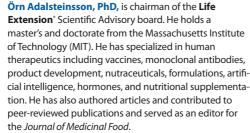
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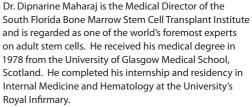
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BY WILLIAM FALOON

# Why Blood Tests Are Not Saving More Lives

Back in the 1950s, a scientist uncovered the link between **LDL cholesterol** and **atherosclerosis**. 1,2

Critics argued that **heart attacks** occurred in some people who did <u>not</u> have excess **LDL**.

What was not known in early years is that there are <u>different</u> types of **LDL** particles.

If your **LDL** surface contains high levels of a protein called **apolipoprotein B**, it is more likely to damage arterial walls and set the stage for **atherosclerosis**.

In some populations, those who maintain <u>low</u> lifetime **apolipoprotein B** levels have an approximately **90%** decreased risk of **coronary artery** disease.<sup>3</sup>

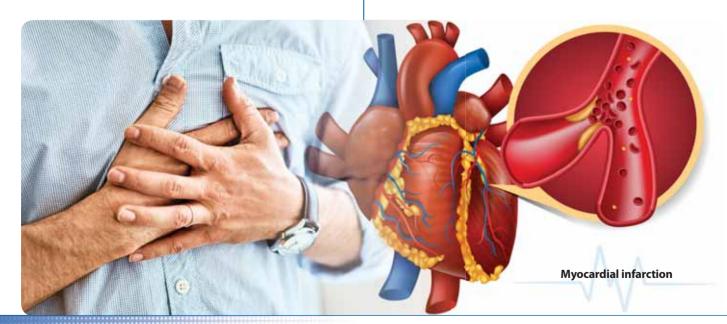
What impresses us is evidence showing **regression** of **arterial plaque** when **apolipoprotein B** blood levels are reduced.<sup>4</sup>

Despite intensive educational efforts, **apolipoprotein B** blood tests are not routinely incorporated into primary care medicine. The tragic result is a failure to prevent heart attacks, strokes, and other occlusive arterial diseases.

For *Life Extension*® readers, this problem was resolved when **apolipoprotein B** was added to comprehensive **blood tests** they undergo each year.

We know today that **atherosclerosis** is a result of a myriad of risk factors. These include elevated **glucose**, **insulin**, **homocysteine**, **triglycerides**, and **inflammation**, <sup>5,6</sup> along with **hormone** imbalances.

This article reveals how coronary artery stenting, bypass surgery, and sudden death can be reduced via inclusion of **apolipoprotein B** in annual **blood test** panels.



**Heart attack** incidence peaked in the year 1968, accounting for 37% of all deaths in the United States.<sup>7,8</sup>

The epidemic was alarming, as it often targeted otherwise healthy people at the peak of their careers.

Advances in **prevention** and **treat**ment led to marked reductions in sudden death and improved survival in those stricken.

While heart disease remains the leading killer, it is responsible for 25% of American deaths today compared to 37% in 1968.9,10

As **blood testing** evolves to better detect artery-damaging factors, further declines in coronary artery disease and ischemic stroke are likely.

#### **Role of Lipoproteins**

**Apolipoprotein B** is linked to the initiation and propagation of atherosclerosis.11

High levels of apolipoprotein B penetrate the inner arterial wall (the endothelium) and set the stage for blockage of blood flow.12-14

Those whose **blood test** shows elevated cholesterol and apolipo**protein B** are especially at risk. 15-17

To put this in perspective, higher apolipoprotein B levels have been associated with a 60% increase in coronary heart disease when total cholesterol and HDL are in safer ranges.18

In people with higher levels of cholesterol and apolipoprotein **B**, along with <u>lower</u> **HDL**, there is a 160% increased incidence of coronary disease.18

Elevated **apolipoprotein B** is a more reliable marker for cardiovascular disease than LDL, HDL, and total cholesterol. 19-22

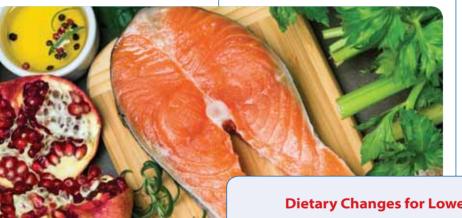
The good news is that apolipopro**tein B** can be easily tested in one's blood and reduced to safe ranges if need be.

#### **Eat Walnuts**

A myriad of studies associates walnut consumption with reduced risk of coronary heart disease.<sup>23-26</sup>

A study assessed a range of cardiovascular risk markers in response to a diet containing about 1.52 ounces of walnuts daily for eight weeks compared to a typical Western diet.<sup>23</sup>

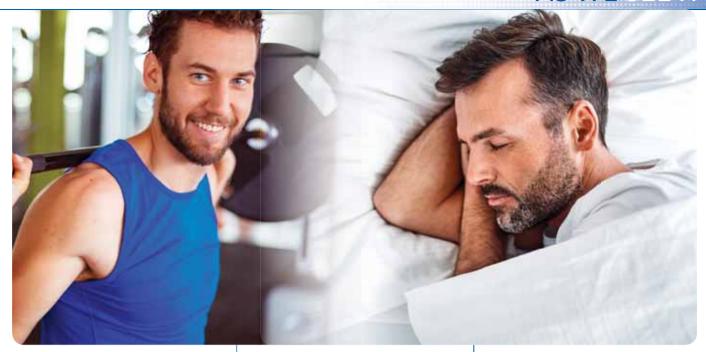
Two benefits of walnut consumption in this trial were a modest non-HDL cholesterol reduction and a more pronounced 5.5% reduction in apolipoprotein B.23



#### **Dietary Changes for Lowering Apolipoprotein B**

Those who follow a more **Mediterranean** style diet have a reduced incidence of cardiovascular disease and lower apolipoprotein B blood levels.<sup>29,30</sup> Here are some additional dietary suggestions:

- Eat far fewer saturated fats (such as butter, palm oil, coconut oil, meat fats, and milk fats); trans fats (such as margarine, vegetable shortening, and partially hydrogenated oils); and dietary cholesterol.
- Eat far more fiber-rich foods (especially soluble fiber from beans, yams, tree nuts, oats, barley, and berries).
- Eat vegetable proteins (such as tofu and beans) in place of meat and poultry.
- Eat fewer added sugars and refined grains (such as white flour).
- Eat more cold-water fish.



These findings led the researchers to conclude that the observation of reduced cardiovascular events in people who eat walnuts may be explained in part by reduced non-HDL cholesterol and apolipoprotein B.23

A follow-up analysis looked at previous studies that evaluated consumption of tree nuts and found reductions in apolipoprotein B in type II diabetics who consumed higher amounts of tree nuts.<sup>27</sup>

#### **Avoid Sugar**

An investigation looked at associations of apolipoprotein B blood levels with education, lifestyle factors, and dietary patterns.28

The intake of **sucrose** and foods containing added sugar (such as pastries, sweets, jam and sugarsweetened beverages) correlated with higher apolipoprotein B levels.28

This study identified other unhealthy factors such as smoking, obesity, and low physical activity with unfavorable lipoprotein profiles.

#### **Exercise Training**

A one-year study of overweight, healthy men found a reduction in apolipoprotein B (but not LDL cholesterol) in response to exercise training.31

This led the authors of the study to postulate that exercise might make **LDL** less atherogenic by reducing apolipoprotein B.

#### **Importance** of Sleep

A Chinese study looked at the relationship of sleep duration and apolipoprotein B blood levels using data from 7,381 subjects.32

The participants were divided into three categories according to sleep duration: less than six hours, seven to eight hours, and more than nine hours.32

The study found that **short sleep** duration (under 6 hours) in women was associated with 1.75-fold greater odds of elevated apolipoprotein B compared to women who got 7-8 hours of sleep per night.32

Longer sleep duration in men was associated with decreased **apolipoprotein B**. The study authors concluded:

"Sleep hygiene management could serve to treat and prevent cardiovascular diseases by altering unfavorable apolipoprotein profile."32

#### **Thyroid Hormones** and Lipoproteins

People with insufficient thyroid hormones can have elevated cholesterol, LDL, triglycerides and apolipoprotein B.33-35

In some cases, people are prescribed statin drugs to lower cholesterol when the appropriate use of a thyroid hormone medication would normalize their lipid profile.

One of several studies on this topic concluded by stating:

"A reduction in lipid and lipoprotein levels after thyroid hormone replacement in our study cohort resulted in a less atherogenic profile."36

#### **Effects of Fish Oil**

The effects of **fish oil** (**1,800 mg** of EPA and **1,200 mg** of DHA a day) were studied on 10 healthy males for four weeks.<sup>37</sup>

The result was about a **30%** reduction in production of **apolipoprotein B** and reductions in other vascular risk factors like triglycerides and VLDL.<sup>37</sup>

Note that this **EPA/DHA** dose of **3,000 mg** a day is much *higher* than many fish oil studies that sometimes use <u>less</u> than **1,000 mg** a day.

#### Acute Risk of Excess Apolipoprotein B

A review of factors that underlie **acute heart attack** revealed a startling finding.

Amongst those with the greatest risk were current **smokers** and people with elevated blood ratios of **apolipoprotein B** to apolipoprotein A1.<sup>38</sup> (Apolipoprotein A1 is the major protein component of beneficial HDL.)

This comprehensive analysis identified other known causes of **heart attack** such as **hypertension** and

diabetes. It then listed protective factors including daily consumption of fruits and vegetables, exercise, and modest consumption of alcohol.<sup>38</sup>

Study participants who were non-diabetic, did not smoke, and had no lipid abnormalities, including high blood ratio of **apolipoprotein B** to apolipoprotein A1, showed large reductions in **acute heart attack** risk.<sup>38</sup>

A separate study found that the **cruciferous vegetable** extract **I3C** (indole-3-carbinol) reduced **apolipoprotein B** secretion from liver cells as much as **56%**. <sup>39</sup> This may be a mechanism by which plant foods like broccoli reduce cardiovascular risks.

#### Drugs That Lower Apolipoprotein B

Prescription drugs that lower **cholesterol** and **LDL** <u>also</u> reduce blood levels of **apolipoprotein B**.<sup>40</sup>

A potentially beneficial mechanism of **statin** drugs and the new **PCSK-9 inhibitors** (like Repatha®) is that they can lower LDL down to around **30 mg/dL**. Normal target goals for **LDL** are below **70-100 mg/dL**.<sup>41</sup>

Some people tolerate statin drugs, while others encounter unbearable side effects.<sup>42</sup>

We are intrigued by the potential of drugs like **Repatha®** to potentially reverse atherosclerosis by driving **lipoproteins** down to the levels of those seen in healthy teenagers. The problem is that even at a reduced price of **\$5,900** a year, Repatha is cost-prohibitive and seldom covered by insurance.

I know most readers of *Life Extension Magazine* seek **non-drug** solutions whenever possible.

With the availability of **apolipoprotein B** blood tests, one can choose to utilize some of the natural interventions described so far and verify efficacy with follow-up **blood tests**.

#### Apolipoprotein B and Fasting Insulin—No Added Cost!

Those with high **apolipoprotein B** blood levels are at greater risk for coronary artery disease.<sup>47</sup>

Last year we added the **apolipoprotein B** to the **Male** and **Female Blood Panels** at <u>no</u> additional cost.



#### What is Apolipoprotein B?

**Apolipoprotein B** is found on all cholesterol particles except **HDL**.<sup>43</sup>

This includes **LDL** and **VLDL** (very-low-density-lipoproteins).

High levels of **VLDL** increase the synthesis and secretion of **apolipoprotein B.**<sup>44</sup>

Higher **apolipoprotein B** generally equates to a higher amount of glycated and oxidized **LDL** particles, which are initiators of arterial plaque.<sup>45,46</sup>

If an **apolipoprotein B** blood test result shows high levels, steps can be initiated to correct this.

This year we added **fasting** insulin to the Male and Female **Panels** and kept our same low price. High fasting insulin levels may indicate that type II diabetic complications are manifesting.

High insulin levels may also interfere with one's ability to lose weight.

#### **Annual Lab Test Super Sale**

The high cost of conventional blood testing precludes many people from availing themselves of this life-saving diagnostic.

We resolved this 23 years ago by enabling our readers to order lowcost **blood tests** direct, and then to visit a **drawing station** in their area at their convenience.

Results come back in less than a week and are emailed and mailed to vou.

If you have any questions, our Wellness Specialists are available to assist seven days a week at no charge.

Once a year, we **discount** prices of all lab tests. This serves as a convenient reminder to have one's annual tests performed and save up to **50%** in the process.

This year's lab test super sale expires on June 3, 2019.

To view the extensive array of tests included in these comprehensive panels, please see the full description by turning this page.

To order the new **Male** or **Female** Blood Panels (that include apolipoprotein B) for \$199, call **1-800-208-3444** or log on to:

www.LifeExtension.com/blood

#### **What Are Optimal Blood Levels of Apolipoprotein B?**

A number of published studies have identified what apolipoprotein **B** blood levels place people at greater risk of cardiovascular events.

The chart below is for people without preexisting vascular disease, diabetes or other major risk factors. The chart lists optimal apolipoprotein B as under 80 mg/dL.

Those at high risk of arterial occlusion, however, should attempt to reduce apolipoprotein B to under 60 mg/dL. This will likely require dietary modifications and use of certain drug therapies.

When it comes to ideal apolipoprotein B levels, the lower the better. The following reference ranges reflect Life Extension's general guidance for **apolipoprotein B** status when you get your blood test results back:

Optimal	<80 mg/dL <60 mg/dL (For those with high risk of arterial occlusion)
Borderline	80-99 mg/dL
High	100-120 mg/dL
Very High	≥120 mg/dL

If your apolipoprotein B levels are not in optimal ranges, there are natural and pharmaceutical approaches to lowering them described in this article.

For longer life,

William Faloon, Co-Founder Life Extension Foundation **Buyers Club** 

**P.S.** Turn this page to view all tests included in the Male and Female **Blood Panels**.

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#### AS WE SEE IT

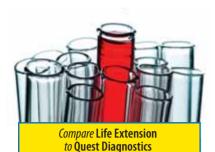
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#### **Comprehensive Blood Tests at Low Super Sale Prices**

The value of the **Male** and **Female Blood Panels** improves as <u>more</u> tests are added at no additional cost. Last year, **apolipoprotein B** was added to better measure **vascular** risk. This year, **fasting insulin** has been added to provide a clearer picture of one's metabolic status. If elevated, steps to reduce **fasting insulin** can help assist with **weight loss** and achieving better **glucose** control.



#### **OUEST DIAGNOSTICS**

TEST NAME	RETAIL	PRICE
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GGT	\$	50.62
Iron	\$	46.12
Lipid Panel Standard	\$	148.00
Complete Blood Count	\$	45.50
Free/Total Testosterone	\$	283.46
Estradiol	\$	223.85
DHEA-S	\$	213.72
Vitamin D 25-Hydroxy	\$	241.84
Homocysteine	\$	227.22
C-Reactive Protein		
(high sensitivity)	\$	59.00
Apolipoprotein B	\$	25.00
Insulin	\$	29.41
HbA1C	\$	74.25
TSH	\$	130.49
PSA	\$	148.48
Draw Fees	\$	21-22
TOTAL	\$2	,086.77

#### **LIFE EXTENSION \$199**

Shop around... NO ONE can beat our Annual Lab Test Sale prices!

#### **MALE PANEL**

#### **CARDIAC MARKERS**

Apolipoprotein B
Homocysteine
C-Reactive Protein (high sensitivity)

#### **METABOLIC PROFILE**

Glucose Insulin NEW

Hemoglobin A1c

Kidney function tests: creatinine, BUN, uric acid, BUN/creatinine ratio Liver function tests: AST, ALT, LDH, GGT, bilirubin, alkaline phosphatase Blood minerals: calcium, potassium, phosphorus, sodium, chloride, iron Blood proteins: albumin, globulin, total protein, albumin/globulin ratio

#### **LIPID PROFILE**

**Total Cholesterol** 

LDL (low-density lipoprotein)
HDL (high-density lipoprotein)
Triglycerides

#### **COMPLETE BLOOD COUNT (CBC)**

Red Blood Cell count including: hemoglobin, hematocrit, MCV, MCH, MCHC, RDW White Blood Cell count including: lymphocytes, monocytes, eosinophils, neutrophils, basophils

#### Platelet count

#### **CANCER MARKER**

**PSA (Prostate Specific Antigen)** 

#### **HORMONES**

Free and Total Testosterone
DHEA-S
Estradiol (an estrogen)
TSH (thyroid function)

Vitamin D (25-hydroxyvitamin D)

#### **FEMALE PANEL**

#### **CARDIAC MARKERS**

Apolipoprotein B
Homocysteine
C-Reactive Protein (high sensitivity)

#### **METABOLIC PROFILE**

Glucose Insulin NEW

Hemoglobin A1c

Kidney function tests: creatinine, BUN, uric acid, BUN/creatinine ratio Liver function tests: AST, ALT, LDH, GGT, bilirubin, alkaline phosphatase Blood minerals: calcium, potassium, phosphorus, sodium, chloride, iron Blood proteins: albumin, globulin, total protein, albumin/globulin ratio

#### **LIPID PROFILE**

**Total Cholesterol** 

LDL (low-density lipoprotein)
HDL (high-density lipoprotein)
Triglycerides

#### **COMPLETE BLOOD COUNT (CBC)**

**Red Blood Cell count including:** 

hemoglobin, hematocrit, MCV, MCH, MCHC, RDW

White Blood Cell count including:

lymphocytes, monocytes, eosinophils, neutrophils, basophils

**Platelet count** 

#### **HORMONES**

Progesterone

**Estradiol** (an estrogen)

Free and Total Testosterone

**DHEA-S** 

**TSH** (thyroid function)

Vitamin D (25-hydroxyvitamin D)

#### Lab Test Super Sale • March 25 – June 3, 2019.

#### Retail price: \$400 • Your Price: \$199

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Blood testing services are available only in the continental United States and Anchorage, AK. Not available in Maryland. Restrictions apply for residents of MA, NY, NJ, RI, and PA.

