

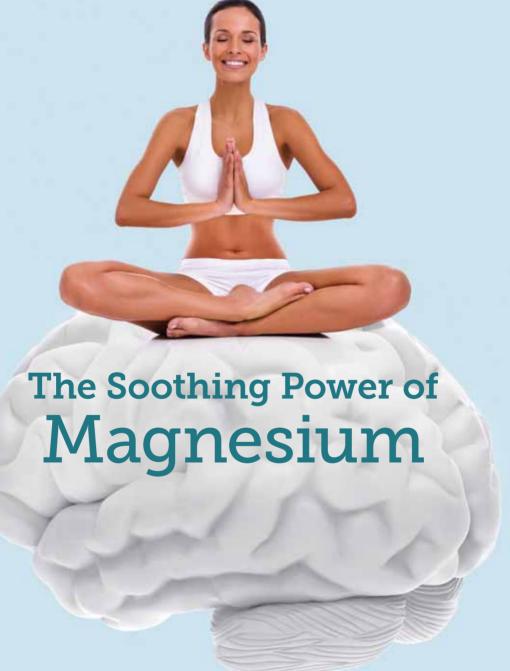
The Science of a Healthier Life®

LifeExtension.com

August 2022



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Surging Increase of Fatty Liver Disease

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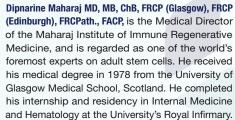
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A Silent Disorder Goes Epidemic



WILLIAM FALOON

In 1999, I reviewed an article for publication in this magazine that described a liver disease that I had not heard of.

The authors told me this "silent" disorder would lead to widespread illness and premature death.

Move forward 20 years, and nonalcoholic fatty liver disease has emerged as the leading cause of chronic liver disease.1

The underlying culprits are **obesity** and **insulin** resistance.2,3

In 1980, obesity was rare in the United States, affecting only 13% of adults according to some reports.4

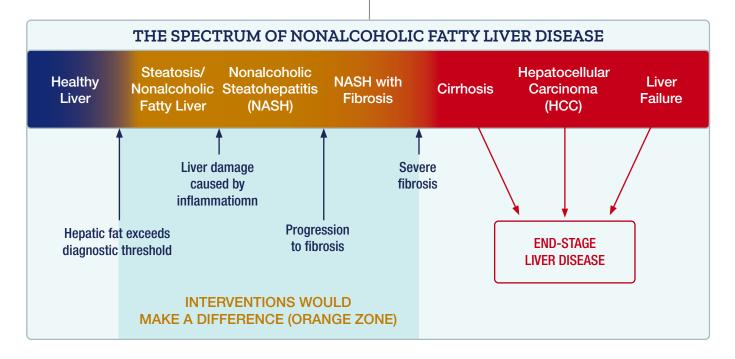
In the most recent study period (2017-2018), a startling **42.4%** of American adults were **obese**.⁵ Nonalcoholic fatty liver disease afflicts up to 75% of overweight adults, approximately 75% of type II diabetics, and about 90% of obese individuals.2,6

People with **nonalcoholic fatty liver disease** are seeking our assistance in combating its multiple morbidities.

The most effective treatment for early-to-midstage nonalcoholic fatty liver disease (NAFLD) is weight loss. Adjuvant approaches can also help.

Once NAFLD fatty liver disease advances to cirrhosis, the only long-term treatment option is a liver transplant.8

An article in this month's issue describes ways to help protect liver health.



Nonalcoholic fatty liver disease or NAFLD begins when excess fat accumulates in liver cells. Over time, it can progress to advanced fibrosis and cirrhosis.9

As the condition is generally asymptomatic or has very mild symptoms initially, many people do not know they have it unless a blood test detects abnormal levels of liver enzymes.

When detected in time, NAFLD is a reversible condition, meaning those afflicted can avoid progressing to advanced-stage disease that requires a liver transplant.10

Fatty Liver Disease and Cancer

Liver failure is the most common manifestation of advanced nonalcoholic fatty liver disease (NAFLD).6

Another NAFLD risk is **primary liver cancer** called hepatocellular carcinoma. 13 Five-year survival for this malignancy is less than 20%.14

A study published in 2018 evaluated 296,707 NAFLD patients and compared them with 296,707 matched **controls**. (This is a huge number of people to evaluate.)

This study found a more than 7.6-fold higher risk of hepatocellular carcinoma among NAFLD patients compared to controls.¹⁵

NAFLD patients with cirrhosis had an even greater incidence of hepatocellular carcinoma.

A medical-record review from this study found that approximately 20% of NAFLD patients with hepatocellular carcinoma had no evidence of cirrhosis.

This finding reveals **hepatocellular carcinoma** (HCC) risk in NAFLD patients who have not advanced to latestage (cirrhosis) liver disease.15

This means people can skip past cirrhosis of the liver and develop primary liver cancer.

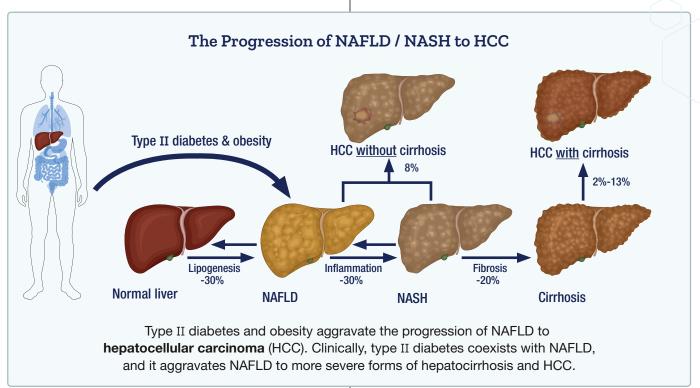
Diagnosing NAFLD

If you have a history of any of the following medical conditions you are at much greater risk of having some degree of NAFLD:6

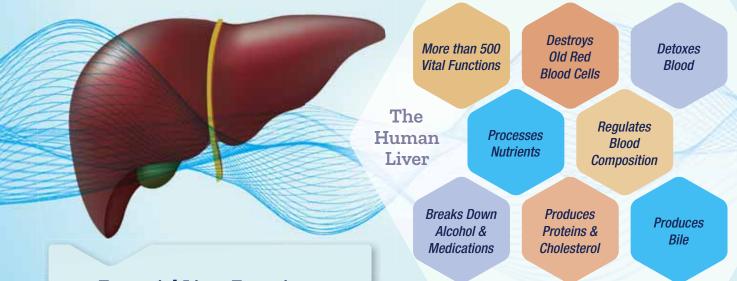
- **Excess body weight (abdominal obesity)**
- Insulin resistance or type II diabetes
- Metabolic syndrome

Most patients with NAFLD, especially early in the disease, are asymptomatic, and are only identified when their disease becomes advanced.

Abnormal liver test results are frequently the earliest evidence of NAFLD, which emphasizes the value of routine laboratory testing to identify asymptomatic NAFLD patients.



THE PROGRESSION OF NAFLD / NASH TO HCC BY MDPI, BASEL, SWITZERLAND CC BY 4.0



Essential Liver Functions

More than **500** processes require healthy liver function, such as:11

- Making bile to carry away waste, break down fats during digestion
- Producing albumin and other blood plasma proteins
- Converting excess glucose into glycogen for storage
- Making proteins to carry fats through the body
- Balancing blood glucose levels
- Regulating blood levels of amino acids
- Processing hemoglobin
- · Converting poisonous ammonia to
- · Clearing the blood of drugs and other toxic substances
- Regulating blood clotting
- Resisting infections by making immune factors and removing bacteria from the bloodstream
- Clearing bilirubin

In advanced liver disease, when liver function is impaired, the result is systemic havoc that may manifest as fatigue, abdominal pain, and fluid accumulation, along with increased cardiovascular and diabetic risks.12

In particular, the liver enzymes alanine aminotransferase (ALT) and aspartate aminotransferase (AST) can be elevated in up to 50% of patients with NAFLD.16

Triglycerides can also be elevated in NAFLD but may not be in the early course of disease.