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#### Reference

\* Int Angiol. 2014 Feb;33(1):20-6.

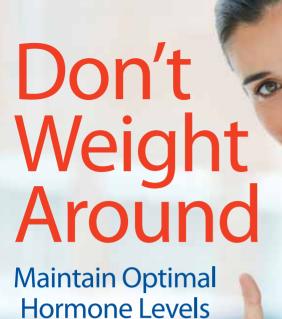
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**7-Keto**<sup>®</sup> is a metabolite of **DHEA** ideal for weight management.

Maintaining youthful DHEA levels can help support immune function, circulatory health, mood, libido, and more.

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NØN GMO

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7-Keto\* brand is a registered trademark of InterHealth N.I. U.S. Patent 7,199,116. Bio-Curcumin\* and BCM-95\* are registered trademarks of Dolcas-Biotech, LLC. U.S. Patent Nos. 7,883,728, 7,736,679 and 7,879,373.

### Risk of Kidney Injury Linked to Common Medications for Acid Reflux

Over-the-counter and prescription medications to treat and prevent symptoms of **heartburn** and acid reflux have been found to have adverse effects, including kidney damage, according to a study in *Scientific Reports*.\*

These medications, called proton pump inhibitors (PPIs), are available under popular brand names such as Prilosec OTC® and Nexium®. Originally developed to treat occasional episodes of heartburn and acid reflux symptoms, PPIs were effective and considered to be safe. But over the years people have continued to use them for prolonged periods.

"Recently, PPI use has come under scrutiny due to growing evidence of renal, cardiovascular, autoimmune, and neurologic adverse effects," the study's authors stated.

The researchers examined more than 10 million records from the FDA's Adverse Event Reporting System, and asserted, "...we provided evidence of kidney injury and electrolyte imbalances in an alarming number of patients taking PPIs."

**Editor's Note:** The authors asserted that, "The observed increased risks of renal and electrolyte adverse effects of PPIs warrant more careful consideration in clinical practice. The risk-benefit ratio should be considered for the individual patient with respect to the adverse effects."

\* Sci Rep. 2019; 9: 2282.



### Heredity Less of a Factor in Longevity

Estimates of the heritability of longevity in humans have been higher than warranted, according to an article published in the journal *Genetics*.\* Heritability is a measure of how much of a variation in a trait can be explained by genetic differences as opposed to lifestyle and other factors.

Using data obtained from a genealogy company, the researchers estimated heritability by examining the similarity of lifespan among relatives. In addition, similarities were also observed between people who were related only by marriage and did not share households.

The researchers found that heritability of lifespan was no more than **7%**, in contrast with previous estimates of up to **30%**. It was determined that past estimates had failed to account for the tendency of humans to select partners with traits similar to their own, a process known as assortative mating.

Editor's Note: "What assortative mating means here is that the factors that are important for life-span tend to be very similar between mates," said lead author Dr. J. Graham Ruby. "When we failed to take assortative mating into account, our own nominal estimates were similar to those of the literature."

\* Genetics. 2018 Nov;210(3):1109-1124.

460166



# CoQ10 Benefits Patients with Chronic Kidney Disease

Supplementation with the antioxidant **coenzyme Q10** (CoQ10) provided several metabolic benefits for patients with chronic kidney disease, according to a review and meta-analysis published in the journal *Current Pharmaceutical Design*.\*

Researchers selected seven randomized, controlled trials that included a total of 384 men and women with chronic kidney disease. Metabolic profiles evaluated during the trials included measurements of triglycerides, total cholesterol, low density lipoprotein (LDL) cholesterol, high-density lipoprotein (HDL) cholesterol, glucose, insulin, insulin resistance, C-reactive protein, malondialdehyde (MDA), a marker of oxidative stress, and creatinine (which is elevated in kidney disease).

The daily dosage of **CoQ10** in the trials ranged from **30 mg** to **200 mg** administered for periods of **4** to **12 weeks**. Pooled analysis of the subjects determined that CoQ10 supplementation was associated with a decrease in total cholesterol, LDL-cholesterol, malondialdehyde and creatinine (a standardized mean difference of -1.65, meaning that **CoQ10** improved creatinine levels in those with *chronic kidney disease*).

If **creatinine** is elevated above **1 mg/dL**, this serves as an indictor of kidney impairment. (The standard reference range for <u>elevated</u> **creatinine** starts at **1.27 mg/dL**, but **Life Extension**® views any level above 1.00 as an indicator to cut back on pain relieving (NSAID) drugs, reduce consumption of meat, or look at other risk factors such as hypertension and diabetes.)

Blood levels of **creatinine** are included in most **CBC/ Chemistry blood test** panels.

**CBC/Chemistry** tests are included in the comprehensive **Male** and **Female Panels** many readers of this magazine have done annually.

A far more accurate **blood test** to assess **kidney function** is called **cystatin-C**.

To order any of these **blood tests** call 1-800-208-3444 or visit www.LifeExtension.com/blood.

**Editor's Note:** Circulating concentrations of CoQ10 have been decreased in patients with chronic kidney disease, suggesting that CoQ10 supplementation may be an ideal antioxidant therapy for these patients, the authors noted.

\* Curr Pharm Des. 2018;24(31):3710-3723.

# HEALTHY FLORA



One of the most exciting fields of scientific research is focused on the trillions of bacteria that live in our bodies.

Researchers have discovered that these **flora** play an enormous role in immunity as well as overall health.

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FLORASSIST



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#### **FLORASSIST® Immune Health**

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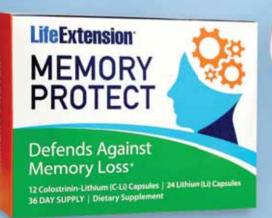
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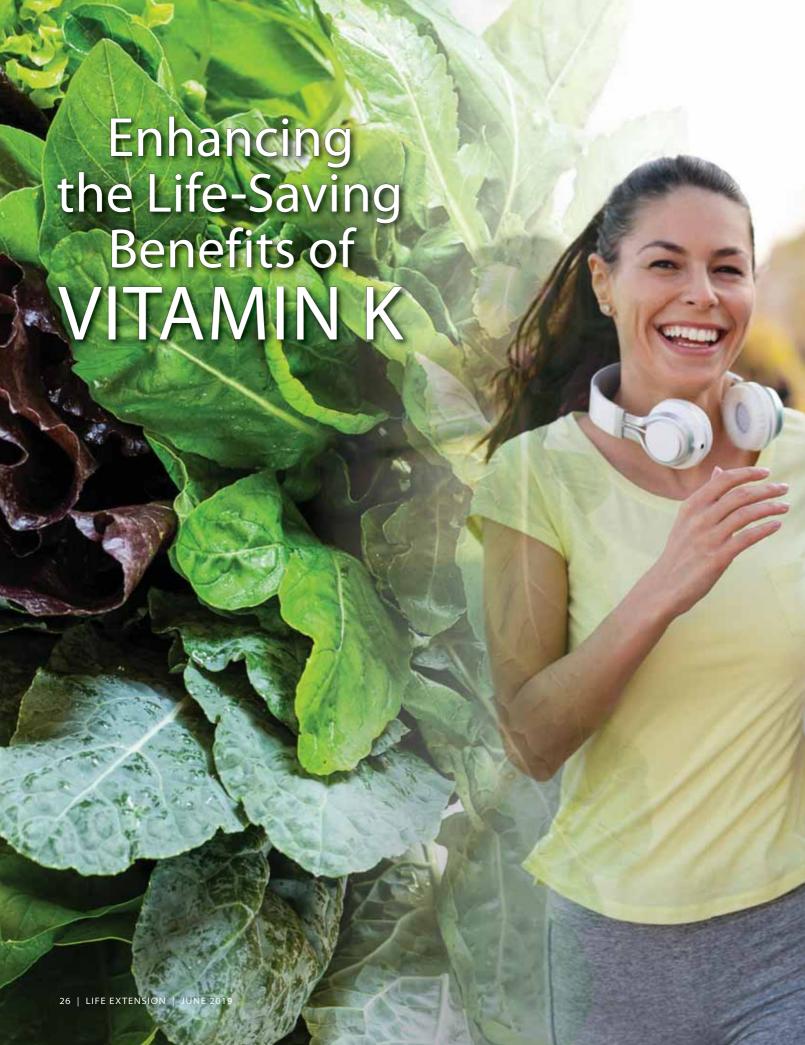
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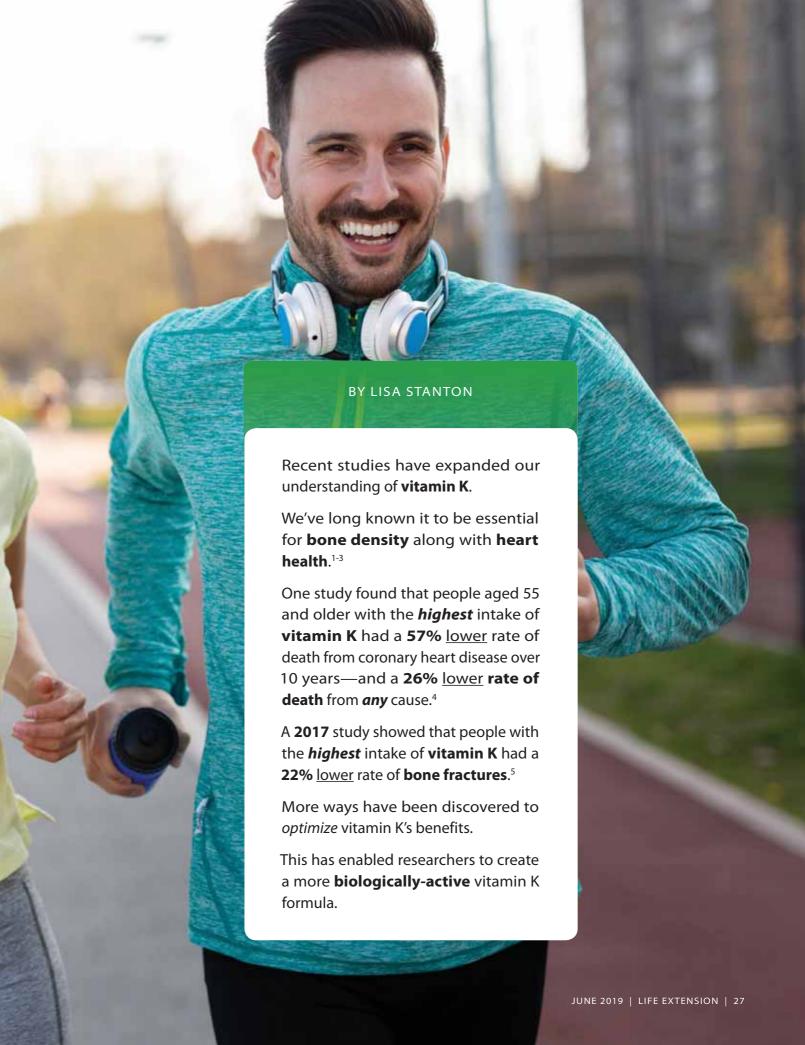
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#### Vitamin K in Food

Scientists have identified different forms and amounts of **vitamin K** that are found in foods.

This work provides crucial insights into the types of vitamin K we should be consuming.

It also reveals limitations in relying on food to supply the types and levels of vitamin K associated with peak bone and cardiovascular health.

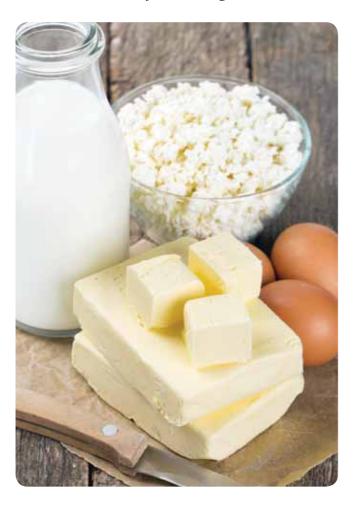
#### Forms of Vitamin K

Vitamin K in food comes in two general forms:

- 1) Vitamin K1 (or phylloquinone), found primarily in leafy green vegetables.
- 2) Vitamin K2, found in natto or fermented soy, and in animal products such as eggs, meat, milk, and cheese.6

Vitamin K2 has several subtypes denoted by a number, such as MK-4, MK-6, and MK-7.

**MK** stands for **menaquinones**, which are forms of vitamin K that vary in their organic structure.



One epidemiological study showed that some forms of menaquinone, such as MK-7 and MK-9, appear to be most powerful in preventing coronary heart disease.7

**MK7** is a long-acting form of vitamin K that has been available as a dietary supplement for many years. **MK-9** has only recently become available in supplement form.

#### Limits to Dietary Intake of Vitamin K

Studies show that there are problems in relying on diet to supply all these forms of vitamin K.

Vitamin K1 in foods has low bioavailability.8 This means that even if you eat a large amount of leafy green vegetables, you may not absorb adequate amounts of vitamin K1.

Vitamin K2 is found in highest concentrations in many foods that people try to limit, such as foods high in saturated fat.9 One would have to consume massive amounts of cheese to achieve the optimal K2 levels that are supported by human clinical trials. And low-fat versions of these foods often have much less vitamin K2 content, or even none at all.9

Supplementation is a more efficient and practical way of increasing vitamin K intake. But what are the optimal doses of the various types of vitamin K that we should be consuming daily?

#### **New Studies, New Dosing**

Several recent clinical trials in humans have demonstrated that adequate vitamin K supplementation can have a significant impact on bone and cardiovascular health.

#### **Vitamin K1 Dosing**

A one-year, randomized, controlled trial evaluated vitamin K1 supplementation in adults with calcifi**cation** in their **aortic valve**, one of four valves that regulate blood flow through the heart.<sup>10</sup> Degenerative heart valve disease such as this can lead to impaired heart function and heart failure if it progresses.<sup>11</sup>

Subjects received either 2,000 mcg of vitamin K1 daily or a placebo. Those who received vitamin K1 benefited from a reduction in the progression of aortic valve **calcification**, based on both imaging and biochemical markers. Disease progression was cut by more than half by vitamin K1 supplementation.<sup>10</sup>

It's a remarkable result. One would have to eat almost 14 cups of spinach a day to achieve this level of vitamin K1 intake.12

What You Need to Know

**Aortic stenosis** is a narrowing of the heart's aortic valve. Some people have this problem due to a congenital defect (i.e. bicuspid valve), but most develop it as a result of aging, with calcification and/or fibrotic scarring of the valve.

Age-related aortic stenosis usually starts in the sixth decade of life, and early treatment can reduce the risk of progression of the narrowing, which can lead to heart failure if left untreated.

The only definitive treatment is surgery, either valve repair or valve replacement. An exciting new approach called **transcatheter aortic valve replacement (TAVR)** can allow valve replacement without the need for open-heart surgery. However, **TAVR** may pose a greater risk of stroke in some patients.

#### Vitamin K2 Dosing: MK-4

Research has helped establish dosing levels that maximize the benefits of the MK-4 form of vitamin K2 on bone health.

Two recent trials used a dose of **1,500 mcg** of MK-4 per day in healthy, postmenopausal women, and compared them to a group receiving a placebo. <sup>13,14</sup> Postmenopausal women are at high risk for developing loss of bone density, osteoporosis, and subsequent risk for bone fractures. <sup>15</sup>

Both studies demonstrated a significant improvement in bone density in the group receiving vitamin K2. They had higher levels of the active form of **osteo-calcin**, indicating more new bone formation.<sup>13,14</sup>

In one of the studies, subjects receiving MK-4 also had a stabilization of their forearm **bone mineral density**, while those in the placebo group continued to show progressive *loss* of bone density over the year of the study.<sup>14</sup>

#### Vitamin K2 Dosing: MK-7

Another group of researchers followed 244 healthy postmenopausal women for three years and examined the effects of the **MK-7** form of vitamin K2 on both bone and cardiovascular health. The results of their findings were published in two separate publications.<sup>16,17</sup>

Participants took either **180 mcg** of MK-7 daily or a placebo. Again, researchers found improvements in markers of both bone *and* blood vessel health in the supplemented group. <sup>16,17</sup>

Additionally, the ratio of inactive to active osteocalcin improved by **58**%. (Only <u>active</u> osteocalcin helps to synthesize bone.) Furthermore, bone mineral content, bone mineral density, and bone strength were all significantly improved in the group taking MK-7.<sup>17</sup>



#### Broad-Spectrum Vitamin K

- Vitamin K is an essential vitamin found in two general forms, K1 and K2.
- Research has revealed that vitamin K is important not only for blood clotting but also for bone and cardiovascular health.
- Research has also helped clarify the ideal doses of vitamin K1 and K2 needed to optimize bone and heart health.
- Due to poor bioavailability and high fat content of foods containing the most vitamin K, obtaining adequate vitamin K in the diet is problematic.

#### Vitamin K and Clotting: What You Need to Know

Although vitamin K is critical to the normal process of blood clotting, supplementation with vitamin K is not associated with any increased risk of abnormal, harmful clotting, such as that associated with heart attack and stroke.

Even studies with very high doses of vitamin K have demonstrated its safety, without any adverse events. 10,22,23

But there is one extremely important caveat to consider when starting a vitamin K supplement.

Powerful anticoagulant medications such as warfarin (brand name Coumadin®) and other related drugs act by blocking vitamin-K-dependent pathways, decreasing the ability of the body to produce several important blood clotting factors. These drugs are used in patients who are at high risk of dangerous, abnormal clotting. For example, they may be used in patients being treated for atrial fibrillation, heart valve disease, deep vein thrombosis, and/or pulmonary embolism.

Vitamin K acts as an antagonist to the anti-clotting effect of warfarin and similar drugs. Therefore, taking higher levels of vitamin K can interfere with the desired clinical effect and increase the chance of clotting in these patients.

Patients taking warfarin (Coumadin®) or related medications should consult their prescribing **doctor** before taking any vitamin K supplement.

But vitamin K does not in any way interact with newer/novel, oral anticoagulants like Xarelto® (rivaroxaban), Pradaxa® (dabigatran), or Eliquis® (apixaban). These newer, oral anticoagulants work by inhibiting venous clotting through thrombin or Factor Xa, independent of vitamin K. Vitamin K supplementation can be used safely with these drugs since there is no potential for interaction. In addition, there are available antidotes to bleeding with the newer drugs. Dangerous or life-threatening bleeding with Pradaxa® can be reversed by Praxbind® (idarucizumab). Recently (May, 2018), Andexxa® (andexanet alfa) was approved to reverse life-threatening bleeding with Xarelto® and Eliquis®.



In terms of blood vessel health, vitamin K2 supplementation helped to preserve the flexibility of arteries by reducing arterial stiffness. While those receiving a placebo had worsening arterial stiffness, those taking vitamin K not only preserved arterial flexibility, but decreased stiffness by about 6%.16

Those individuals with more advanced arterial stiffness at the start of the study improved the most. 16 This indicates that vitamin K not only preserves blood vessel health, it also appears to be able to help *reverse* existing blood vessel disease.

#### **Novel Vitamin K2 Subtypes**

Food sources of vitamin K2 provide an assortment of other MK forms as well, including **MK-6** and **MK-9**. Although clinical trials have not yet been performed for most of these forms, observational studies suggest many benefits.

For example, a study followed a group of more than 16,000 individuals to evaluate the impact of intake of specific vitamin K forms on risk of coronary heart disease.7 It found that those who consumed higher levels of vitamin K reduced their risk of heart disease. Much of this effect could be attributed to the longer MK forms, such as MK-7 and MK-9.

The study also found that the risk of coronary heart disease was reduced by 9% for every additional 10 mcg of vitamin K2 consumed per day in individuals aged 49 to 70.7

#### Vitamin K's Role in Bones and Blood Vessels

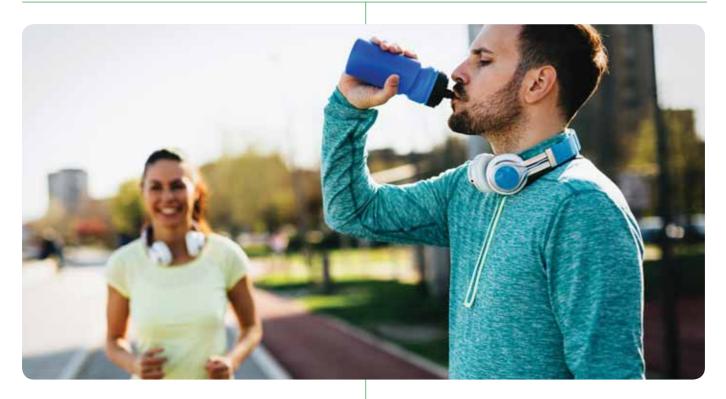
The effect of vitamin K on the production of **blood clotting** factors crucial for normal clotting has been well understood. But research has just recently revealed its impact on bones and blood vessels.

Several proteins have been discovered in both bone and blood vessels that are **vitamin K-dependent**. This means that they require adequate levels of vitamin K to function.<sup>2,3</sup>

Vitamin K is essential for producing active **osteo**calcin, a bone hormone involved in new bone formation and often used as a biochemical marker of overall skeletal health.

Vitamin K supports the deposition of **calcium** in

In blood vessels vitamin K has the opposite effect of helping to prevent excess calcium deposition. Calcification in arteries is common in older age and is associated with arterial stiffening, atherosclerotic plaque, and increased risk for heart and kidney disease.3



#### Vitamin K and **Cardiovascular Disease**

Several other studies have shown benefits of vitamin K for cardiovascular health.<sup>2-4,7,18</sup>

A study of vitamin K intake in 564 postmenopausal women found that higher vitamin K2 levels were associated with protection from coronary artery calcification. 18 Those with the *highest* intake had a **20**% lower rate of calcification than those with the lowest intake.

And in a study that followed 4,807 adults aged 55 and older, for up to 10 years,4 several negative cardiovascular outcomes were shown to be less common in those subjects with the *highest* intake of vitamin K2, compared to those with the lowest intake.

The rate of new diagnoses of coronary heart disease during follow-up was 41% lower in those with the *highest* intake of vitamin K2. Most dramatically, death due to coronary heart disease was 57% lower in those with the highest intake, and **death by** any cause was 26% lower.4

#### **Reduced Fracture Risk**

Several other studies have found that various forms of vitamin K supplementation improve the osteocalcin status of participants, an important marker of new bone formation and overall bone health.<sup>13,14,17,19-22</sup>

But it is important to ask whether the **rate of bone fractures** is reduced with increased vitamin K dietary intake.

A study published in the journal *Medicine* in **2017** investigated just that question.5 Researchers performed an extensive meta-analysis, pooling data from close to 81,000 individuals. Overall, they found there was a 22% lower rate of fractures in those individuals with the highest intake of vitamin K, confirming a protective effect of vitamin K against fractures.

In addition, the scientists found a dose-response relationship, with a 3% lower rate of fracture for every **50 mcg** of vitamin K consumed per day.<sup>5</sup>

#### **Summary**

Vitamin K is an essential nutrient that is being recognized for more health benefits than just aiding blood clotting.

Scientific research demonstrates that adequate intake of vitamin K is crucial for optimal bone, heart valve, and blood vessel health.

New studies reveal the importance of new **forms** of vitamin K that are associated with reduced risk of age-related outcomes.

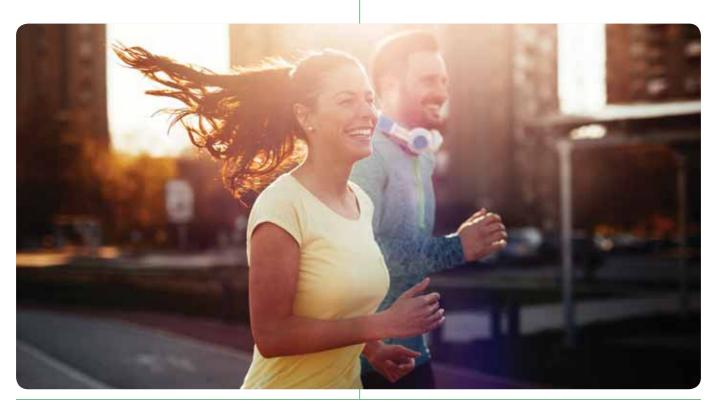
Using this knowledge, scientists have created a broad-spectrum vitamin K formula, with beneficial MK-6 and MK-9 compounds, along with K1, MK-4, and MK-7.

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

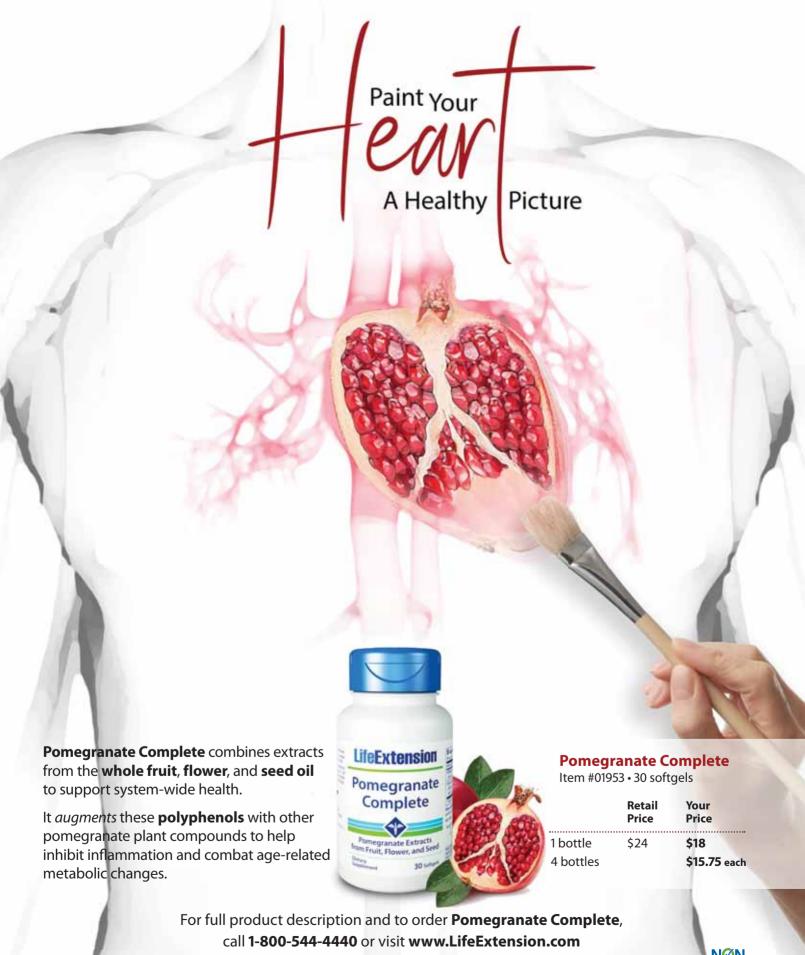
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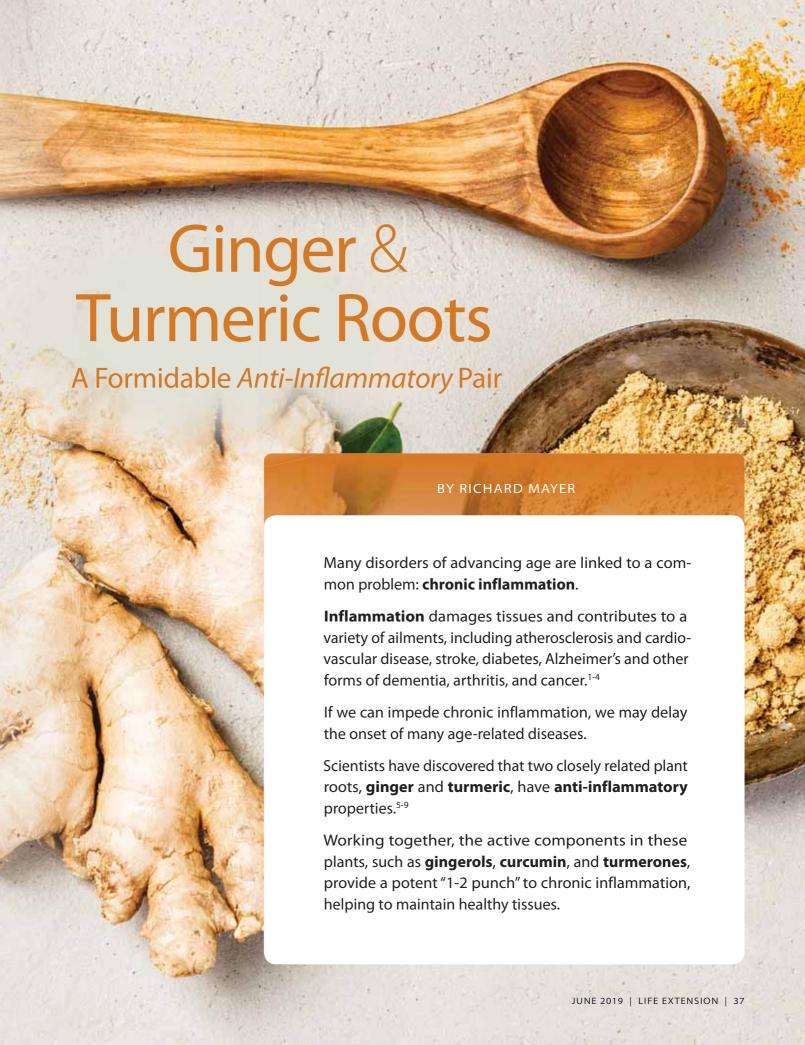


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### **Inflammation Contributes to Aging**

Chronic, low-grade inflammation doesn't just increase the risk of disease—it has been identified as one of the key contributors to the aging process itself. 1,3,4,10,11

Inflammation plays such a central role in agerelated ailments that scientists have coined the term **inflammaging** to describe its effects.<sup>1</sup>

There are many different causes of this type of chronic inflammation.

With advancing age, many cells become **senescent**. That means they no longer divide or support tissues. Senescent cells also spew out harmful compounds that activate inflammation and cause tissues to age more rapidly.<sup>4</sup>

In overweight or obese individuals, excess fat cells release compounds that promote inflammation throughout the body.<sup>1,3</sup>

Other contributors to inflammation include diet, gut microorganisms, pollution, hormone imbalances, stress, and more. 1,3,4,10

### **Plant-Based Solutions to Inflammation**

For centuries, **ginger** and **turmeric** have been recognized as powerful anti-inflammatory plants. They both contain beneficial compounds that reduce inflammation and offer other health benefits.

They also have antiviral and antibacterial properties that help to bolster the immune system's defense against pathogens. Ginger and turmeric have also demonstrated anti-cancer properties.<sup>12,13</sup>

### Ginger: An Anti-Inflammatory Powerhouse

Ginger is a flowering plant whose underground stem or rhizome, also known as **ginger root**, has been used in cooking and as a natural medicine.

**Ginger root** is the source of a cocktail of beneficial compounds, including **gingerols**, that have free-radical scavenging and anti-inflammatory properties.<sup>14</sup>

**Leukotrienes** contribute to dangerous inflammatory reactions. Compounds in ginger have been found to *inhibit* enzymes that produce **leukotrienes**. <sup>5,6</sup>

Ginger also works to reduce inflammation by affecting several other compounds and cellular pathways, including:

- Reduction in levels of inflammatory mediators, such as tumor necrosis factor-alpha,
   C-reactive protein, interleukin-1β and interleukin-6. 15-20
- Inhibition of **nuclear factor kappa B** (**NF-kB**) pathways, which are involved in the *initiation* of chronic inflammation. <sup>16,21,22</sup>

### Turmeric: A Golden Anti-Inflammatory

Turmeric, a close relative of the ginger plant, has a bright yellow-orange rhizome. Like ginger, turmeric is a source of both spices and medicine.

**Curcumin** and other **curcuminoids** are compounds derived from turmeric that have been the most-studied. There is a robust research record

showing their efficacy in fighting inflammation. Most extracts of turmeric on the market primarily contain isolated curcumin.

Other compounds found in turmeric, such as turmerones, offer their own health benefits, many of which help to augment the effects of curcumin.<sup>23</sup>

Turmeric's anti-inflammatory effects are, like ginger's, inhibition of leukotrienes and other proinflammatory pathways, as discussed on the previous page. Turmeric has additional effects that go beyond what ginger can do, including:

- Support of **sirtuin 1** expression, which is associated with healthy aging and longevity,<sup>24</sup>
- Neuroprotective capabilities, by reducing neuroinflammation and by protecting against the toxic effects of **beta-amyloid** accumulation associated with Alzheimer's disease, and<sup>25-27</sup>
- Activation of **heme oxygenase-1** and other pathways of protection from oxidative damage.24,28

In addition, **STAT3** signaling, which is associated with inflammation and development of some cancers, has been shown to be suppressed by turmeric.<sup>29,30</sup>

### **Specific Conditions for which Ginger and Turmeric are Beneficial**

Although turmeric and ginger contain different compounds, they offer overlapping anti-inflammatory properties and complementary effects. In other words, they complement each other.