These three screening tests (ALT, AST, and triglycerides) are included in Life Extension®'s comprehensive CBC/Chemistry Panel for only \$35.

The CBC/Chemistry Panel can be ordered 24-hours a day by calling 1-800-208-3444 or logging on to: www. LifeExtension/blood

Most of you live close to a drawing station so you can walk in for the CBC/Chemistry Panel at your convenience. Fast for eight hours before your blood draw.

Diagnosing Advanced Liver Disease

Symptoms of advanced NAFLD and cirrhosis can include loss of appetite, pain in the upper right abdomen, fatigue, muscle loss, and/or fluid accumulation in your belly (ascites).17

A physician's exam can reveal an enlarged liver, spleen, and other clinical signs with advanced NAFLD.

There are a variety of **imaging** (MRI, ultrasounds, CT) and advanced blood tests that can detect suspicious liver abnormalities. These diagnostic tests are often useful to identify the degree of fatty liver damage.

One such blood test for those at risk is called NASH FibroSure. It is designed to assess what degree of liver damage has occurred.

The cost of this blood test is a prohibitive \$595. For those with symptoms/signs of NAFLD, it might be best to have your physician prescribe this test and hopefully have it covered by your health insurance.

Nonalcoholic Fatty Liver Disease Liver cells **HEALTHY LIVER** Fat cells **FATTY LIVER**

For definitive diagnosis, liver biopsies can establish the stage, and degree of structural damage in patients with NAFLD.

Reversing Damage Caused by NAFLD

If caught in time, healthy dietary and lifestyle choices can stop and help reverse fatty accumulation in your liver cells.

The single most effective way to control or reverse NAFLD is weight loss.

Those who wait for **NAFLD** to reach advanced stages have few treatment options beyond being placed on a liver transplant waiting list.

Prevention remains the most effective strategy!

For longer life,

William Faloon, Co-Founder

Life Extension®



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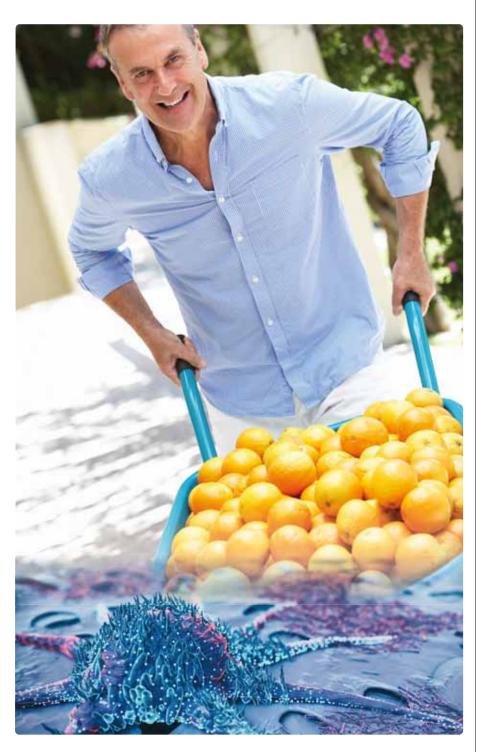
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In the News



Greater Vitamin C Intake Linked to **Lower Cancer Risk**

A comprehensive review that included 57 meta-analyses evaluating the association between vitamin C consumption and cancer found that higher intake of the vitamin was linked to a lower risk of many cancers, according to an article published in Frontiers in Nutrition.*

When digestive system cancers were evaluated, the highest intake of vitamin C, compared to the lowest, was associated with a 42% decrease in the risk of esophageal cancer, a 34% lower risk of stomach cancer and a 30% lower risk of pancreatic cancer.

Additionally, among individuals whose vitamin C intake was highest, lung cancer risk was 17% lower than in the group with the lowest intake.

Editor's Note: "Vitamin C consumption was associated with lower incidence of...total cancer occurrence," the authors stated.

* Front Nutr. 2022 Jan 20;8:812394.

Vitamin D Supplementation Reduces Palliative Cancer Patients' Need for Opioids

A reduced need for pain relief as well as less cancer-related fatigue were found in palliative cancer patients receiving supplemental vitamin D, compared to a placebo group, the journal Cancers reported.*

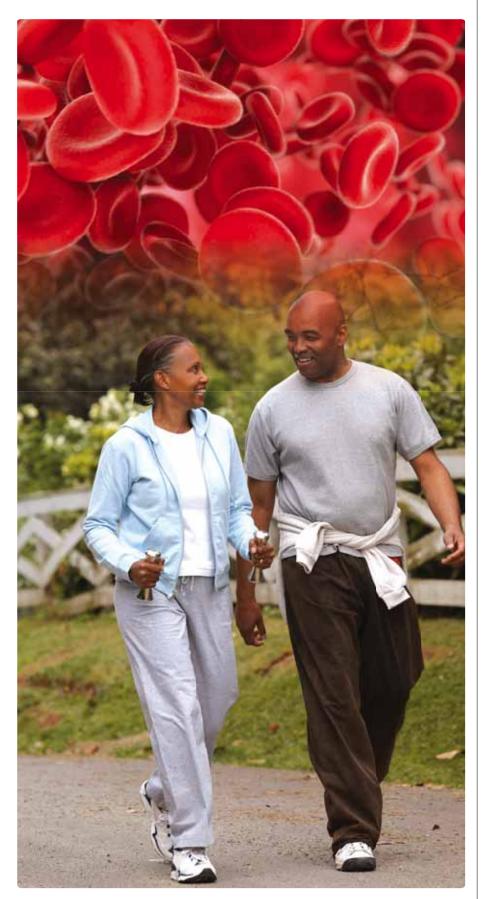
At the start of the double-blind. randomized placebo-controlled study, 244 cancer patients, all deficient in vitamin D, were given either high-dose vitamin D (4,000 IU/day) or a placebo for 12 weeks. Their opioid doses were assessed throughout the 12 weeks as a measurement of pain.

Those taking vitamin D had significantly lower increases in opioid doses during the study. This indicates a reduced need for pain relief. They also reported less cancer-related fatigue compared to the placebo group.

Editor's Note: "Correction of vitamin D deficiency may have positive effects on pain and fatigue in palliative cancer patients," the researchers concluded.

* Cancers. 2021; 13 (15): 3707.





Higher CoQ10 Levels Benefit Older Folks

A study found greater physical capacity and a reduction in blood factors related to cardiovascular disease among participants with higher plasma levels of CoQ10, a factor in the production of energy within the mitochondria of the cells, according to an article in Antioxidants (Basel).*

Participants included men and women aged 65 and older, with a majority being women.

Higher CoQ10 levels were significantly associated with lower cardiovascular disease risk. Having greater levels of CoQ10 also correlated with better physical activity.

Authors suggest the combination of CoQ10 with physical activity as an important therapy for the prevention of sarcopenia and the maintenance of physical capacity.

Editor's Note: "It is probable that supplementation with CoQ10 could improve physical capacity in addition to the known effects in the cardiovascular system. These relationships suggest that CoQ10 could be considered an important component for maintaining independence and health in aged individuals," the authors stated.

* Antioxidants (Basel). 2022 Jan 29;11(2):279.