Together, these two plant roots and their extracts offer superlative anti-inflammatory effects. Scientific studies have shown that ginger and turmeric can help fight a wide range of age-related disorders and conditions that include practically every system and organ in the body.

The following are a few of the conditions for which extracts of ginger and turmeric can be beneficial.

### **Metabolic Disorders**

Obesity, diabetes, and metabolic syndrome are increasingly common and are risk factors for many other problems, such as cardiovascular disease.31,32 Inflammation is strongly associated with these disorders, and is a primary driver of the health problems to which they lead. 33

Research shows that both ginger and turmeric help alleviate many of the metabolic abnormalities that characterize these disorders. 15,18,22,34-39

### Curcumin and Ginger Fight Inflammation

- Chronic inflammation increases throughout the body as we age.
- Many sources contribute to this harmful inflammation, which does damage to our organs and tissues.
- This long-term damage builds up to cause age-related disease and loss of function, contributing to the development of almost all age-related disorders, from cardiovascular disease to cancer.
- Natural compounds in ginger and turmeric work by means of several different complementary mechanisms to deliver powerful anti-inflammatory effects.



Mice fed high-fat and/or high refined-carbohydrate diets normally suffer from dangerous changes in body composition and metabolism, including obesity, abnormal blood lipid levels, and systemic inflammation. Compounds in ginger prevent these changes, by helping to maintain healthy weight and lipid levels, and reducing inflammation.34,35

In animal studies, ginger also provided protection to organs (kidneys and brain) commonly affected in diabetics.36,37

Human studies show that supplementation with ginger and turmeric can reverse many of the metabolic abnormalities seen in prediabetics. 15,18,22,39 Daily supplementation with ginger powder helps to control blood sugar levels, triglycerides, and LDL ("bad") cholesterol, while raising healthy HDL cholesterol and improving insulin sensitivity. 15,18,22

### Cardiovascular Disease and Stroke

The abnormalities associated with metabolic disease and diabetes are strong risk factors for blood vessel disease that can lead to heart disease, stroke, and other related conditions.

Preclinical studies show that both ginger and turmeric reduce the risk for these diseases by improving glucose and lipid control. They also contribute to blood vessel health by blocking inflammation that leads to **atherosclerosis** (hardening and narrowing of the arteries).24,28,40-42

Studies in animal models that include ginger and turmeric have shown that both compounds reduce vascular inflammation and protect blood vessels from the damage that high blood pressure and abnormal cholesterol levels can cause. 40,41 This suggests a reduction in the risk for atherosclerosis and its dangerous consequences, including heart attacks and strokes.

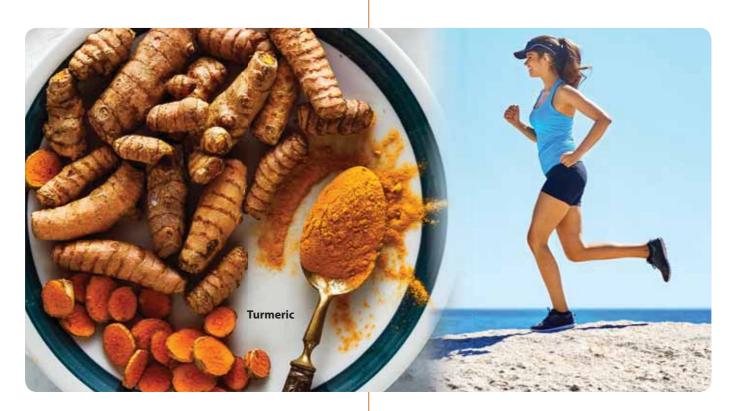
### **Osteoarthritis**

A common cause of loss of mobility in older age, **osteoarthritis** occurs when the protective cartilage on the ends of the bones wears down. Human studies have now documented the ability of curcumin and ginger to help reduce its symptoms. 43,44

Improvement of several clinical markers of arthritic disease has been observed, including lower levels of pro-inflammatory compounds, improved physical function, and better quality of life. 19,43,44

Curcumin and ginger have also been shown to be beneficial for rheumatoid arthritis, an autoimmune form of arthritis that starts when the immune system targets the joint linings.45-47

One study assessed treatment with a combination of ginger and turmeric for rheumatoid arthritis in a rat model. The ginger-turmeric combination was found to help preserve the structure of joints, reduce inflammation, and prevent many other changes that can occur in rheumatoid arthritis such as impaired kidney function and anemia.46



### Alzheimer's, Parkinson's, **Cognitive Decline and Dementia**

Turmeric and curcumin are recognized for their ability to protect against many of the age-related changes that occur in the brain and contribute to loss of cognitive function. They protect against biochemical pathways of neuroinflammation, which are involved in the development of neurodegenerative diseases like Alzheimer's and Parkinson's disease. In preclinical models they reduce the damage done by abnormal proteins like beta-amyloid.<sup>26,27,48</sup>

Ginger has similar benefits. Not only do compounds found in ginger reduce brain damage caused by amyloid abnormality and inflammation, they have been found to inhibit acetylcholinesterase, an enzyme that breaks down the neurotransmitter acetylcholine. 49,50 This helps boost brain levels of acetylcholine, which is essential for processing memory and learning. Most pharmaceutical drugs used to treat the symptoms of Alzheimer's disease do just this.

Taken together, the combination of ginger and curcumin may help both *prevent* the progression of these common brain diseases and reduce their symptoms.

#### Cancer

Inflammation is a major contributor to the formation of most cancers. By fighting inflammation along with other mechanisms, ginger and curcumin have been shown in many studies to help inhibit the growth of tumors. Together they promise to be significant weapons in the fight against cancer.<sup>2,14,29,51-56</sup>

### **Summary**

Chronic, low-grade inflammation has many causes. It is a major contributor to the loss of function and risk for disease that become more prevalent as we age.

In fact, almost all age-related changes and disorders can be tied to inflammation, from arthritis and diabetes to dementia and cancer.

Natural compounds found in the rhizomes of two related plants, ginger and turmeric, have demonstrated anti-inflammatory properties. They act by means of multiple, complementary mechanisms to reduce harmful inflammation while protecting normal immune function.

Experimental, animal, and human studies have demonstrated significant benefits. •

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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# CoQ10 Protects Skin Mitochondria from Aging

BY KATHY WISE

As we age, we tend to tire more quickly. This loss of energy is reflected not only in how we feel but how we look.

When cells are unable to maintain and repair themselves properly, it leads to skin wrinkles, sagging, and other outward signs of age.<sup>1,2</sup>

A new study<sup>3</sup> shows that we can **fight skin aging** by using a familiar and vital nutrient:

Coenzyme Q10

### The Role of Mitochondria

According to the *mitochondrial theory of* aging, energy loss is a causative factor in degenerative aging.4

**Mitochondria** are the tiny power plants of our cells. They burn food molecules and generate adenosine triphosphate (ATP), which delivers energy to the cells.

Over time, the mitochondria are damaged.<sup>3-6</sup> There are many causes, including oxidative stress, toxins, and infections. And some of the oxidative stress comes directly from the mitochondria's own energy-generating activity.7

Damaged mitochondria are less efficient and produce less ATP from food. That leaves us with an energy deficit.

In our skin, insufficient ATP means inadequate cell maintenance and repair. One result is visibly wrinkled and sagging skin.<sup>1,2</sup>

Studies showed long ago how CoO10 can heal mitochondria and boost mitochondrial function.8-10 Scientists at a top university in Germany set out to examine how this would impact aging skin.

### **Testing Aging Skin for** CoO10 Benefits

Researchers took skin biopsy samples from donors of varying ages to measure the effects of mitochondrial function on skin aging.3

They kept the skin samples alive in cell culture medium and tested each one for mitochondrial function, including oxygen consumption rate (more consumption

means more mitochondrial activity) and ATP production (more ATP means more active mitochondria).

Next, they supplemented the culture medium of the biopsies with either CoO10 or a control solution and measured the impact on mitochondrial function. Oxygen consumption and ATP production were again measured.

### CoO10 **Restores Energy**

Two major findings emerged. First, researchers demonstrated the mitochondrial theory of **aging**—and showed that it applies to human skin-by demonstrating that oxygen consumption and ATP production declined with age.3

Samples from older people had lower rates of mitochondrial activity and, as a result, lower rates of energy production. In fact, the scientists found a reliable and severe decline in mitochondrial activity of about 10% per decade.3

Second, the study revealed a role for CoQ10 in the prevention of age-associated mitochondrial dysfunction.

The researchers found that when CoO10 was added to the skin biopsies, there was a marked increase in mitochondrial activity. This was measured by examining the oxygen consumption rate, since mitochondria use oxygen to generate ATP.

Skin biopsies supplemented with CoQ10 ramped up their oxygen consumption rate by close to 30% compared with the unsupplemented ones.3

The study indicates that supplementation with CoQ10 can significantly restore mitochon**drial energy** production. This has important implications for reducing skin aging. It shows that by restoring mitochondrial energy production, skin tissue repair and **maintenance** are improved. This gives CoQ10-nourished cells a boost in fighting the ravages of aging.

This finding is important for those concerned about skin aging. It has implications for other organs and systems, all of which rely on mitochondria to function properly.

### **Human Study Shows** CoQ10 Helps Combat Visible Skin Aging

Researchers at the Institute of Cosmetics in Liubliana, Slovenia, set out to examine the potential benefits of CoO10 on visible signs of skin aging as measured by skin thickness, hydration, elasticity, and wrinkles. The study was done during the cold winter months when skin problems tend to be most evident.11



Thirty-three women aged 45-60 were enrolled and told to continue their routine skin-care and dietary habits.

This group was subdivided randomly to receive one of the following:

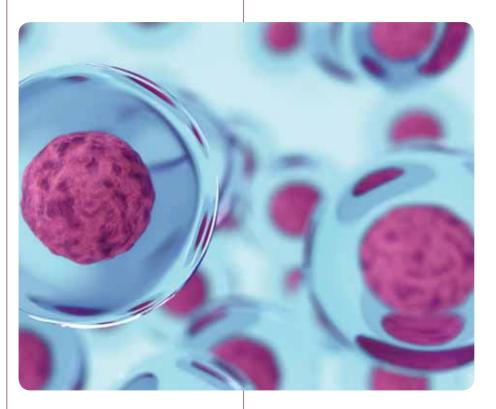
- Placebo.
- **50 mg** each day CoO10 supplement, or
- **150 mg** each day CoQ10 supplement.

The study lasted 12 weeks, and skin quality measurements were conducted at baseline, and at six and 12 weeks of supplementation.

### The study showed that:11

- Wrinkles (measured in the area around the eyes) did not change in the placebo group, while both CoQ10supplemented groups had significant *reductions in* wrinkles.
- **Skin elasticity** decreased by nearly 25% over 12 weeks in placebo subjects, but remained unchanged in both CoQ10 groups, resulting in significant differences between treated and placebo subjects.
- **Smoothness:** Expert evaluation (blinded to study group) revealed significant improvement in CoQ10-treated groups' **skin smoothness** by 70%-82%, with no significant changes in the skin of placebo recipients.

This human study is among the first to demonstrate the clinical benefits of oral CoQ10 supplements for reducing **skin aging**.



### How It Works and Why It Matters

Cells require efficient, cleanrunning mitochondria to function properly-and when mitochondria suffer, so do the tissues in which they reside.

At the core of the energy-converting process is the *electron* transport chain, a sort of power line that transfers electrons from food-derived molecules (mainly carbohydrates and fat) to ATP molecules. Cells then use ATP to power themselves.

As this process occurs, toxic byproducts are produced, including free radicals.6,12-14 The resulting oxidative damage causes the mitochondria to gradually deteriorate. In skin tissue, that leads to visible defects in skin quality. 15-17

CoQ10 plays an essential role in the electron transport chain, facilitating the efficient transfer of

electrons. It also acts as a free radical scavenger, reducing oxidative damage.

But CoQ10 levels drop with advancing age. This leaves cells relatively unprotected, and causes mitochondria to burn their fuel inefficiently, wasting energy and producing more toxic byproducts.6,12-14

In the skin biopsy study, adding CoQ10 acted as a "fuel additive" that enhanced electron transport to overcome the consequences of ailing mitochondria.3

What all this means is:

- a) CoQ10 supplementation may be an important step in maintaining healthy skin into advanced age, and
- b) Other tissues in the body are likely to receive similar benefits from CoQ10.

### **Summary**

One aspect of aging involves accumulated cellular damage and dysfunction.

One of the biggest contributors is the increase of **dysfunctional mitochondria** in our cells.

A study shows that skin cells supplemented with CoQ10 have healthier mitochondria. Adding to this is evidence from a **clinical trial** showing oral CoQ10 supplementation improved the appearance and youthfulness of aging skin.

These findings suggest that CoQ10 supports youthful function in tissues throughout the body. •

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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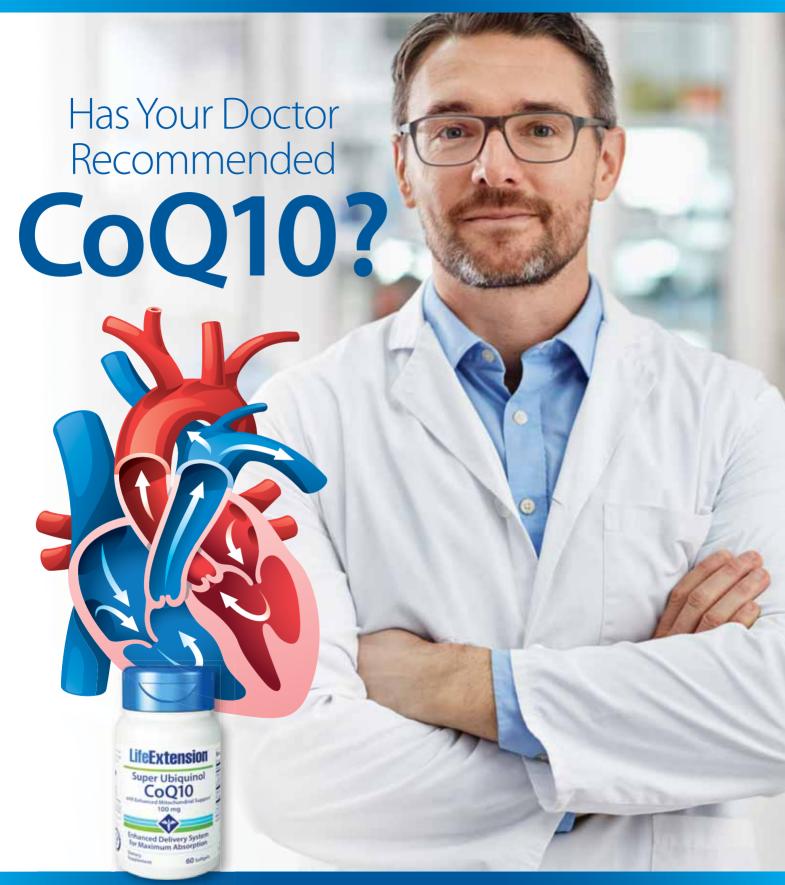
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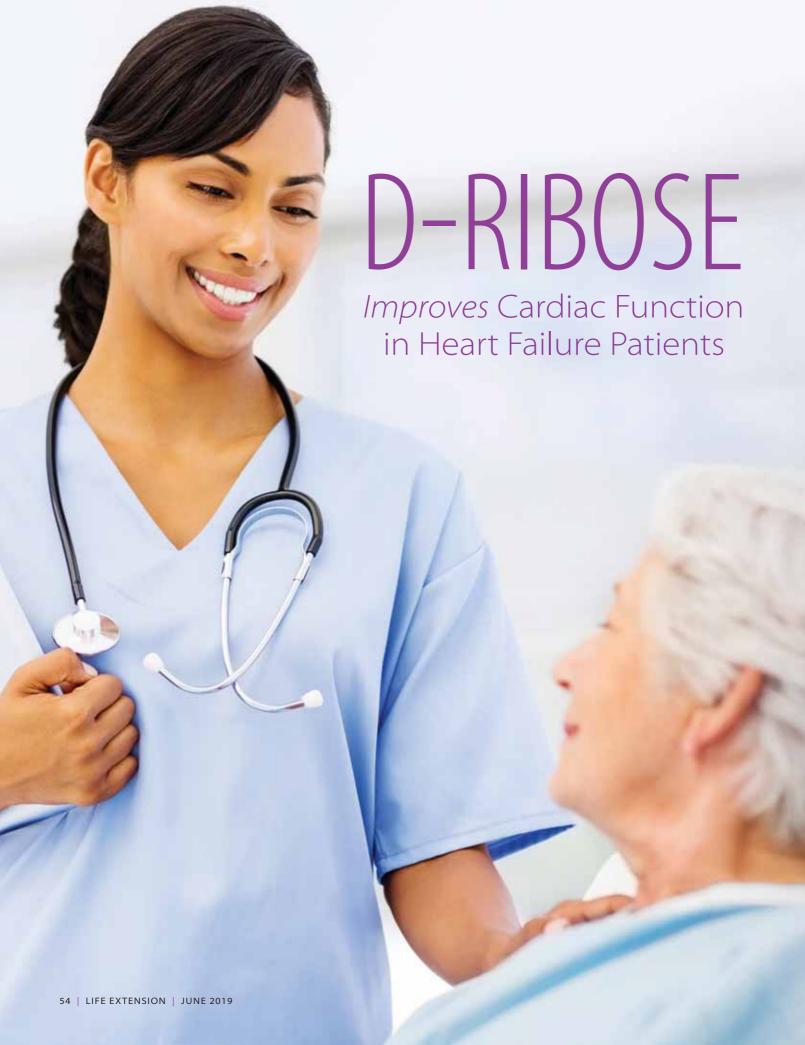
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### BY JULIE STEIN

When it comes to heart failure, the statistics are chilling.

More than 870,000 new cases are diagnosed in the U.S. each year in people aged 55 and older.1-4

One in five people who are currently 40 years old will go on to develop heart failure in their lifetimes. By 2030 it is expected that more than eight million Americans will be affected by the disease.1-3

**Heart failure** occurs when the heart fails to pump enough blood to support the whole body. 1 It is a debilitating and progressive condition.

Clinical trials have shown that supplementation with **D-ribose**, which is produced in the body, can **improve heart function** following damage to the heart muscle.

It helps the heart muscle manage energy, offering a potential solution to the slow-motion cardiac energy crisis that is heart failure.5-7

But how does **D-ribose** achieve these results?

The answer has to do with the way energy is released and converted into adenosine triphosphate (ATP). This universal "energy currency" then transfers energy to cells in a form they can use to do their cellular work.

Here, we review the compelling clinical studies that reveal the power of D-ribose in heart failure, and then explain precisely how it works to feed the energy-hungry, failing heart.

### **D-Ribose Improves Heart Function**

D-ribose is a building block of *ATP*, 8 the energy source for every cell. Research has demonstrated that D-ribose supplementation can support the heart's ability to extract energy and store it in the form of ATP, improving the function of the heart even if it's been damaged.

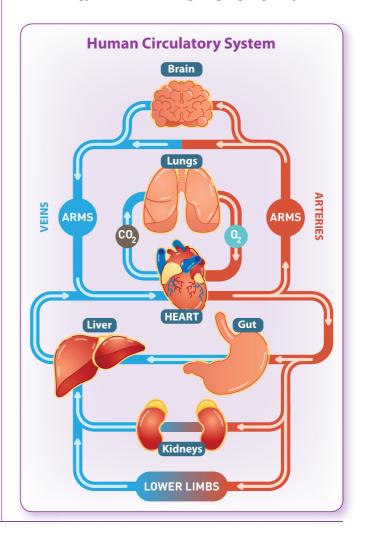
Human studies show the following potential of D-ribose to reduce the symptoms of heart failure when given orally at a dose of **5 grams three times daily**:

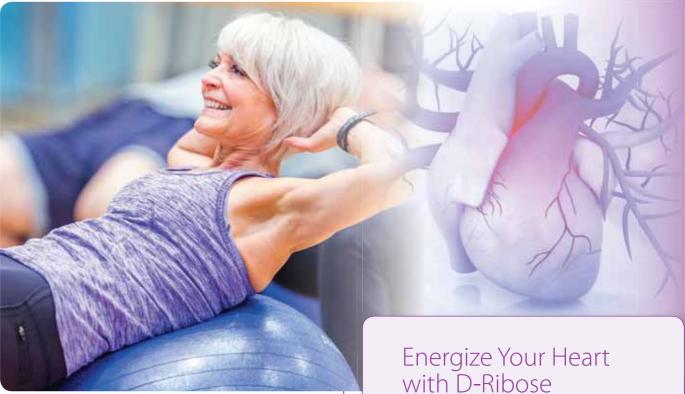
- Improved blood flow through the heart. Heart failure results in slower flow of blood through the heart, raising the amount of work the heart muscle must do to push the blood through the body. Fifteen patients with chronic coronary artery disease and heart failure were randomly assigned to receive **D-ribose** or a **placebo** for three weeks, and after a one-week "washout" period, were switched to the opposite therapy. During the D-ribose supplement phases subjects had significantly enhanced filling of the left ventricle and other signs of reduced cardiac loading that *eased* the workload on the ailing heart muscle, and significantly improved measures of quality of life. No such changes were seen during placebo phases.
- Improved blood flow in the body. Heart failure reduces the amount of blood the heart can pump with each beat, slowing down the body's whole circulation, imperiling organs that need ample blood supply. Eleven patients with New York Heart Association Class II-IV (mild to severe) heart failure were supplemented with D-ribose for six weeks.<sup>5</sup> At the end of the six-week supplementation period, **improved blood flow** in tissues was shown by Doppler ultrasound in 64% of patients. This improvement lasted through nine weeksthree weeks *after* the end of supplementation.
- Improved exchange of oxygen and CO, through the lungs. Heart failure increases how hard the heart-lung system must work to rid the body of carbon dioxide and take in oxygen. Sixteen people with New York Heart Association Class III-IV heart failure (among the most severe) were treated in a trial of D-ribose for eight weeks.<sup>6</sup> All patients had a significant improvement in their breathing parameters, which are so often impaired in heart failure.

D-ribose supplementation significantly increased ventilatory efficiency (how much air needs to be moved to eliminate metabolicallyproduced carbon dioxide from the body) by 16%. Ventilatory efficiency is a strong predictor of survival in heart failure patients, so this demonstrates a direct reduction in the risk of dying.6 Measures of oxygen uptake efficiency, oxygen flow through the lungs, and amount of oxygen pumped per heartbeat also improved, by 25%, 19%, and 16%, respectively, demonstrating marked increases in the ability of the heart-lung system to take up oxygen and rid the body of carbon dioxide.

Together, these and other studies show the value of adding a D-ribose supplement, 5 grams three times daily, to improve heart failure symptoms and quality of life in heart failure patients.

A deeper look into what happens in the muscles of a failing heart has opened an entirely new chapter in *metabolic cardiology*, the study of how the heart's own energy flow affects its pumping capacity. 9,10





### D-Ribose Supports Cardiac Energy Metabolism

After a hard workout or physical labor, our muscles are tired and weak. That's because, at the cellular level, they've used up much of the available molecules of ATP that provide the energy for the cellular work of moving. We rest after exercise or hard work in order to give our muscles time to burn *more* glucose to make *more* ATP.

The same situation occurs in the heart, with one vital difference: The heart muscle *can't* stop to rest and re-energize itself with more ATP. When ATP levels fall, the heart muscle simply operates with increasing weakness, pumping less blood with each beat.

If we can figure out a way to *boost* ATP levels, we can re-energize the heart muscle and may prevent or even *reverse* the symptoms of heart failure.

To understand how D-ribose benefits the heart, we need to look at the makeup of ATP itself.

The ATP molecule consists of three major parts, as seen in **Figure 1** on the next page.

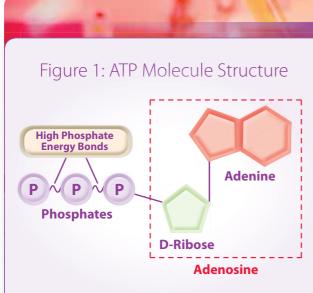
*Adenine* is a nucleotide, one of the building blocks of DNA.

*Adenosine* is simply adenine chemically bonded to a molecule of *D-ribose*.<sup>8</sup> D-ribose is a structural component of the vital ATP molecule.<sup>9</sup>

The *D-ribose* portion of adenosine binds tightly to a series of *phosphate molecules*, with each

# with D-Ribose Heart failure means the heart muscle

- Heart failure means the heart muscle is failing to pump enough blood to meet the body's metabolic requirements.
- Its incidence is rising, but there are no good, long-term medications available that truly treat heart failure and reverse its debilitating effects.
- The field of metabolic cardiology suggests that fundamental energy requirements go unmet in muscle cells of a failing heart, leading to significant drops in ATP, the energy-carrying molecule that supports all cellular work.
- One source of low ATP in heart failure is a diminished supply of the sugar *D-ribose*, which ailing heart cells cannot manufacture fast enough to meet energy requirements.
- Studies now show that supplemental D-ribose, 5 grams three times daily, can significantly improve heart muscle function in heart failure, resulting in better pumping action and reduced workload on heart and lungs.



Structure of the high-energy molecule <u>adenosine triphosphate</u>, or ATP.<sup>12</sup> Large red portion is adenine, which, when bound to D-ribose, is called adenosine. Opposite side of D-ribose portion binds tightly to up to three phosphate groups, with each chemical bond holding large amounts of energy that can be released when phosphates are broken away, first to adenosine **di**phosphate (one phosphate gone), and then to adenosine **mono**phosphate (AMP, with two phosphates gone).

chemical bond between the phosphate groups containing energy extracted from the metabolism of glucose and fat. ATP, with **three** such bonds, contains more stored energy than adenosine monophosphate (AMP), which has just **one**.

A maximum of *three* high-energy phosphate groups can be bound to a single adenosine molecule, producing adenosine *tri*phosphate, or ATP. When cells require energy, they release phosphates from ATP to carry out useful work by harvesting the newly-released energy.<sup>8,11</sup>

But in heart failure, a *cellular energy deficiency* arises in which D-ribose production falls in heart muscle cells. <sup>12</sup> This leads directly to a *decrease* in **ATP** production in the very cells that need energy the most. Without ample **D-ribose**, cells cannot capture metabolic energy, and almost literally starve themselves into weakness. <sup>8</sup>

The obvious solution, then, is to provide extra *D-ribose* in the form of an oral supplement, which bypasses the overwhelmed cellular *D-ribose-producing machinery and permits creation of many new ATP molecules.***<sup>8</sup>** 



And it works. One of the first studies of supplemental D-ribose was done in dogs back in 1989. It showed that ATP levels are depressed by about **50**% following loss of blood flow (*ischemia*) in the heart muscle. But when the dogs were supplemented with D-ribose intravenously, during and after the ischemic event, ATP levels <u>rebounded</u> to **85**% of their normal levels within 24 hours, a benefit not seen in the unsupplemented animals.<sup>13</sup>

In other words, **D-ribose replenishes low cardiac muscle energy levels**, improving cardiac function during ischemia.<sup>6</sup>

The evidence for D-ribose as a heart muscle energizer is so great that scientists have proposed treating it as a *conditionally essential nutrient*. This means that, under certain conditions, like heart failure, D-ribose is so important that its levels *must* be supported by external supplementation.<sup>14</sup>

### Summary

Heart failure is increasing, with an estimated **eight million** sufferers expected by **2030**.

Currently, drugs for heart failure are only marginally effective and act by forcing the heart muscle to work *harder*, even as it is being deprived of sufficient ATP to perform extra work.

An alternative approach, called *metabolic car-diology*, offers a more natural and physiologically sound way to improve heart muscle performance.

Researchers have found that we can improve the conditions that lead to ATP production within the heart muscle itself by providing D-ribose as a vital "substrate" for ATP production.

Studies in animals, and now in humans, have shown that increasing available D-ribose, the "scaffold" on which ATP is built, can dramatically restore energy to the heart, enabling it to carry out its vital work with greater power and efficiency.

Regular supplementation with D-ribose is one proven means of feeding the starving heart muscle and producing real improvements in cardiovascular function.

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



### What Is Heart Failure, Anyway?

"Heart failure," despite its dramatic-sounding name, is not a sudden, fatal event.

The term refers to a situation in which the heart muscle cannot keep up with the body's demands for ample blood flow. In other words, the heart "fails" to provide adequate supplies of blood. As a result, a host of organs and systems run on insufficient fuel, leading to gradual, systems-wide malfunctions.

Heart failure doesn't appear out of nowhere. Generally, either the heart muscle is damaged (by a heart attack, for example), or the workload faced by the heart muscle rises sharply (as with severe high blood pressure).15

Regardless of the cause, heart failure typically results in an overstretched, thickened, and enlarged left ventricle (the main pumping chamber), which overburdens the heart's ability to work efficiently.9,16

The outcome is that there is a reduction in the volume and pressure the ailing heart muscle can pump with each contraction. That causes both an insufficient forward flow, and a backup of blood that can't get back to the heart for recirculation.

These factors produce the common symptoms of heart failure, which worsen as the condition becomes more severe.17,18

In early heart failure, patients may experience no symptoms at all. As heart failure progresses, people may be comfortable at rest, but find that ordinary physical activity produces fatigue, heart palpitations, trouble breathing, or chest pain.

In later stages, symptoms markedly limit physical activity. In the most severe stage (Stage D, in the American Heart Association's classification), people with advanced heart failure may have symptoms at rest and are unable to carry out any physical activity without discomfort or a sharp increase in symptoms. These patients require aggressive medical therapy just to survive.

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### New Study to Evaluate D-Ribose, CoQ10 Combination in Heart Failure

Metabolic cardiology focuses on preventing and treating cardiovascular disease at the cellular level, by preserving and promoting the production of ATP. It holds the promise of reducing death and disability from heart failure.

The approach described in this article, providing supplemental D-ribose to boost available ATP in heart muscle, is one of the most important contributions of metabolic cardiology to date. But it is not the only one.

In a planned study by researchers at the University of Kansas School of Nursing and Department of Molecular and Integrative Physiology, subjects with heart failure will be randomly assigned to receive either a placebo, D-ribose (15 grams/daily), a form of coenzyme Q10

(CoQ10) called ubiquinol (600 mg/day), or a combination of both D-ribose and ubiquinol.3

The study will continue for 12 weeks, and will measure multiple outcomes, including symptoms, vigor, echocardiographic measurements of heart chamber sizes and function, and walking endurance. Measures of ATP production in mitochondria will also be taken.

This is just one example of a marked shift in researchers' attitudes and approaches to heart failure. Instead of drugs that "whip" the heart muscle to do more work with less energy, supporting cardiac energy levels with D-ribose and CoQ10 offers the chance to genuinely improve heart muscle function.

### **Heart Failure Protocol**

Decades ago Life Extension® identified studies in Japan showing heart failure patients improving in response to 3,000 mg a day of the amino acid taurine.19

Japanese research on CoQ10 led a pioneering American cardiologist, Peter Langsjoen, MD, to develop a protocol for heart failure patients that involved high-dose **ubiquinol** (**580 mg** on average and after a few months reduce the ubiquinol dose to about 400 mg daily).20

Dr. Langsjoen's dosing of ubiquinol is based on each patient's blood levels of CoQ10. Life Extension offers CoQ10 blood testing to help those who need to achieve higher blood concentrations. To inquire about CoQ10 blood testing, call 1-800-208-3444 (24 hours) or visit www.LifeExtension.com/blood

For healthy people not supplementing with CoQ10, Dr. Langsjoen suggests starting with 300-400 mg per day for the first month to fully saturate your cells. People can then back down to a daily dose of 100-200 mg per day to maintain high cellular CoQ10 levels.21,22

A recent animal model study indicates that heart failure patients might benefit by boosting the **NAD**<sup>+</sup> levels.<sup>23</sup> This can be accomplished by taking **250-500 mg** a day of **nicotinamide riboside**. NAD<sup>+</sup> is required for healthy cell energy functions.

There are also indicators that heart failure patients might benefit from taking senolytic compounds that selectively remove senescent cells. The accumulation of senescent cells damages healthier heart cells by spewing protein-degrading enzymes and igniting persistent inflammatory reactions.

So an updated heart failure protocol might include:

- 5,000 mg of D-ribose powder 3 times daily (15,000 mg total per day)
- 3,000 mg a day of taurine (a low-cost supplement)
- 200-600 mg of ubiquinol (dose can be based on blood test results)
- 250-500 mg a day of nicotinamide riboside
- 500 mg a day of magnesium
- 1,000 mg a day of carnosine
- · Senolytic protocol of your choice

These approaches are NOT meant to substitute for conventional prescription drugs that are extending lives of heart failure patients. Their efficacy is evidenced by a 44% decline in sudden death rates of heart failure patients that occurred between 1995-2014 with more intelligent use of these drugs by conventional cardiology.<sup>24</sup> This period is also associated with surging use of **CoQ10** that heart failure patients often take.

The next page features some Power Point slides that were used to describe recent findings on the potential of NAD+ and senolytics to improve heart muscle function, along with a listing of prescription drugs often prescribed for heart failure patients.

**Note**: Take **CoQ10** with a fatty meal to further improve **absorption**.



### Recent Findings on NAD+ and Senolytics Relating to Heart Failure



### Aorta

Weakened heart muscle cannot pump enough blood into the aorta. Blood pools in the heart resulting in heart failure.

January 8, 2019 Heart Failure Fact Sheet

- Heart failure happens when the heart cannot pump enough blood and oxygen to support other organs in your body.
- About half of people who develop heart failure die within 5 years of diagnosis.
- 8 million Americans suffer heart failure.

 $https://www.cdc.gov/dhdsp/data\_statistics/fact\_sheets/fs\_heart\_failure.htm$ 



February 8, 2019

Newcastle University
Scientists are Killing Zombie
Cells to Reverse Age-Related
Damage in the Heart

"... senescent cells – also known as zombie cells – form in the heart during aging and lead to heart failure."