Light to Moderate Coffee Drinking Can be Good for You

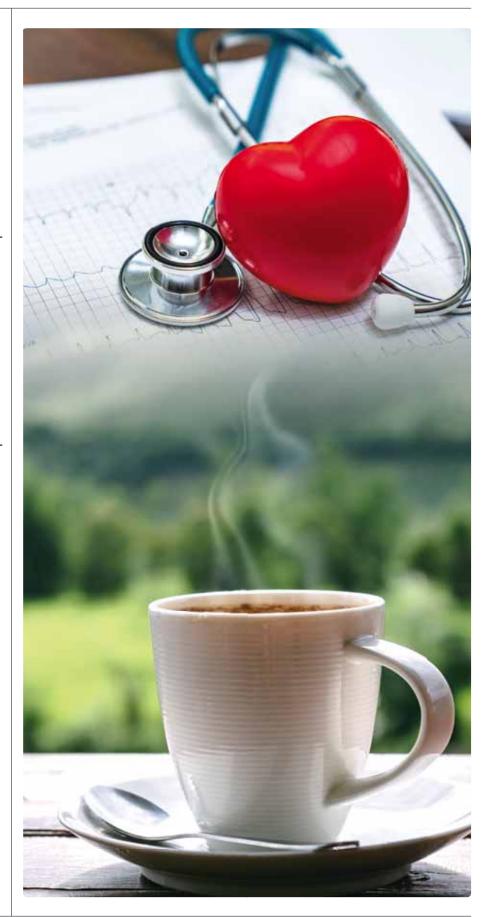
The European Journal of Preventive Cardiology published the largest observational study to date showing that drinking coffee was associated with a reduced risk of all-cause mortality, cardiovascular mortality, and stroke.*

For 11 years, researchers tracked data from 468,629 individuals who did not have heart disease at the start of the study.

Compared to non-coffee drinkers, light-to-moderate drinkers (0.5-3 cups per day) were found to have 12% lower risk of all-cause mortality, 17% lower risk of dying from cardiovascular disease, and 21% reduction in the incidence of stroke.

Editor's Note: The researchers concluded that, "This favorable impact might be partly explained by lower Arterial Stiffness Index and subclinical alterations in cardiac structure and function."

* Eur J Prev Cardiol. 2022 May 6;29(6): 982-991.



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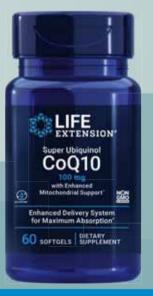


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BY STUART SANCHEZ

In a **2022** report, the **American Psychological Association** noted that **73%** of U.S. adults say they feel "overwhelmed by the number of crises facing the world right now."¹

Chronic stress has been linked to:2,3

- Anxiety and depression
- Cardiovascular disease
- Obesity
- Menstrual problems
- Sexual dysfunction
- Gastrointestinal problems
- Hair and skin conditions

People suffering from **stress-related** symptoms, depression, or anxiety often have <u>low</u> blood levels of **magnesium**.⁴

In a follow-up analysis of a clinical study, it was found that approximately **44**% of participants screened for stress were **magnesium** deficient.⁵

Preclinical studies found that a specific <u>form</u> of magnesium called **magnesium acetyl taurate** *increased* brain tissue levels of magnesium more effectively than other forms of magnesium tested.^{6,7}

Magnesium acetyl taurate was recently tested in women with premenstrual syndrome over a duration of three menstrual cycles, who had inadequate magnesium intake from food.8

Researchers found improved scores for **symptoms of stress** including nervous tension, anxiety, irritability, headache, fatigue, and depression in women supplementing with **magnesium acetyl taurate** twice daily.

The Magnesium-Stress Connection

Decades ago, researchers first noticed a link between magnesium and stress.9,10

Many of the frequently-reported symptoms of stress fatigue, irritability, anxiety, headache, and upset stomach-are the same symptoms commonly found in patients with magnesium deficiency.4

As scientists explored the connection, they discovered that it goes both ways:4,11

- The body's responses to stress lead to a loss of magnesium in the urine, over time creating a magnesium deficiency.
- Low magnesium levels make people more susceptible to stress, increasing release of stress hormones like adrenaline and cortisol, which can be harmful if their levels remain elevated.

This can create a vicious circle. As low magnesium levels make the effects of stress more severe, that further reduces magnesium levels, making people even more susceptible to stress, and so on.4

On the other hand, maintaining adequate magnesium levels helps protect against stress and other conditions.

Magnesium is an important cofactor for the synthesis of serotonin, a neurotransmitter closely linked to positive

mood and feelings of calm. Most antidepressant and antianxiety medications act at least partially by modulating serotonin neurotransmission.4,11

Magnesium can also inhibit release of the stress hormone cortisol from the adrenal gland.12

Unique Brain-Targeted Magnesium

Scientists identified a specific form of magnesium that, taken orally, can rapidly increase brain levels of magnesium.

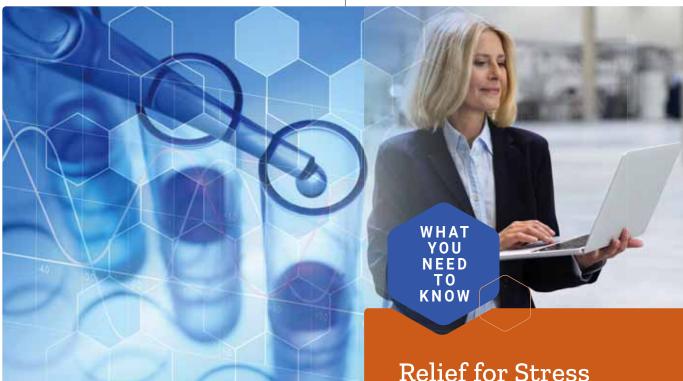
Magnesium acetyl taurate is magnesium combined with a form of the amino acid taurine. The combination makes it easier for the magnesium to cross the blood/ brain barrier.

Studies have found that this form of magnesium was more easily *absorbed* into the **brain** than the other forms of magnesium tested.

In one study, rats were given either magnesium acetyl taurate or two other common forms of magnesium, magnesium sulfate and magnesium oxide.⁶ Brain tissue and blood magnesium levels were significantly higher after eight hours in the group receiving magnesium acetyl taurate.

Another preclinical study pitted magnesium acetyl taurate against four other common forms of magnesium: magnesium sulfate, oxide, citrate, and malate.7





Again, brain levels of magnesium were significantly higher in the group receiving magnesium acetyl taurate than with control or any of the other forms of magnesium tested.

This study also found that magnesium acetyl taurate was associated with decreased anxiety indicators in rodents.

This form of magnesium has also shown promise in an early study in humans.8

Easing Stress

Researchers enrolled adult women with symptoms of premenstrual syndrome who had inadequate magnesium intake in their diet.8 Premenstrual syndrome can cause symptoms similar to those of stress.

After receiving 385 mg of magnesium acetyl taurate twice daily over a series of three consecutive menstrual cycles, scores for numerous symptoms were significantly reduced, including those for nervous tension, anxiety, irritability, headache, fatigue, and depression.

Intake of this form of magnesium may promote calm and help people cope with symptoms of stress and anxiety, whatever the cause.

Relief for Stress and Anxiety

- Chronic stress is associated with cardiovascular disease, obesity, anxiety, depression, and more.
- The close link between stress and magnesium in the body has gained interest by researchers. Having low magnesium levels increases susceptibility to stress and its negative health consequences.
- One specific form of magnesium, magnesium acetyl taurate, was found to be superior to other forms tested in raising brain levels of magnesium.
- In a human study, taking 385 mg of magnesium acetyl taurate twice daily reduced premenstrual syndrome symptoms that are similar to those of stress, including anxiety, irritability, headache, fatigue, and depression.



The Importance of Magnesium

Magnesium is an important essential mineral in the body.4,5

It is involved in most major metabolic and biochemical processes and serves as a cofactor ("helper molecule") for more than 300 different enzymatic reactions.^{4,5}

Low magnesium has been tied to numerous health problems, including cardiovascular disease, diabetes, osteoporosis, depression, and anxiety. 11,13

Suboptimal levels of magnesium are more common than most people realize.

It has been estimated that 64% of all men and 67% of women in the U.S. have inadequate dietary intake of magnesium. More than 80% of people over the age of 71 have an inadequate dietary intake of magnesium.14

Making matters worse, excessive sodium intake, high alcohol and caffeine intake, and some medications (including proton pump inhibitors for acid reflux) can further contribute to lower magnesium levels.4

Summary

Chronic stress can be extremely harmful, increasing the risk cardiovascular disease, obesity, and other conditions.