"Newcastle scientists, in collaboration with researchers in the Mayo Clinic... have not only discovered how this process takes place in the heart, but also how it can be reversed or treated."



May 21, 2018 NAD+ May Improve Heart Function

"Stabilizing the intracellular NAD+ level represents a promising therapeutic strategy to improve myocardial bioenergetics and cardiac function."

"In this issue of Circulation,
Diguet et al report exciting data
suggesting that supplementation
with a NAD+ precursor, nicotinamide
riboside, reduces cardiac dysfunction
in preclinical models of heart failure."



Sharp Decline in Heart Failure Death Rates

- Clinical trials spanning 1995 to 2014 show:
- 44% decline of sudden death rates in heart failure patients.
- Marked improvement in just 19 years!



"Declining Risk of Sudden Death in Heart Failure." New England Journal of Medicine; 2017;377(1):41-51.

### Proper Medication Use Buys Heart Failure Patients Extra Years

Combinations of the following drugs treat heart failure under cardiologist supervision:

- Beta Blockers (carvedilol, metoprolol)
- ACE Inhibitors (lisinopril, captopril)
- Angiotensin Receptor Blockers (losartan)
- Digoxin (Lanoxin)
- Hydralazine and Nitrates (Apresoline, Nitrobid, Imdur, Isordil)

Without innovative approaches, heart failure eventually results in death.

https://wa.kaiserpermanente.org/healthAndWellness/index.jhtml?item=%2F common%2FhealthAndWellness%2Fconditions%2FheartDisease%2FchfMed ications html

### Senolytics to Treat Heart Failure Need Human Validation

### What we learned from this 2019 published study:

- Senescence involving cardiomyocytes (i.e. cardiac muscle cells) appears to be associated with age-linked fibrosis and hypertrophy of the heart.
- Pharmacologic and genetic removal of p16-positive (senescent) cells in mice appears to ameliorate (or improve) the level of age-induced fibrosis and hypertrophy in senescent cardiac muscle cells, and may support regeneration of cardiomyocyte.

Clinical trial on existing senolytic compounds urgently needed!

EMBO J. 2019 Mar 1;38(5). pii: e100492.

### What Are Senolytics?

- "Seno" refers to senescent cells
  - "Lytic" refers to destruction

Senolytics selectively destroy senescent cells.



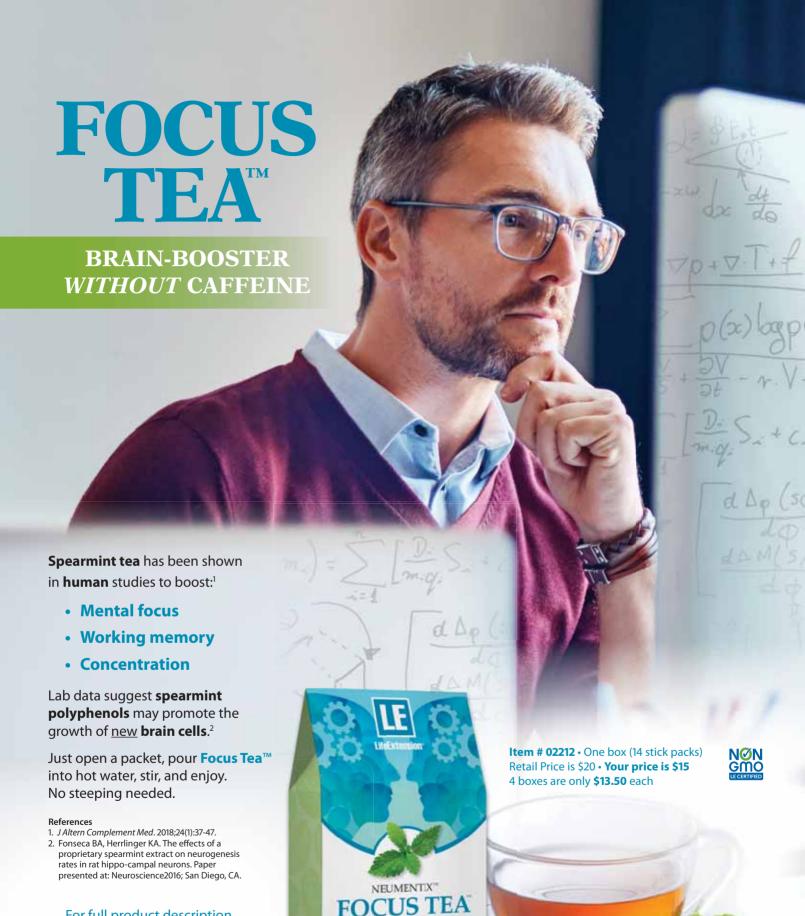
February 11, 2019

"Scientists believe it may be possible to reverse the heart damage caused by aging"

"We saw that removing senescent cardiomyocytes from the hearts of aged mice, both genetically and using drugs, was able to restore cardiac health – essentially removing the damage caused by aging.

These data provide critical support for the potential of using medicines to kill zombie cells. If this is validated through clinical trials it would provide us with a new way of treating cardiac diseases."

Rhys Anderson et al. Length-independent telomere damage drives post-mitotic cardiomyocyte senescence, The EMBO Journal (2019). DOI: 10.15252/embj.2018100492t



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# 

**Whey protein** is widely assumed to be a supplement just for athletes seeking to add additional muscle mass.

That's just a small part of the picture. Studies reveal that whey has anti-aging benefits.

Research shows that whey reduces **muscle** wasting in the elderly, inhibits weight gain, may help prevent cardiovascular disease, and more.<sup>1-11</sup>

Whey is a food source studied for maximizing production of **glutathione**, 12,13 one of the body's main internally produced, free radical scavengers. 14,15

Glutathione levels drop with age, and this could play a role in neurodegenerative disease, reduced immunity, and a host of other age-related conditions. 16-20

BY MICHAEL DOWNEY

Whey, made from the liquid part of milk that separates during cheese production, is not just a source of protein, but also of nutrients including branched-chain amino acids, immunoglobulins, and bio-active protein sub-fractions such as lactoferrin.

These benefits show why whey is increasingly viewed as a food that can prevent frailty and promote healthy longevity.

### Whey Keeps Muscles from Weakening

About 45% of older individuals in the general population, and more than 84% in residential care facilities, are *not* adequately nourished.<sup>21,22</sup> This happens, among other reasons, because older people often have reduced appetites and eat less, while nutrients aren't absorbed as well as we age. 22-24

An insufficient intake of quality protein can lead to loss of muscle mass. For most people, this muscle loss begins around age 40, with an estimated 8% loss of muscle mass per decade. After age 70, muscle mass decreases by about 15% per decade.<sup>24</sup>

Approximately 5%-13% of people aged 60 or over experience age-related muscle-wasting that is so severe, it increases the risk of falls and disability.<sup>25-27</sup>

Inadequate protein consumption among older individuals is associated with reduced strength, decreased bone mass, low immunity, cognitive impairment, and delayed recovery from wounds and surgery.<sup>28</sup> In fact, low protein intake is associated with **frailty**, <sup>29</sup> when the body is so weak it becomes unable to cope with stress or injury. Frailty is a strong predictor of mortality in the elderly.<sup>21,30</sup>

Loss of muscle mass is not inevitable but does require some active countermeasures to prevent it.

Whey protein delivers an abundance of **branched**chain amino acids (BCAAs), essential nutrients that reduce muscle breakdown and stimulate the building of protein in muscle.31

Of the three BCAAs found in whey, *leucine* is the most metabolically active, helping to build muscle by activating a signaling pathway that controls the body's anabolic (growth-promoting) drive. <sup>2,32-35</sup> Aging muscle ordinarily becomes resistant to leucine stimulation, but taking leucine-rich whey can overcome this, stimulating muscle synthesis.33-37

One study focused on hospitalized elderly men and women who had been classified as frail and assessed the effects of whey on rehabilitation outcomes. The intervention group received whey supplements daily for the duration of their hospital stay.6

Compared to patients who didn't receive whey. those who took it showed significant improvements in grip strength and knee extensor force. Scientists concluded that whey improves the nutritional status and rehabilitation outcomes among frail and elderly people.6

### Whey Boosts Lean Muscle Mass

Whey has valuable potential to help prevent agerelated muscle loss. But the benefits go even further. Recent studies have shown that whey significantly increases lean muscle mass—and not just among athletes.

In a randomized, controlled trial, researchers divided 81 healthy, older women, aged 65-80, into three groups for a 24-week program. One group



exercised twice weekly, the second group took whey protein supplements but didn't exercise, and the third group took whey protein after exercising.<sup>4</sup>

The increase in skeletal muscle mass was significantly *higher* for the whey-and-exercise group than for either of the two other groups. There was also a significant increase in grip strength and walking speed.<sup>4</sup>

Researchers next turned to inactive older individuals, conducting a study to assess whey's effects on muscle during recovery from periods of *inactivity*, such as hospitalization. In this controlled trial, men and women in their late sixties consumed a diet in which **45**% of their protein intake came from either whey or a source of animal peptides.<sup>1</sup>

After a week of habitual activity and energy balance, they spent another week in energy restriction, during which 500 fewer daily calories were consumed per day. The number of steps they took daily was then reduced for another two weeks. Finally, they returned to normal activity for the last week (recovery).<sup>1</sup>

During the periods of reduced activity, lean leg mass declined in both groups. But during the recovery week, lean leg mass increased—only in the whey protein group. 1 The branched-chain amino acids found in whey play an essential role in supporting lean muscle mass, especially among older adults.<sup>38,39</sup>

### Whey Helps Stop **Age-Related Weight Gain**

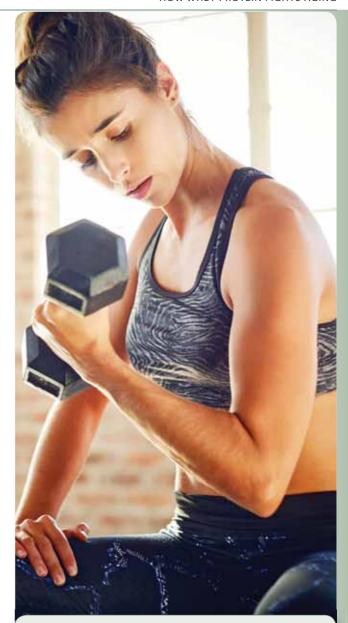
Whey doesn't just affect the muscles. As we age, metabolism slows, and we start to put on weight more easily, increasing our risk of everything from heart disease to diabetes to strokes. This used to seem like an inevitable part of life.

Not anymore. Research shows that whey is an effective way to fight fat, helping to maintain weight and lean body mass as we age.40

Scientists reviewing many previous studies concluded that whey may even be an effective therapeutic treatment for obesity.40

When other groups of researchers set about trying to understand why whey protein supplementation works in this way, they discovered that the proteins, peptides and minerals in whey boost satiety (the feeling of fullness), influence glucose homeostasis (the regulation of blood sugar levels in the body), and optimize lean body mass, all of which help fight weight gain.41-45

In one recent study, scientists assigned 100 men with what's known as **sarcopenic obesity**, aged 70 or over, to one of three groups. Sarcopenic obesity is characterized by low lean mass and high fat mass. One group served as a control and received no treatment.



### Whey Protein and Aging

- Whey protein is not just for athletes seeking to build muscle mass.
- Whey supplementation has been shown to help prevent the loss of muscle mass in aging individuals, inhibit weight gain, and reduce risk factors for cardiovascular disease.
- This potent protein helps prevent frailty, obesity, and heart disease, while promoting longevity.



Another used an exercise technology called whole-body electromyostimulation (WB-EMS), in which muscles are contracted with electrical impulses. The third took **1.7 grams** to **1.8 grams** of whey protein per kilogram of body weight, daily.<sup>10</sup>

After 16 weeks, total body fat, trunk body fat, and waist circumference were significantly reduced in the group receiving electrical stimulation *and* were also reduced in the group taking whey supplements, but not in the control group. Whey protein supplementation, the study concluded, can help treat obesity.<sup>10</sup>

Also, in a meta-analysis of randomized, controlled trials conducted in overweight and obese participants, investigators reported that there was a significant **improvement** in both body weight and total fat mass in those who supplemented with whey protein.<sup>11</sup>

### Whey Helps Prevent Heart Disease

By contributing to an overall diet and exercise fitness program, whey can protect against a long list of diseases linked to increased weight gain. But whey protein has also been shown to *specifically* target a key risk factor for **cardiovascular disease**.

Hypertension is one of the main factors contributing to cardiovascular disease. <sup>46</sup> Research shows that whey-based peptides may help reduce this risk factor. <sup>47,48</sup> (Peptides are chains of amino acids that are smaller than proteins.) And food-derived peptides like the kind found in whey are far safer than antihypertension drugs.

In one study, researchers asked 27 adults with **mild hypertension** to eat a high-fat breakfast and lunch along with **28 grams** of whey protein. This was later repeated with **28 grams** of calcium caseinate (a protein produced from casein in skim and sometimes **1%** milk). It was again repeated with **27 grams** of maltodextrin (a type of carbohydrate made from starch).<sup>5</sup>

Compared with the other supplements, whey was found to *reduce* systolic blood pressure, by an average of **15.2 mmHg**, for up to five hours after ingestion. Compared to maltodextrin, whey *improved* arterial stiffness. These results all show whey's potential to improve cardiovascular risk factors.<sup>5</sup>

Another group of scientists reviewed whey's effects on the cardiovascular risk factors of *overweight and obese* patients. They found whey protein supplementation resulted in a reduction in body weight *and* reduced multiple risk factors for cardiovascular disease in those patients. Improvements were found in systolic blood pressure, diastolic blood pressure, glucose levels, and in HDL and total cholesterol.<sup>11</sup>

### **Summary**

Whey protein isn't just for athletes and bodybuilders. Whey supplementation has now been shown to help prevent several common effects of aging, including loss of lean muscle mass and excessive weight gain. It also lowers cardiovascular risk and blood pressure. It is a powerful tool to prevent frailty and heart disease and to boost longevity. •

#### What Type of Whey Protein Supplement is Right for You?

Whey protein is commonly available in three forms:

- Concentrate,
- Isolate, and
- Isolate with added creatine and glutamine.

Whey concentrate is whey, but with the water removed. It is rich in both branched-chain amino acids (BCAAs) and glutamine. The concentrate form of whey is a powder that mixes easily for a smooth-textured protein shake. High-quality whey concentrates contain no animal growth hormones. Most whey concentrates contain about 80% protein. It is the ideal protein to help build muscle size and aid in recovery. Overall, whey concentrate may be the most economical form of protein for the human body to digest and use, which is why it is among the world's most popular sports supplements.

Whey isolate goes through additional processing steps and filtration that remove some of the carbohydrate, lactose, and fat, providing a purer protein as the end product. Whey isolate contains about **90%** protein. In high-quality whey isolate, 98% of the protein content is in the bioavailable, un-denatured form similar to concentrates and it retains essential bioactive subfractions in their natural ratios. It, too, is free of animal growth hormones. (Those who are lactose-intolerant should note that, like whey concentrate, whey isolate contains a small amount of lactose.)

Whey isolate with added creatine and glutamine is a premium isolate option for those seeking *greater* strength and exercise performance.

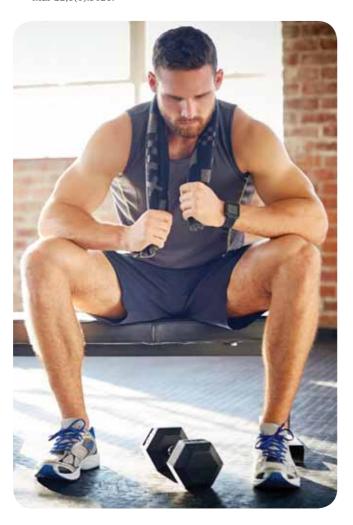
- Creatine supports energy storage by tissues, increases levels of ATP (the molecule that stores and transports energy within cells) in mitochondria, and helps maintain healthy muscle mass, which generally declines with aging.<sup>49-51</sup> Studies show that creatine helps build muscle and strength in explosive, short-duration activities such as resistance exercise training. 52,53
- Glutamine is considered a conditionally essential amino acid<sup>54</sup> that is abundant in muscles, but levels are reduced after prolonged and highintensity exercise, which can affect the immune system if supplies are not replenished.<sup>55-58</sup> Glutamine encourages recovery after intense exercise, increases glycogen fuel synthesis, and helps inhibit protein breakdown in muscle tissue. 59-61 It can inhibit blood ammonia accumulation during prolonged exercise, 62,63 a key factor in physical fatigue.<sup>64</sup> Glutamine is also essential to gut integrity, as it provides metabolic fuel for digestive tract cells, helps regulate cellular reproduction, aids in the maintenance and repair of the lining of the gastrointestinal tract, and has anti-inflammatory properties. 65,66



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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#### References

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# Topical Peptides Rebuild Youthful Skin

# BY ROBERT GOLDFADEN AND GARY GOLDFADEN, MD

**Peptides** are amino acids naturally present in **young** skin.

As we age, there is **decreased** production of these **peptides**.

When applied to older skin, **peptides** promote growth, organization, and maintenance of cells. The result is restoration of more youthful skin structure and function.<sup>1,2</sup>

An advance in topical skin rejuvenation has been the discovery of specific peptides called **growth factors**.<sup>3</sup>

These growth factor **peptides** have been found to repair and reverse chronological and environmental skin damage.<sup>4</sup>

#### Growth Factors: Master Regulators of Skin Repair

Young skin efficiently repairs damage from internal and external insults, thanks to the inner workings of **growth factors**.

When skin is damaged, there is a natural release of **growth factors** that initiate and coordinate reparative processes. These growth factors accomplish this by communicating with other cells, like keratinocytes and fibroblasts. 5-7

Scientists have zeroed in on <u>five</u> key **growth factors** that modulate skin repair and protect against premature aging.

Here is how each **growth factor** functions:

- **Fibroblast growth factor (FGF)** stimulates proliferation of dermal fibroblasts responsible for producing major skin matrix constituents such as collagen, elastin, and fibronectin. 8,9
- Vascular endothelial growth factor (VEGF) helps boost the delivery of oxygen and vital nutrients. 10,11
- Acidic fibroblast growth factor (AFGF)
  increases the growth and differentiation of
  epidermal keratinocytes to renew the skin's
  surface.<sup>12,13</sup>
- Epidermal growth factor (EGF) attenuates the strong inflammatory and oxidative stress response following skin damage.<sup>14</sup>
- Insulin-like growth factor (IGF-1) triggers new production of collagen and protects against its breakdown by downregulating collagenase. 15,16

These **growth factors** control cellular growth, proliferation, and differentiation, to give skin the capacity to renew itself completely in our youth. This is why young individuals are able to temporarily combat the detrimental effects of sun exposure, air pollution, and advanced glycation end products (AGEs).

As we grow older, decreased production of **growth factors** compromises the skin's ability to preserve structural integrity and function.<sup>17,18</sup> This finding prompted researchers to investigate the impact on aging skin of replenishing **growth factors**.

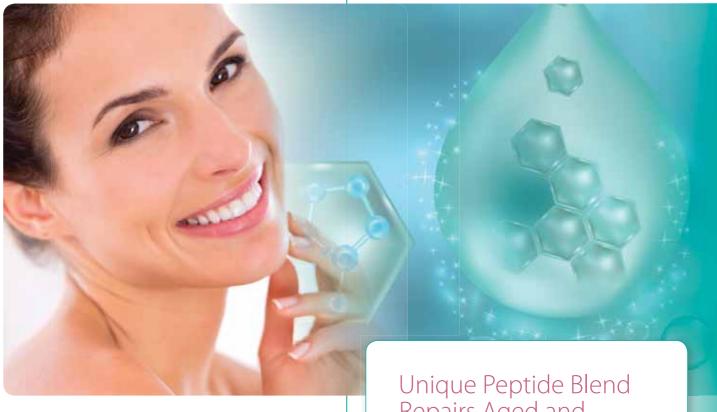
#### **Clinical Efficacy of Topical Growth Factors**

Human trials show that topical preparations with one or more (but not all) of the **growth factors** described so far function to boost **collagen** and **elastin** synthesis, promote **epidermal thickening**, and **quell inflammation**. Findings from these studies reveal improvements in skin that was aged and photodamaged. <sup>19-23</sup>

When all five growth factors are *physiologically balanced* in one topical formula, the results are more impressive. In one human study, female participants who applied a topical cream containing all five **growth factors** daily for four weeks experienced the following:<sup>4</sup>

- Decreased wrinkles under the eyes by **46**%
- Reduced appearance of crow's feet by 21%
- Increased skin elasticity by 47%
- Enhanced skin hydration by 64% (improved by 83% immediately after applying)





#### **Accelerate Epidermal Regeneration**

Epidermal stem cells are responsible for selfrenewing the skin's surface.

These stem cells manufacture basal keratinocytes that travel upward to form the "brick and mortar" structure that acts as a barrier against the external environment.<sup>24,25</sup>

As we age, epidermal turnover rate slows down as stem cell activity decreases. This causes thinning of the epidermis, making it more susceptible to external threats, and allowing vital moisture to escape.

**Sea fennel** is a flowering plant whose stem cells allow it to adapt and thrive in salty coastal areas of Europe. When researchers tested the impact of sea fennel (Crithmum maritimum) extract on aged human skin cells, they found that it increased the viability of skin cells by 120% compared with untreated controls. This rapidly increased the density and thickness of the epidermis, enabling the extract-treated group to achieve complete epidermal regeneration significantly faster than the non-treated group.<sup>26</sup>

These in-vitro findings led to a human study in which 12 healthy volunteers with a mean age of 56 applied **sea fennel** extract or a placebo daily to their forearm skin for two weeks before undergoing a toxic protocol that induced epidermal damage.

# Repairs Aged and Damaged Skin

- Growth factors are dynamic peptides that activate cell-signaling pathways to orchestrate the repair of chronological and environmental skin damage.
- Decreased concentration of growth factors as we age compromises the skin's healing capacity, setting the foundation for wrinkles, age spots, and sagging skin.
- Topical replenishment of five key growth factors has been shown to stimulate epidermal and dermal regeneration to reverse visible signs of aging.
- Two plant extracts—sea fennel (Crithmum maritimum) and date palm fruit (Phoenix dactylifera L.)—further support skin renewal and maintenance.
- All of these compounds have been incorporated into one unique, topical formula that results in healthier, more youthful and radiant skin.



Researchers observed that within 96 hours, the group treated with **sea fennel** extract had significantly lower transepidermal water loss than the placebo group, indicating faster recovery of epidermal structure and function.27

Research also shows improvements in skin radiance, tone, and firmness with topically applied sea fennel extract.28

#### **Fade Hyperpigmentation and Wrinkles**

Date palm (*Phoenix dactylifera* L.), a plant native to southern Asia and North Africa, confers anti-inflammatory activity due to its abundance of polyphenols, tannins, and carotenoids.29

Hyperpigmentation is a common skin disorder that stems from excessive production of the pigment melanin in response to cumulative sun exposure. It manifests as age spots, dark patches, and melasma that contribute to older looking skin.<sup>30</sup>

In a clinical trial, female volunteers topically applied a facial cream with date palm extract daily for eight weeks. Compared to baseline, researchers observed an aproximate 25% reduction in melanin output, thereby demonstrating **skin lightening** effects. This was accompanied by a decrease in erythema, as well as increases in skin elasticity and hydration, which left participants with even-toned, youngerlooking skin.31

Date palm fruit extract has also delivered antiwrinkle activity. Topical application of a facial cream containing date palm extract twice daily for five weeks reduced the total surface area of wrinkles by 27.6% and the depth of wrinkles by 3.52% compared to a placebo in a group of women aged 46 to 58.32 These potent wrinkle-reducing effects might be attributed to the presence of phytoestrogens which induce collagen formation for dermal renewal.33,34

#### **Summary**

Scientists have identified five key growth factors that <u>activate</u> natural repair pathways in the skin to offset damage from chronological aging and external stressors.

Clinical research verifies that replenishing growth factor peptides restores skin elasticity, tone, and hydration, while reducing wrinkles.

These growth factors have been combined with sea fennel and date palm fruit to support more youthful skin structure.

Gary Goldfaden, MD, is a clinical dermatologist and lifetime member of the American Academy of Dermatology. He is the founder of Academy Dermatology in Hollywood, FL, and Cosmesis Skin Care. Dr. Goldfaden is a member of the Life Extension® Medical Advisory Board. All Cosmesis products are available online.

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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### A DAILY FOOT MASSAGE LOWERS BLOOD PRESSURE AND LOWERS TRIGLYCERIDE LEVELS

Pusan National University, South Korea

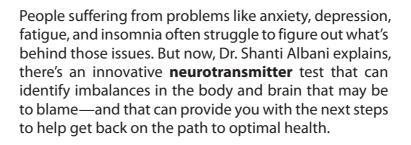


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A Window to the Brain



LE: Back in 2016, Life Extension® made neurotransmitter testing available to our customers. This test was, and has continued to be a popular test. What accounts for the popularity of the Neurotransmitter Panel?

Dr. Albani: Neurotransmitter testing provides insight into imbalances that may be occurring in the body and brain and can shed light on related symptoms. Basic blood work is great for evaluating blood sugar, cholesterol, and organ function, but often turns up little when individuals are trying to determine why they have symptoms such as anxiety, insomnia, depression, or poor cognition. Neurotransmitter testing is a window to our brain function that no other testing can provide. Ideally, people would also test their hormones along with their neurotransmitters, since hormones are also key players in mood and energy.

LE: It's fascinating that we can measure neurotransmitters in the urine. Could you explain how this works?

Dr. Albani: Neurotransmitters are the chemicals that facilitate the transmission of signals from one neuron to the next across a synapse, or from a neuron to target cells such as muscles or glands. They are produced in the brain, but also in other areas of the body. For example, our intestines have a vast neural network and produce neurotransmitters to help coordinate digestion and intestinal motility. The urinary Neurotransmitter Panel provides a measurement of wholebody neurotransmitter production. Studies and clinical experience have also established that the levels measured in urine correlate with both mental and physical symptoms.

LE: You mentioned testing hormones. How are hormones and neurotransmitters related?

Dr. Albani: Both hormones and neurotransmitters powerfully influence mood and energy, so I encourage people to have their hormones and neurotransmitters tested together. Hormones can also influence neurotransmitters. and vice versa. For example, the hormone estrogen can increase activity of the neurotransmitters serotonin and dopamine, while the hormone progesterone increases the calming action of the neurotransmitter GABA. For their part, neurotransmitters can influence hormone production via their communication with the pituitary gland in the brain, which in turn controls hormone production.

When functioning properly, neurotransmitters and hormones work in harmony to sustain good mental and physical health. However, alterations in neurotransmitters and hormones can play a significant role in contributing to symptoms such as cognitive disorders, depression, anxiety, diminished drive, fatigue and sleep difficulties, cravings, addictions, and pain.

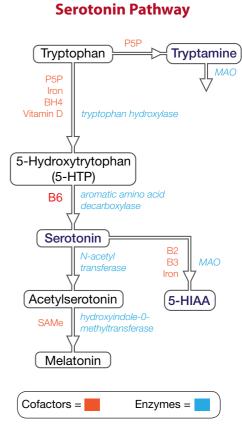
LE: Once people test their neurotransmitters and identify imbalances, how does this information assist them on the path to health?

**Dr. Albani:** This is the exciting part! Fortunately, vitamins, minerals, amino acids, hormones, and botanical medicine can be used to rebalance our brain chemistry. Our understanding of neurotransmitters has advanced so much that we know which nutrients are needed as cofactors for the enzymes that create, or break down, each neurotransmitter. Vitamin B6, for instance, is needed for the conversion of the amino acid 5-HTP into serotonin, so individuals who have low serotonin should ensure they are getting enough B6. And since we know that neurotransmitters are created from amino acids, we can use amino acid therapy to support neurotransmitter production in a targeted way. For example, we know that tryptophan is the amino acid precursor to serotonin. Therefore, taking tryptophan supports serotonin production. Our understanding of how hormones and botanical medicines influence brain biochemistry can also help us to balance neurotransmitters.

**LE**: Have there been any upgrades to the Neurotransmitter Panel?

Dr. Albani: Yes. There's a Basic Neurotransmitter Panel, which tests for nine neurotransmitters: PEA (phenylethylamine), dopamine, norepinephrine, epinephrine, serotonin, glutamate, GABA (gamma-aminobutyric acid), glycine, and histamine. But we now also offer a Comprehensive Neurotransmitter Panel. Comprehensive Neurotransmitter Panel tests for the nine neurotransmitters I mentioned, along with the neuroactive amino acid taurine and several neurotransmitter precursors and metabolites, namely, tyrosine, tyramine, DOPAC, 3-MT, normetanephrine, metanephrine, tryptamine, and 5-HIAA.







#### **Neurotransmitter Panel – Comprehensive**

Analyte	Result	Unit per Creatinine	L	WRI	Н	Reference Interval
Phenethylamine (PEA)	8	nmol/g				32 - 84
Tyrosine	24	μmol/g				32 - 80
Tyramine	2.1	μmol/g	Δ			2.0 - 4.0
Dopamine	149	μg/g				125 - 250
3,4-Dihydroxyphenylacetic acid (DOPAC)	2388	μg/g				390 - 1500
3-Methoxytyramine (3-MT)	126	nmol/g		<b>A</b>		90-210
Norepinephrine	14.9	μg/g				22-50
Normetanephrine	207	μg/g				85 - 300
Epinephrine	1.2	μg/g				1.6 - 8.3
Metanephrine	103	μg/g		Δ		45 - 119
Norepinephrine / Epinephrine ratio	12.4				7	< 13
Tryptamine	0.67	μmol/g		A		0.20 - 0.90
Serotonin	66.7	μg/g	Δ			60 - 125
5-Hydroxyindolacetic acid (5-HIAA)	2539	μg/g				2000 - 8000
Glutamate	16	μmol/g				12.0 - 45.0
Gamma-aminobutyrate (GABA)	4.8	μmol/g				2.0 - 5.6
Glycine	1121	μmol/g		<u> </u>		450 - 2200
Histamine	18	μg/g				14 - 44
Taurine	864	μmol/g		A		320 - 1000
Creatinine	61	mg/dL	Δ			30 - 225

#### **Neurotransmitter Panel – Comprehensive**

**LE**: What is the advantage of the Comprehensive Neurotransmitter Panel? How does measuring taurine and the additional neurotransmitter precursors and metabolites provide further benefits?

Dr. Albani: The Comprehensive Neurotransmitter Panel provides significantly greater detail on where in the creation or metabolism of our neurotransmitters there may be an imbalance. For example, the Basic Neurotransmitter Panel may identify that an individual has low dopamine. But the Comprehensive Neurotransmitter Panel can help identify if that is because of low levels of the amino acid tyrosine, because of over-activity of the MAO or COMT enzymes (which break down dopamine), or because of some other factor (see dopamine pathway figure on the

next page). This level of detail can allow us to be even more targeted with our therapeutic suggestions for neurotransmitter balance.

Ultimately, both neurotransmitter tests are extremely valuable, but the Comprehensive Panel provides more detail for those who want or need it.

LE: Who should have this test done?

Dr. Shanti: This test is especially valuable for individuals who have mental and emotional symptoms that evade identification with conventional testing. Some examples of symptoms which can indicate neurotransmitter imbalances are:

- Depressed mood
- Anxiety
- Fatigue
- Poor sleep

- Loss of mental focus
- Attention deficit disorder (ADD) and attentiondeficit/hyperactivity disorder (ADHD)
- Addiction or dependency
- Loss of appetite control
- Compulsive behavior
- Cravings
- Low libido
- Altered pain response
- Poor mental performance

Many individuals who suffer from these symptoms may compound their health issues by using food, caffeine, chocolate, alcohol, nicotine, medications, and other neuroactive substances to try to obtain relief from their symptoms. When functioning properly, there is a balance between the excitatory and inhibitory neurotransmitters.

#### **Dopamine Pathway Tvramine Tyrosine** P5P Iron BH4 tyrosine hydroxylase Vitamin D L-Dopa **DOPAC** aromatic amino acid decarboxylase Dopamine | Vitamin C dopamine B-hydroxylase Copper Norepinephrine } Cofactors = Enzymes =



Restoring this balance can help individuals enjoy life again and feel less dependent on the substances I just mentioned to maintain daily function.

LE: If people are interested in testing their neurotransmitters, how should they proceed?

Dr. Albani: Many conventional doctors are not familiar with urinary neurotransmitter testing. Some innovative, forward-thinking physicians may offer the test through specialty labs. For customers who are unable to obtain the test through their doctor, Life Extension has partnered with a lab that uses the gold standard testing method of liquid chromatography combined with tandem mass spectrometry (LC-MS/MS) to provide neurotransmitter testing.

LE: How is the test collection done?

Dr. Albani: Collection is easy and requires only a single, morning urine sample. The test comes in the form of a home collection kit, so the urine collection is noninvasive and can be done in the comfort of your own home. Once the collection is complete, it is sent to the lab using the pre-paid shipping label provided.

LE: Once someone has the test results, how can they be used to effect health improvements?

Dr. Albani: All test results come with an explanation of what the results may mean, along with nutrient suggestions to balance neurotransmitters. In addition to the useful information provided with the results, individuals can work directly with their doctors or call to speak with one of our trained Life Extension Wellness Specialists. Our team of wellness specialists can provide individualized suggestions to people for

nutrients to balance neurotransmitters, based on their test results and on how they feel. The information they provide also helps people have more detailed conversations with their own doctors. •

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

Dr. Shanti Albani obtained her medical degree in Naturopathic Medicine in 2003 from the National University of Natural Medicine in Portland, Oregon. She practiced medicine for many years in central

Mexico specializing in gastrointestinal disorders and hormone balance. During this time, she also owned a nutrition store and taught courses in bioidentical hormone replacement therapy for physicians. She has worked at **Life Extension** since 2010 and is currently the Manager of Clinical Information.