The close link between stress and levels of magnesium in the body has sparked the interest of researchers.

A specific form of magnesium called magnesium acetyl taurate improves the bioavailability of magnesium and was found to be more effective at raising brain levels of this essential mineral than the other forms of magnesium tested. •

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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1. Sleep Med Rev. 2005 Feb;9(1):41-50. 2. Lancet. 1995 Aug 26;346(8974):541-4. 3. Neurol Res. 2017 Jun;39(6):559-65.

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Calm-Mag provides **50 mg** of elemental magnesium in each daily capsule. It may be used with other forms of magnesium.

References: 1. Nutrients. 2020 Nov 28;12(12). 2. Synapharm - Company supplied data. 2021.

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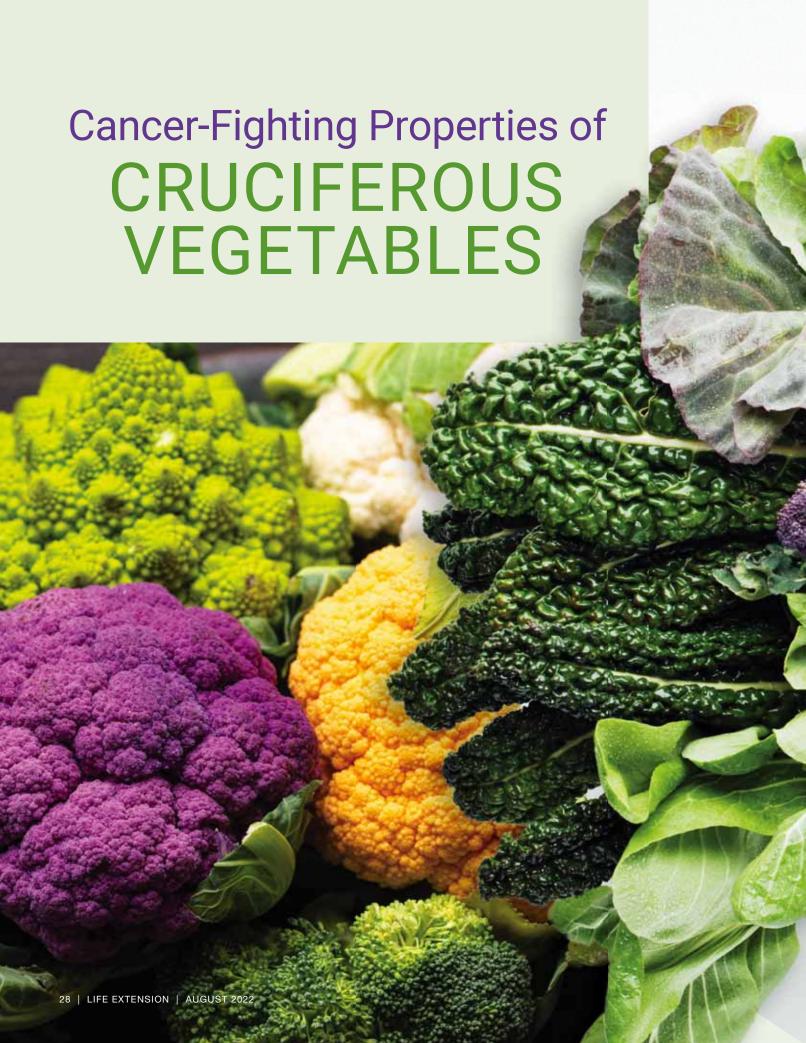
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The anti-cancer effects of **cruciferous vegetables** like **broccoli** have long been researched.

Studies show that *higher* intake of these **vegetables** is associated with a reduced risk of many cancers.^{1,2}

Two established cancer-fighting plant compounds found only in **cruciferous vegetables** are:

SULFORAPHANE and DIM (3,3'-DIINDOLYLMETHANE) Research suggests these plant compounds target **six** different **pathways** to impede the development of cancer and slow progression of existing cancer.

DIM is readily **bioavailable** to the body, but it is challenging to obtain **sulforaphane** from mature broccoli.

This article updates readers on the cancer-fighting properties of **sulfora-phane** and how to deliver it to the small intestine for *systemic* **absorption**.

Benefits of Cruciferous Vegetables

The cruciferous vegetables include:

- Broccoli
- Cabbage
- Cauliflower
- Kale
- Brussels sprouts
- Collard greens
- Bok choy
- Arugula
- Watercress
- Radishes.

Vegetables in this family contain a wide range of nutrients, including flavonoids, carotenoids, and minerals.^{3,4}

Dietary intake of **cruciferous vegetables** like broccoli and cauliflower has been demonstrated to reduce **cancer risk**.⁵⁻⁷

High intake of **cruciferous vegetables** is also associated with <u>better</u> **survival rates** in patients already diagnosed with cancer.^{8,9}

Two unique compounds present in the cruciferous family, **sulforaphane** and **DIM (3,3'-diindolylmethane)**, modulate pathways involved in cancer development and progression.¹⁰⁻¹³

They work in multiple ways to block the development of cancer *and* to make it difficult for cancer cells to grow and survive.

Promising Sulforaphane Studies

A study published in **2021** evaluated the effect of **sulforaphane** on human **glioblastoma** cells.¹⁴ Glioblastoma is an aggressive cancer of the brain or spinal cord that is often incurable.

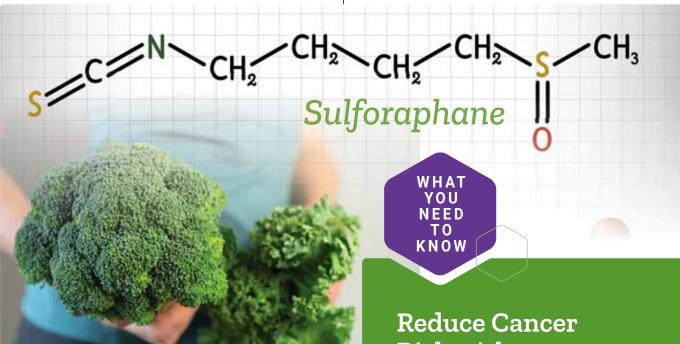
Sulforaphane **stopped the growth of cancer cells** in their tracks. It also caused the cancer cells to begin to **die off**.

Similar results have been seen in other human cancer cell lines, including **prostate** and **breast** cancers. 15-21

Sulforaphane has shown promise in animal studies as well.²² In one, rats were given a potent carcinogen. In the group of animals that did *not* receive sulforaphane, **68**% developed tumors.

In rats given **sulforaphane**, just **39**% developed tumors. And in treated rats that *did* develop tumors, the tumors were smaller *and* slower growing.





One recent study indicates that **sulforaphane** may be effective when used alongside certain conventional anti-cancer drugs.

Scientists tested stomach cancer cells, some of which were sensitive to the anti-cancer drug lapatinib and some which were resistant to the drug.23

Combining lapatinib and sulforaphane stopped the growth and spread of both types of cancer and killed off the cells—even cells previously **resistant** to the drug.

Improving the Delivery of Sulforaphane

Sulforaphane is an unstable compound that rapidly degrades into non-active substances if it isn't quickly absorbed or if the vegetable is cooked. Interestingly, it isn't even present in cruciferous vegetables themselves.