# Brussels Sprouts

#### BY LAURIE MATHENA

Surveys have put Brussels sprouts at the top of the list of most-hated vegetables in the U.S.<sup>1</sup> But if you've been shunning these "miniature cabbages," it's time to give them a second chance.

#### **Cancer**

Brussels sprouts contain cancer-fighting compounds called *glucosinolates*, which break down in the body into indoles and isothiocyanates. These compounds have been shown to inhibit the development of numerous types of cancer in animals, including bladder, breast, colon, lung, liver, and stomach cancers.<sup>2,3</sup>

#### **Diabetes**

Studies have shown that a diet high in cruciferous vegetables like Brussels sprouts decreases the risk of **type II diabetes**.<sup>4,5</sup> Brussels sprouts contain alpha-lipoic acid,6 which has been shown to increase insulin sensitivity in type II diabetics.7

The best way to prepare Brussels sprouts is by lightly steaming them. Not only will this produce the best flavor and retain the most nutrients, but it will help prevent the unpleasant sulfur smell that can make people turn their noses up at this nutritious vegetable.

- 1. Available at: https://www.hy-vee.com/health/ healthy-bites/brussels-sprouts-the-most-hatedvegetable-in-america.aspx. Accessed April 1, 2019.
- 2. Drug Metab Rev. 2000 Aug-Nov;32(3-4):395-411.
- 3. Nutr Cancer. 2001;41(1-2):17-28
- 4. Prim Care Diabetes. 2016 Aug;10(4):272-80.
- 5. J Diabetes Investig. 2016 Jan;7(1):56-69.
- 6. Frontiers in pharmacology. 2011;2:69-
- 7. Hormones (Athens). 2006 Oct-Dec;5(4):251-8.



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#### Cabbage Stir-fry with Coconut

MUTTAI KOSE VELLAI PORIYAL • Preparation time: 5-7 minutes • Cooking time: 7 minutes • Serves: 4

This is a traditional stir-fry from south India where such dishes are described as Poriyal. Much less oil is used than in the north and the cabbage is lightly cooked. Freshly grated coconut adds a very pleasant sweetness to it.

PREPARATION: Heat the oil in a skillet (frying pan) over medium heat. When the oil is hot, but not smoking, add the curry leaves, if using, mustard seeds and dal and fry for 30 seconds, or until the mustard seeds sputter and the lentils are golden. Add the cabbage, ginger, and green chiles and fry for 2 minutes, or until well-blended with the spices. Add the salt, then cover and cook over low heat for 3 minutes, or until the cabbage is tender but still crisp. Add the coconut, stir to combine, and remove from the heat to serve.

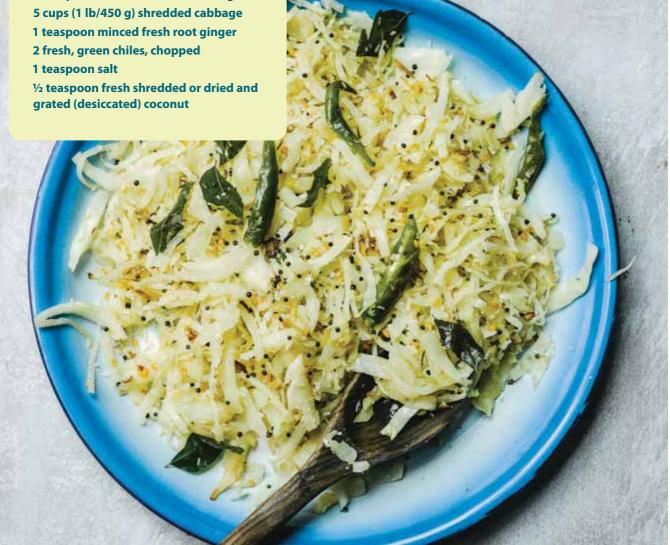
#### **INGREDIENTS**

2 tablespoons vegetable oil

4-6 curry leaves (optional)

1 teaspoon black mustard seeds

2 teaspoons urad dal lentils (black gram)



#### Split Red Dal

MASOOR DAL • Preparation time: 15 minutes, plus soaking time • Cooking time: 20-25 minutes • Serves: 4

Masoor or Egyptian lentils are arguably the most celebrated dal. There is a story about a temperamental chef in Lucknow who was so enraged with his employer—a Nawab addicted to chess—who kept the chef waiting and allowed the specially cooked dal to get cold. The chef left in a huff after pouring the dal on the stump of a tree. The next day, the dried wood sprouted fresh shoots and the Nawab realized what he had lost.

#### **INGREDIENTS**

- 1½ cups (9 oz/250 g) masoor dal (split red lentils), rinsed and drained
- 1 teaspoon ground turmeric
- 1 teaspoon ghee or vegetable oil
- 4 dried red chiles
- 1/2 teaspoon ajwain (carom) seeds, ground
- 1/3 cup (3/4 oz/20 g) chopped cilantro (coriander) leaves
- 4 fresh, green chiles, halved lengthwise and de-seeded

salt, to taste

PREPARATION: Put 5 cups (2 pints/1.2 liters) water in a large, heavy-based pan and add the masoor dal and turmeric. Bring to the boil and remove the scum from the surface with a slotted spoon, then reduce the heat, season with salt, and simmer for about 20 minutes, or until the dal is soft. Stir to break up the dal.

Heat the ghee or oil in a skillet (frying pan) over medium heat, add the dried red chiles and ground ajwain (carom) seeds and stir-fry for about 2 minutes, or until the chiles turn a shade darker. Pour over the dal, then cover and simmer for about 2 minutes. Add the chopped cilantro (coriander) and green chiles before serving.



#### Lentil & Vegetable Stew

SAMBAR • Preparation time: 30 minutes • Cooking time: 1 hour • Serves: 4

No Indian meal is complete without a lentil dish and the south-Indian sambar has an unrivalled pan-India following. This vegetable enriched lentil soup is served with snacks such as Dosa and Idli and, accompanied by rice, can also comprise a separate course in a formal meal. There are many regional variations and it would be a very brave man who would hazard to share an 'authentic' recipe. Until a few years ago, many families laboriously pounded sambar masala following a recipe handed down the generations. Now people are more likely to buy it.

**INGREDIENTS** 

scant 1 cup (7 oz/200 g) arhar/toor dal (yellow split pigeon peas), rinsed, soaked in water for 30 minutes, and drained

7 oz/200 g vegetable drumsticks or green beans

4 fresh, green chiles, slit lengthwise

4 oz/120 g medium shallots, left whole

11 oz/300 g medium tomatoes, quartered

1 teaspoon ground turmeric

1 teaspoon chili powder

1 teaspoon Sambar Masala

2 tablespoons tamarind extract

3 teaspoons chopped cilantro (coriander) leaves salt, to taste

**FOR TEMPERING** 

2 tablespoons vegetable oil

1 teaspoon urad dal lentils (black gram), rinsed

1 teaspoon cumin seeds

1 teaspoon coriander seeds

1 teaspoon black sesame seeds

1 teaspoon black mustard seeds

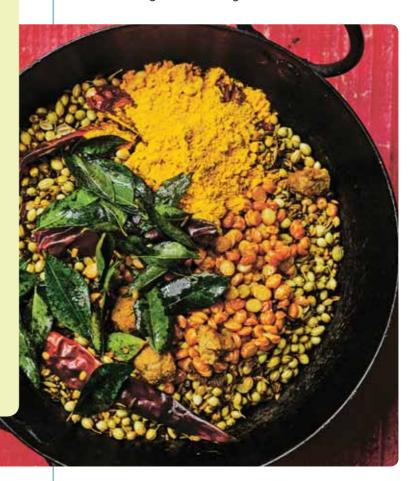
15-20 curry leaves

pinch of asafoetida

PREPARATION: Put the arhar/toor dal in a large, heavybased pan, add the vegetable drumstick or green beans, green chiles, shallots, tomatoes, turmeric, and chili powder, 4¼ cups (34 fl oz/1 liter) water, and season with salt to taste. Bring to a boil, then turn the heat down to medium-low, and simmer for 20 minutes. Stir in the Sambar Masala and continue to simmer, stirring occasionally, for 5 minutes.

Heat the oil for tempering in a skillet (frying pan) over medium heat, add the lentils and seeds, and cook for about 1 minute, until the seeds start to sputter. Add the curry leaves and asafoetida, and stirfry, until the leaves sputter. Pour this mixture over the sambar, then add the tamarind extract and stir well. Simmer on low heat for about 15 minutes.

Finally, add the chopped cilantro (coriander) and adjust the seasoning before serving.



#### Morels in Yogurt Sauce

KANAGUCCHI • Preparation time: 15 minutes • Cooking time: 15 minutes • Serves: 4-6

This recipe from Kashmir is one of the most sublime and subtly spiced of all Indian vegetarian gravy dishes. Morels, locally called gucchi, are a very expensive ingredient and are served on special occasions. They have a delicate flavor that can be easily killed by strong spices.

#### **INGREDIENTS**

2 cups (2 oz/50 g) dried morels

3 tablespoons vegetable oil or ghee

1 onion, finely sliced (optional)

5 cloves garlic, peeled and crushed

1 cup (8 fl oz/250 ml) yogurt, whisked

1 teaspoon ground ginger

2 teaspoons ground fennel

1/2 teaspoon cumin seeds, crushed

1 teaspoon garam masala

salt, to taste

cilantro (coriander) leaves, to garnish

PREPARATION: Soak the morels in hot water for 1 hour, then drain and wash thoroughly to remove any grit. Squeeze dry and set aside.

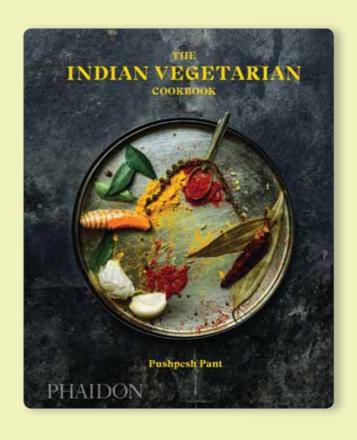
Heat the oil or ghee in a large, heavy-based pan over medium heat. Add the onion, if using, and fry for 2-4 minutes, until translucent but not browned. Add the garlic and fry for 1 minute, then add the morels, seasoning with salt to taste. Cook, shaking the pan from time to time, for about 8-10 minutes, until all the water has evaporated. Add 1 cup (8 fl oz/250 ml) water and bring to the boil. When boiling, pour in the whisked yogurt and add the ginger and fennel, stirring briskly to ensure that the gravy does not curdle. Reduce heat to low and simmer for about 10 minutes, until the gravy reaches a thin, custard-like consistency. Sprinkle over the crushed cumin seeds, garam masala, and cilantro (coriander) leaves to 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.

Reprinted from The Indian Vegetarian Cookbook (Phaidon 2019).

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Venotone

Xyliwhite Mouthwash

Youthful Collagen

#### Pet Care

Cat Mix

Dog Mix

#### **Probiotics**

Bifido GI Balance

FLORASSIST® Balance

FLORASSIST® GI with Phage Technology FLORASSIST® Heart Health

FLORASSIST® Immune Health

FLORASSIST® Mood FLORASSIST® Nasal

FLORASSIST® Oral Hygiene FLORASSIST® Prebiotic FLORASSIST® Throat Health

Jarro-Dophilus® for Women

Theralac® Probiotics
TruFlora® Probiotics

#### **Skin Care**

Advanced Peptide Anti-Glycation Serum

Advanced Growth Factor Serum

Advanced Hyaluronic Acid Serum

Advanced Lightening Cream Advanced Peptide Hand Therapy

Advanced Triple Peptide Serum Advanced Under Eye Serum with Stem Cells

All-Purpose Soothing Relief Cream

Amber Self MicroDermAbrasion

Anti-Aging Face Oil

Anti-Aging Mask

Anti-Aging Rejuvenating Face Cream

Anti-Aging Rejuvenating Scalp Serum

Anti-Oxidant Serum with

Blueberry & Pomegranate Extracts

Anti-Oxidant Facial Mist Hydrator

Collagen Boosting Peptide Serum

Cucumber Hydra Peptide Eye Cream

DNA Support Cream Environmental Support Serum

Eye Lift Cream

Face Rejuvenating Anti-Oxidant Cream Hyaluronic Facial Moisturizer

Hyaluronic Oil-Free Facial Moisturizer

Hydrating Anti-Oxidant Facial Mist

Hydroderm

Lifting & Tightening Complex
Melatonin Advanced Peptide Cream

Melatonin Cream

Mild Facial Cleanser

Multi Stem Cell Hydration Cream Multi Stem Cell Skin Tightening Complex

Neck Rejuvenating Anti-Oxidant Cream

Rejuvenex® Body Lotion

Rejuvenex® Factor Firming Serum Renewing Eye Cream

Resveratrol Anti-Oxidant Serum

Shade Factor™ Shade Factor™ Sunscreen Lotion

Shade Factor™ Sunscreen Spray

Skin Care Collection Anti-Aging Serum Skin Care Collection Body Lotion

Skin Care Collection Day Cream

Skin Care Collection Night Cream

Skin Firming Complex Skin Lightening Serum

Skin Restoring Ceramides Skin Stem Cell Serum

Ultra Eyelash Booster

Skin Tone Equalizer Stem Cell Cream with Alpine Rose

Tightening & Firming Neck Cream Triple-Action Vitamin C Cream

Ultimate MicroDermabrasion Ultimate Peptide Serum

Benfotiamine with Thiamine Beta-Carotene BioActive Complete B-Complex **Biotin** Buffered Vitamin C Powder Fast-C® and Bio-Quercetin Phytosome Gamma E Mixed Tocopherol Enhanced

Ascorbyl Palmitate

Ultra Rejuvenex®

Vitamin C Serum

Vitamin D Lotion

Vitamin K Cream

Circadian Sleep

Glycine

L-Tryptophan

Melatonin IR/XR

**Vitamins** 

Melatonin

Ultra RejuveNight® Ultra Wrinkle Relaxer

Under Eye Refining Serum

Under Eye Rescue Cream Vitamin C Lip Rejuvenator

Bioactive Milk Peptides

Enhanced Sleep with Melatonin

Fast-Acting Liquid Melatonin

Optimized Tryptophan Plus Quiet Sleep Quiet Sleep Melatonin

Enhanced Sleep without Melatonin

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

No Flush Niacin Optimized Folate (L-Methylfolate)

Pantothenic Acid (Vitamin B-5)

Pyridoxal 5'-Phosphate Caps

Super Absorbable Tocotrienols

Super K

Super K Elite Super Vitamin E Vitamin B6

Vitamin B12

Vitamin C and Bio-Quercetin Phytosome

Vitamin D3

Vitamin D3 with Sea-Iodine™

#### Vitamins D and K with Sea-lodine™ Weight Management & Body Composition

7-Keto® DHEA Metabolite

Advanced Anti-Adipocyte Formula Advanced Appetite Suppress

AMPK Metabolic Activator CalReduce Selective Fat Binder **DHEA Complete** 

Garcinia HCA

**HCActive Garcinia Cambogia Extract** 

Integra-Lean®  $Mediterranean\ Trim\ with\ Sinetrol^{\mbox{\tiny TM}}\ -XPur$ 

Optimized Irvingia with Phase 3™ Calorie

Control Complex

Optimized Saffron with Satiereal® Super CLA Blend with Sesame Lignans Waist-Line Control™

Wellness Code Appetite Control

Women's Health Enhanced Sex for Women 50+ Breast Health Formula

Femmenessence MacaPause® Estrogen for Women

Menopause 731™ Progesta-Care®

Super-Absorbable Soy Isoflavones

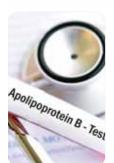


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## WHAT'S INSIDE

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# **LifeExtension**°



# 7 WHY BLOOD TESTS ARE NOT SAVING MORE LIVES

Healthy levels of **apolipoprotein B** can *decrease* coronary artery disease risk by **90%** and help reduce arterial plaque. Conventional blood tests fail to test for **apolipoprotein B** and other important vascular markers.



# 36 INFLAMMATION-FIGHTING PLANTS

Natural compounds found in **ginger** and **turmeric** reduce **inflammation** that contributes to degenerative aging.



#### 54 HEART FAILURE AND RIBOSE

Supplementing with **D-ribose** *reduces* symptoms of **heart failure**. Read about a new, multi-pronged approach for heart failure patients.



# 26 ENHANCING BENEFITS OF VITAMIN K

**Vitamin K** has been shown to lower death rate from *any* cause by **26%**. A new broad-spectrum formula helps support bone and arterial health.



# 47 COQ10 REVERSES SKIN AGING

**CoQ10** restores **mitochondrial function** in aged skin and reduces wrinkles by nearly **25%** while promoting smoother skin by up to **82%**.



## 66 HOW WHEY PROTEIN FIGHTS AGING

**Whey protein** impedes muscle loss, reduces frailty, limits weight gain and promotes lean muscle mass, all critical factors for aging adults.