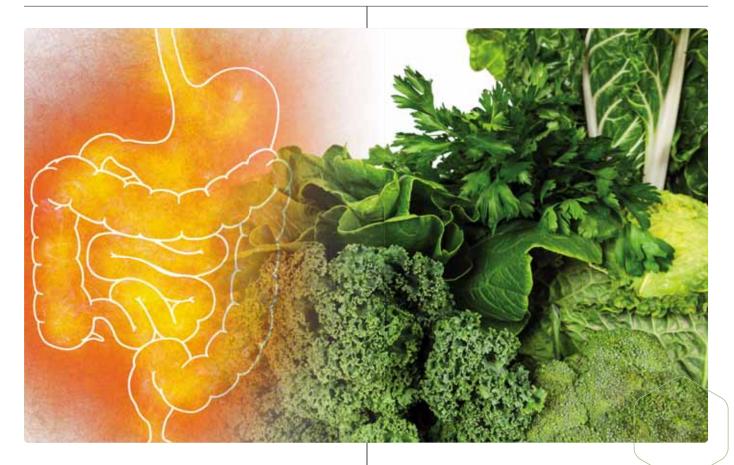
Instead, a precursor called glucoraphanin is stored inside the cells of these plants. In a separate compartment in these cells there is an enzyme called myrosinase.

When these two are combined, the myrosinase converts the glucoraphanin into sulforaphane.

This is what happens when the vegetable is eaten raw. During digestion, sulforaphane is formed. Then, before it degrades, it can be absorbed within the small intestine.2,5,24

Risk with Cruciferous Veggies

- Cruciferous vegetables include broccoli, cabbage, kale, cauliflower, Brussels sprouts, and more. Studies show that diets high in these vegetables protect against many forms of cancer.
- Two nutrients derived *only* from cruciferous vegetables have been shown to possess potent cancerfighting abilities: sulforaphane and DIM (3,3'-diindolylmethane).
- These compounds limit the ability of cancer cells to grow, divide, and spread, and they cause cancer cells to die off.
- Sulforaphane is unstable. Scientists have solved this problem by packaging a sulforaphane precursor with an enzyme that only converts it into sulforaphane in the body. That way it can be rapidly absorbed in the digestive tract.



But maximizing these anti-cancer benefits would require the consumption of massive amounts of raw cruciferous vegetables or cruciferous sprouts.

The challenge for scientists was to find a way to deliver **glucoraphanin** and **myrosinase** separately to the small intestine.

One group of scientists came up with an ingenious solution that copies nature.

They isolated **glucoraphanin** and **myrosinase** from broccoli, then developed a unique delivery system that keeps them separate, *just the way plants do*.

A dual-layered tablet was given an enteric coating to prevent its ingredients from being released until it reaches the small intestine.

With this delivery system, the compound **gluco-raphanin**, and the enzyme **myrosinase** meet and mix in the small intestine. There, they come together to create **sulforaphane**, just as nature planned.

The results have been striking. Research at the Johns Hopkins University School of Medicine demonstrated that **sulforaphane** levels from this **glucoraphanin-myrosinase** mix are **three** to **four times** more bioavailable (absorbable) than those created by glucoraphanin supplementation alone.²⁵

Six Anti-Cancer Mechanisms

DIM and **sulforaphane** act in **SIX** different ways to help prevent the development of cancer and to slow the spread of existing cancer.

IMPEDING CANCER CELL GROWTH

Both compounds can arrest the **cancer cell cycle**, interfering with the ability of tumor cells to grow. 14-17,19,21

They also block the formation of new blood vessels in tumors, starving them of nutrients and oxygen. 16,26-28

Type 2 transglutaminase is a cancer cell survival protein in several forms of cancer. A study published in **2022** showed that **sulforaphane** attaches itself to this protein, *blocking* its activity.²⁹ This further shuts down cancer cells' ability to survive.

KILLING OFF CANCER

When normal cells become damaged, they're supposed to die off through a process known as **apoptosis** (programmed cell death).

Many cancer cells evolve to *shut off* apoptosis. **DIM** and **sulforaphane** have been found to turn apoptosis *back on*, initiating **cancer cell death**. 16,18,20

REDUCING HARMFUL **EPIGENETIC CHANGES**

Cancer can also be caused by epigenetic changes, which happen when genes are turned "on" or "off," making them active or inactive.

Both sulforaphane and DIM reduce epigenetic changes that contribute to tumor development. 11,30-32

STIMULATING CELLULAR PROTECTION

Nrf2 is a protein that regulates cellular protection. Activating Nrf2 turns on different genes that protect cells against stress and injury.33

For example, Nrf2 activates enzymes that help eliminate mutagens and other toxins.34,35

Many of sulforaphane's benefits stem from its activation of the Nrf2 pathway.33

REDUCING CHRONIC INFLAMMATION

DIM and sulforaphane both inhibit the action of nuclear factor-kappa B (NF-kB), a regulatory protein that contributes to chronic inflammation.³⁶⁻³⁸

This anti-inflammatory effect helps prevent cancer and other chronic health conditions.



FIGHTING HORMONE-DRIVEN **CANCER STIMULATION**

Some prostate and breast cancers are stimulated by forms of **estrogen**. DIM shifts estrogen balance away from an estrogen form that promotes tumor cell growth and toward a form that inhibits it.39,40

In women with a history of breast cancer, daily DIM intake increased the proportion of "good" estrogen and reduced the forms linked to faster cancer progression.^{39,40}

In men, higher estrogen levels are associated with prostate enlargement and cancers. In a cell study, DIM prevented estrogen-induced stimulation of prostate cancer cells.41

Summary

Many studies show that higher intake of cruciferous vegetables protects against cancer.

These vegetables are a source of two compounds that have demonstrated strong anti-cancer activity: **DIM** (3,3'-diindolylmethane) and sulforaphane.

Both compounds can stop cancer cell development and growth in their tracks.

Sulforaphane intake has previously been problematic because it is so unstable. A novel two-layer formula enables the nutrient to be bioavailable for absorption into the bloodstream. •

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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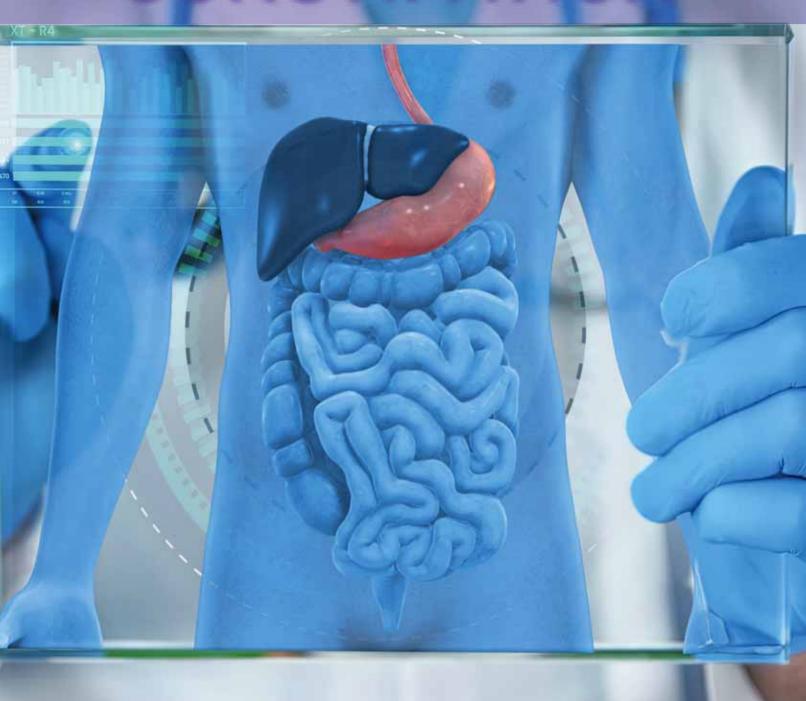
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BANISH CONSTIPATION



and ITS HEALTH RISKS

BY MICHAEL DOWNEY

No one likes to talk about it, but about **15%** of Americans experience **constipation** on an ongoing basis. It can be unpleasant and uncomfortable.

Constipation increases with age, affecting approximately **34**% of women and **26**% of men 65 years and older.¹

The risks go well beyond discomfort.

A 2019 study found that constipation is associated with a higher risk of **stroke**, **coronary heart disease**, and **all-cause mortality**.²

Changes to the gut microbiota associated with constipation might be related to atherosclerosis, a rise in blood pressure, and cardiovascular events.³

There's a unique way to counter those changes: a targeted **probiotic** strain that reduces constipation.⁴

For those who need immediate relief combinations of magnesium + vitamin C or potassium + magnesium + vitamin C are effective while providing the body with healthy nutrients.



Health Issues Linked to Constipation

Constipation can make life miserable. It's associated with a higher risk of heart disease, stroke, and overall mortality, as well as to a disrupted microbiota that may have negative effects throughout the body.^{2,6}

Constipation is also linked to a higher risk of cancer. This may be due to increased inflammation or prolonged contact between stool carcinogens and the tissue lining the colon.5

A Probiotic Solution

Scientists focused on the beneficial bacteria known as **probiotics** as a possible solution.

After investigating more than 2,000 probiotic strains, scientists identified several strains derived from yogurt produced in New Zealand that had probiotic value and the ability to survive conditions similar to those in the human digestive tract.7

Of these, one **specific strain** decreased **colonic transit** time, the time it takes for food to move through the colon. This probiotic is *Bifidobacterium lactis* HN019.4

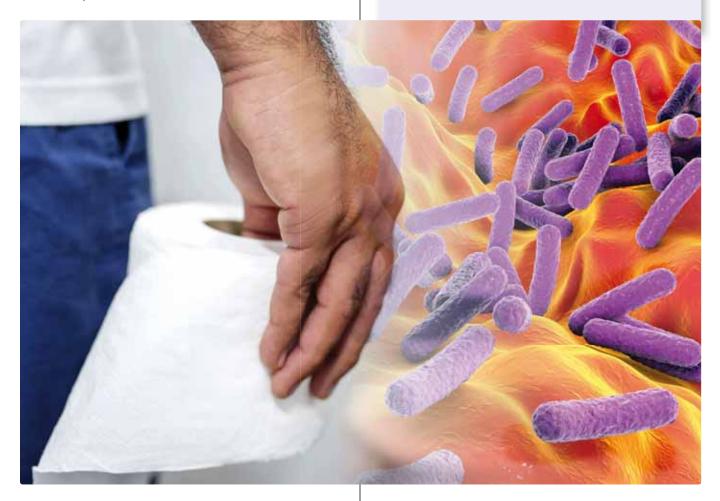
By moving stool along faster, scientists believed that this probiotic could help relieve existing constipation and many symptoms often associated with it, such as abdominal pain and gas.4

Clinical Study of Constipation

In a clinical trial of patients with moderate constipation, daily use of a unique probiotic for two weeks led to a:4

- 42% decrease in constipation,
- 52% decrease in abdominal pain, and
- 48% decrease in nausea.

In addition to restoring regularity, this probiotic could help prevent some of the long-term health effects associated with constipation.





Impressive Clinical-Trial Results

To validate their hypothesis, scientists set out to test B. lactis HN019 in a clinical trial.4

A total of 88 men and women ages 25 to 65 were divided randomly into three groups:4

- One group took 1.8 billion colony-forming units (CFU) of B. lactis HN019 once daily,
- Another took 17.2 billion CFU of B. lactis HN019 once daily, and
- A third group took a daily placebo.

After 14 days, compared to the placebo group:4

- Those who took the lower B. lactis dose had colonic transit times that were 18.5 hours faster, an improvement of 31%.
- Those who took the higher daily dose of 17.2 billion CFU of B. lactis had colonic transit times that were 28.1 hours faster, an improvement of 57%.

Safe, Quick Constipation Relief

- Scientists have identified a probiotic strain, Bifidobacterium lactis HN019. that relieves ongoing constipation without side effects.
- In a clinical trial, oral use of B. lactis HN019 decreased colonic transit time by up to 57%, eased nausea and abdominal pain, and effectively restored normal bowel regularity in just two weeks.
- In addition to reducing quality of life, **chronic constipation** may be associated with serious health issues, including higher rates of heart disease, cancer, and all-cause mortality.

For Immediate Relief

Life Extension long ago published a solution for **constipation** caused by insufficient **peristalsis**.

The term peristalsis refers to a series of organized muscle contractions that moves food through the digestive tract.¹⁵

Insufficient or **ineffective peristalsis** means there is not enough colon contractile activity, or the activity does not occur in the necessary rhythmic pattern needed to completely evacuate one's bowels.

The encouraging news is that if one **drinks** the proper **nutrient mix** on an **empty** stomach (usually first thing in the morning), a **surge of peristalsis** will occur within an hour that cleans out most or all fecal matter.

The most popular nutritional powders used for this purpose contain vitamin C with magnesium and/or potassium.

It is important to **drink at least one to two** glasses of water (8 to 16 ounces) after taking

these powdered nutrient mixes as they will draw water from surrounding tissues into the colon to facilitate passage of feces.

By increasing the volume of **water** in the intestine in combination with **nutrients** that stimulate peristalsis, stools are softened, intestinal muscle contraction is stimulated, and bowel evacuation is prompted.

The most popular formula our customers use is a low-cost effervescent buffered blend comprised of **vitamin C + magnesium + potassium**.

Those with **chronic kidney disease** should not take the high doses of magnesium and potassium used to stimulate peristalsis. This is especially relevant to those with advanced kidney disease (stages 3 and 4).

Maintaining <u>sufficient</u> intake of **magnesium** and **potassium**, however, is critical for all individuals.

Subjects taking the *higher* dose improved their slower-than-normal colonic transit times into the **normal range**—within just **two weeks**.

Digestive discomfort questionnaires completed by the participants showed that in the *higher-dose* probiotic group the symptom frequency was, on average:⁴

- For constipation, decreased 42%,
- For abdominal pain, decreased 52%, and
- For nausea, decreased 48%.

Taking this probiotic did not result in any adverse effects.⁴

How the Probiotic Works

Studies have provided insight into *how B*. *lactis* **HN019** may be able to reduce colonic transit time.^{4,8,9}

Certain types of bacteria in the digestive tract act on food in the digestive tract, producing **short-chain fatty acids**, which are a source of **energy** for cells lining the colon. These fatty acids are essential to optimal gastrointestinal health.

Preclinical models have demonstrated that short-chain fatty acids interact with a protein within certain cells that exist alongside intestinal **epithelial** (surface) cells. This sets in motion a cascade of events that speeds up **colonic transit times**.^{4,9-11}

A clinical study also revealed that use of *B. lactis* HN019 *increased* levels of two other beneficial bacteria, **bifidobacteria** and **lactobacilli**, and *decreased* levels of harmful **enterobacteria**.¹²

The net result of this gut microbiota rebalancing may be stimulation of **peristalsis** (the muscle contractions that move food through the digestive tract) and a shortening of colonic transit time.⁴

More Effective than Constipation Drug

Scientists at Life Extension compared this probiotic's clinical results to those of a prescription constipation medication called prucalopride.

They reviewed multiple clinical studies of prucalopride.13

Reviewers found that **prucalopride** improved colonic transit times by 20%, 13 far less than the 57% improvement seen in the high-dose B. lactis HN019 trial.4

Prucalopride has been associated with multiple side effects, including headaches, abdominal pain, nausea, and diarrhea,14 compared to no adverse side effects with the probiotic.4

This probiotic strain shows greater effectiveness at relieving and preventing constipation than a major prescription drug. This can help prevent serious, longterm health effects.

Summary

Constipation afflicts about a third of people over 60. Beyond the discomfort and unpleasantness, there is even some thought that constipation may have an association with heart disease, cancer, and other serious health risks.

Clinical research showed that taking the probiotic strain Bifidobacterium lactis HN019 decreased colonic transit time by up to 57%, safely restoring regular bowel movements within two weeks.

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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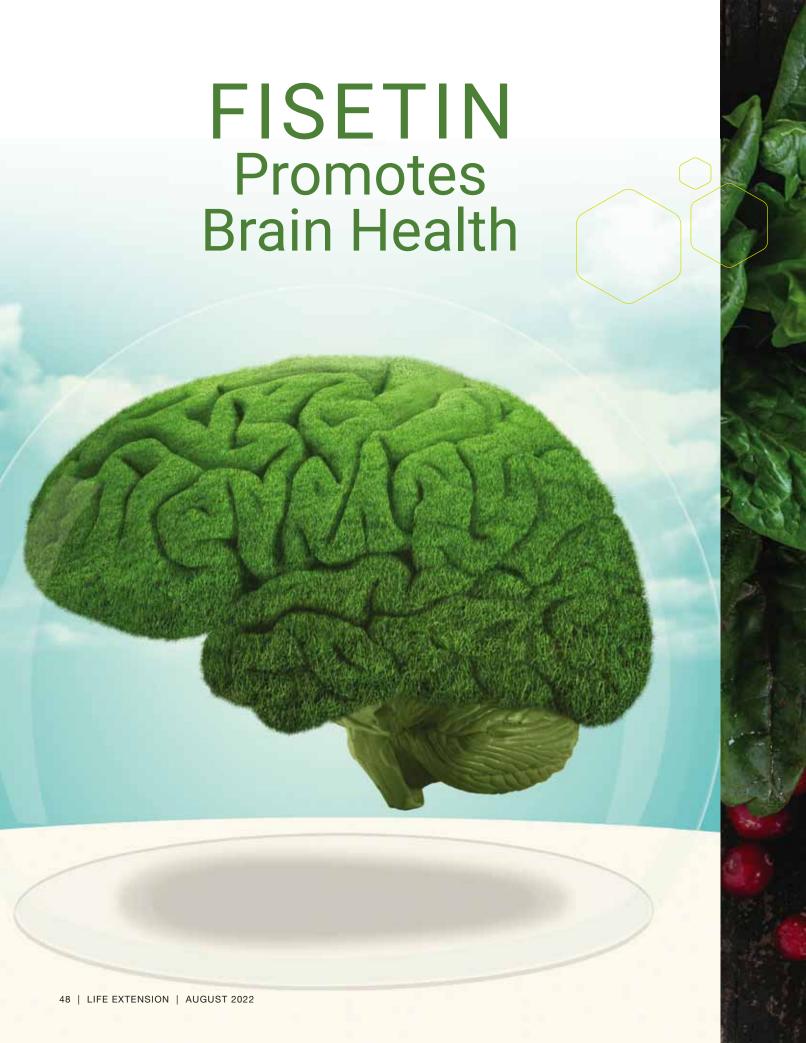
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A Multifunctional Nutrient

The **flavonoids** are a group of **plant compounds** with health-promoting properties.

Fisetin is a flavonoid with dozens of biological mechanisms that support neurological health.2

Fisetin is found in many fruits and vegetables, but only in very small amounts. Most commercially available oral fisetin has poor bioavailability.3

Scientists have overcome this problem by coating the fisetin in a form of fiber found in the fenugreek plant.

This helps protect the fisetin from getting broken down, leading to about 25 times more fisetin being absorbed into the body.4

Promise for Neurological Conditions

In pre-clinical studies, fisetin has shown promise for a range of neurological disorders that include:1,2

- Alzheimer's disease.
- Brain aging and cognitive decline,
- Stroke.
- Neurological complications of diabetes,
- Depression,
- Traumatic brain injury,
- Parkinson's disease,
- Huntington's disease,
- Amyotrophic lateral sclerosis (ALS),
- Neurotoxicity (damage to the nervous system caused by toxins),
- Schizophrenia,
- Vascular dementia, and
- Other forms of neurodegenerative disease.

Additionally, in a clinical trial, fisetin improved response to treatment in ischemic stroke.5

Fighting Against Neurological Disease

A review published in 2021 found that fisetin modulates 37 different biological pathways.2

All these actions could reduce the risk and severity of age-related neurological conditions.

Five ways that fisetin protects the brain and nervous system are:

1. PREVENTING TOXIC BRAIN DEPOSITS

The plaques and tangles deposited in the brain are made of toxic proteins called beta-amyloid and hyperphosphorylated tau. In preclinical models, fisetin prevents their accumulation.6,7

In another study, fisetin increased autophagy, a form of cellular cleaning, which helps clear existing toxic proteins from brain cells.8

2. ANTIOXIDANT ACTIVITY

Oxidative stress is a significant contributor to neurological disorders.

Fisetin is a potent antioxidant that can scavenge harmful compounds before they do damage.1



It has been shown in preclinical models that fisetin increases intracellular levels of glutathione, a freeradical scavenger important for cell survival.9

3. REDUCED CHRONIC INFLAMMATION

Chronic inflammation contributes to most chronic disease in the brain.

In preclinical trials, two forms of fisetin blocked activation of nuclear factor-kappa B (NF-kB), a master regulator of inflammation. 10,111 This action prevents the release of pro-inflammatory compounds.

In a study in a particular strain of mice susceptible to Alzheimer's-type neurological degeneration, feeding them fisetin in early to mid-life prevented learning and memory decline.

Fisetin also limited levels of inflammatory compounds that are elevated in Alzheimer's disease and other neurodegenerative disorders.¹²

4. MEMORY AND LEARNING SUPPORT

A protein called **CREB** helps brain cells adapt to experiences over time, a process known as long-term potentiation.¹³ This is a critical component of memory formation.

Fisetin increases activation of CREB and enhances long-term potentiation in preclinical studies. 13-15

5. BLOCKING GLYCATION DAMAGE

Glycation occurs when glucose attaches to proteins, DNA, and lipids, forming toxic compounds.

Damage caused by glycation has been associated with cognitive dysfunction and neurodegenerative diseases like Alzheimer's disease.16 Glycated betaamyloid is even more damaging to brain function than non-glycated forms.¹⁷

In mice, fisetin was shown to reduce glycation and its damaging effects.18

Summary

The flavonoid **fisetin** acts in multiple ways that may help improve nervous system health and reduce risk for neurological disorders.

Two recent reviews amalgamate mostly preclinical evidence supporting fisetin's abilities.

These include the prevention or management of neurological conditions such as stroke, Alzheimer's, Parkinson's, and cognitive decline. •



A Brain-Protecting Nutrient

- Neurological disorders were identified, as of 2016, as responsible for more disability than any other cause and were the second leading cause of death.
- **Fisetin** is a nutrient found in many fruits and vegetables in small amounts.
- Oral fisetin is rapidly metabolized. Combining it with fiber from fenugreek increases its bioavailability by about 25 times.
- Preclinical research has shown that fisetin may have benefits for neurological conditions including stroke, Alzheimer's disease, Parkinson's disease, brain aging, and cognitive decline. A clinical study has also been completed and others are on the way.

A Potent Senolytic

The discovery of senolytics is one of the most important breakthroughs in the field of aging.

Senolytic compounds can eliminate aged senescent cells from the brain and body. These dysfunctional cells contribute to many conditions of older age, including neurodegenerative disease.19

Removing senescent cells improves tissue function, reduces risk for chronic disease, and extends lifespan in preclinical studies.20-22

In a panel of 10 flavonoids tested in mice genetically predisposed to rapid and premature aging, fisetin was the most potent senolytic.23

Senolytics like fisetin hold promise in the fight against aging and age-related disease.

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

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Increase

AMPK

to Better Manage Body Weight

Most people today consume excess calories.

This results in **mTOR** constantly running at high gear, which is a factor in unwanted **fat storage**.

Studies show that <u>increasing</u> **AMPK** activity turns down excess **mTOR**.¹

Reduce Cell Fat Storage

Scientific studies show that <u>increasing</u> **AMPK** activity can encourage cells to store less fat and burn it as energy.^{2,3}

AMPK Metabolic Activator was formulated based on data showing <u>reduced</u> **belly fat** in response to just one of its ingredients (*Gynostemma pentaphyllum*).³

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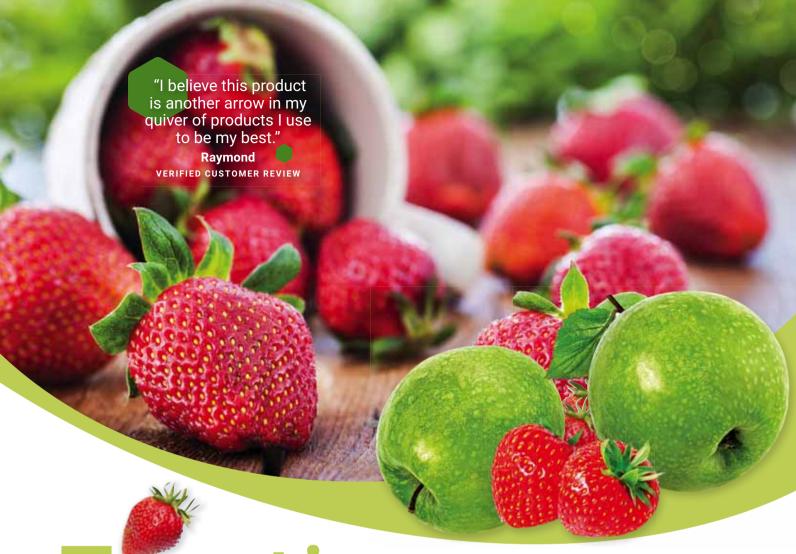
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Caution: High-dose ascorbic acid may cause diarrhea or gastric upset. If this occurs, lower the dose. Those prone to calcium oxalate kidney stones should consult their doctor before using these products.



Fisetin The Longevity Flavonoid

Fisetin, a flavonoid found in strawberries and apples, is currently being studied for its effectiveness as a senolytic in humans.1

In preclinical studies, fisetin:

- Mimics effects of calorie reduction²
- Targets longevity pathways²⁻⁶
- Extends lifespan of mice by about 10%⁷
- Removes senescent cells through senolytic action7
- Suppresses excess mTOR activation⁸

Fisetin is poorly absorbed due to its breakdown in the small intestines.

Bio-Fisetin solves this problem by enclosing fisetin with a compound from the fenugreek herb.

A human trial showed bioavailability of this fisetin compound increased up to 25 times compared to fisetin by itself.9

Just one capsule daily of Bio-Fisetin helps manage senescent cells and may support overall longevity.

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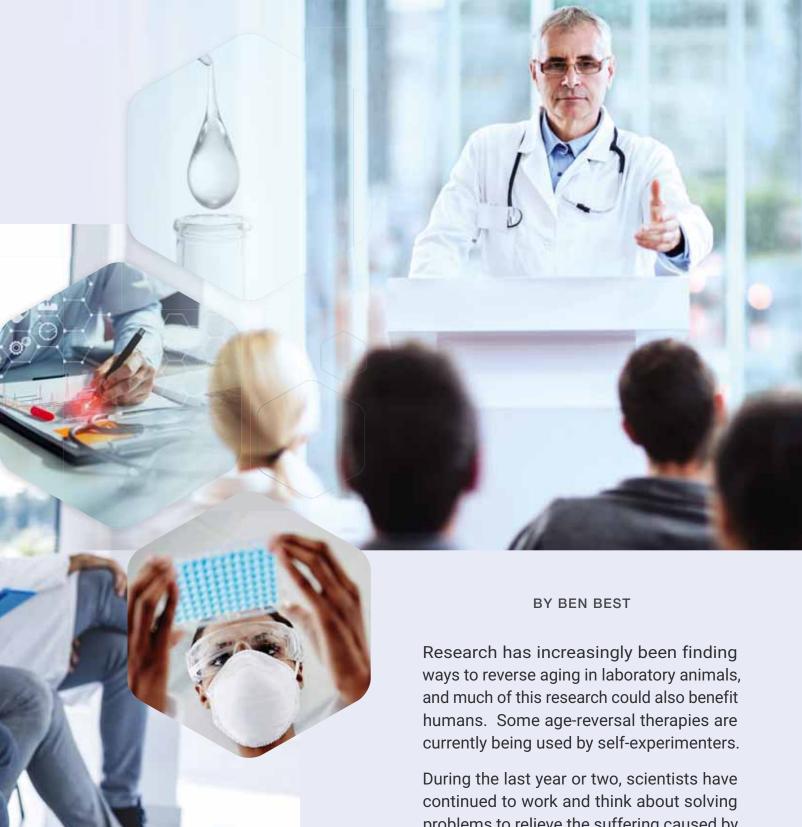




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2021 AGING CONFERENCES





problems to relieve the suffering caused by aging.

This article describes presentations by scientists at 2021 aging conferences which I found most inspiring.

Therapy for Cancer and Senescent Cells

Peter de Keizer, PhD (Associate Professor, UMC Utrecht University, Utrecht, Netherlands) is developing therapeutic interventions against cancer and senescent cells. Senescent cells are cells that cease to perform normal functions, but which produce chemicals (cytokines) that induce chronic inflammation. The increase in senescent cells with age is believed to contribute to most age-related diseases.1 Therapies that eliminate senescent cells have been shown to extend healthy lifespan in mice.^{2,3} Senescent cells can be cleared by immune system cells, but with age, immune cells themselves become senescent, creating a vicious cycle of increasing numbers of senescent cells as age progresses.4

Dr. de Keizer is concerned that there are many different types of senescent cells, and that therapies to eliminate senescent cells only eliminate a few subtypes, but not others.5 In addition, he is concerned that chemotherapy and radiation therapy not only kill cancer cells, but cause many cells to become senescent.6-8 The proinflammatory factors secreted by those senescent cells enhance the ability of surviving cancer cells to spread into other tissues (metastasis).9 Dr de Keizer believes that therapies targeting senescent cells should be used along with therapies against cancer.10

Dr. de Keizer has created a cell-penetrating peptide that potently kills senescent cells. Unlike other senescent cell therapies, his peptide is more selective for senescent cells. 11,12

Dr. de Keizer has co-founded the company Cleara Biotech (www.clearabiotech.com) dedicated to finding even more effective cell-penetrating peptides to kill senescent cells in the context of cancer therapy. Dr. de Keizer has been raising funds from venture capitalists and has hopes to begin human clinical trials in the near future.

SIRT6 Against Cancer and Aging

Vera Gorbunova, PhD (Co-director Rochester Aging Research Center, University of Rochester, Rochester, New York) is an expert on DNA damage, DNA stability, the effects of SIRT6, and the role these factors play in aging and cancer.

Cancer is a disease of altered DNA, which causes cancer cells to cease their normal functions and instead to simply reproduce and spread. Aging too is a disease of altered DNA, causing cells to lose their ability to reproduce and perform their normal functions in organs and tissues.

DNA controls all cell functions, but with age DNA not only becomes less stable,13 but becomes more likely to become damaged.¹⁴ Both the lack of stability¹⁵ and the increasing DNA damage contribute to the chronic inflammation which causes most age-related diseases, including cardiovascular disease, ¹⁶ Alzheimer's Disease,¹⁷ and cancer.¹⁸ DNA damage is a major cause of cellular senescence.1

Humans have seven sirtuin enzymes (SIRT1 to SIRT7), which perform a wide variety of functions important for health and longevity.¹⁹ Dr. Gorbunova is a world expert on SIRT6, which her team has shown maintains DNA stability, 15 and enhances repair of DNA damage.²⁰ Comparing different rodent species of varying lifespans (from 3 to 32 years) her team has shown that the efficiency of the SIRT6 enzyme for DNA repair corresponds to the maximum lifespan of a rodent species.20 Other researchers have shown that SIRT1 has many health benefits and that SIRT6 has been shown to extend lifespan of mice.21

For many years Life Extension® funded Dr. Gorbunova's SIRT6 research. Noting that resveratrol and similar chemicals enhance the enzyme activity of SIRT1, Dr. Gorbunova sought chemicals which could enhance SIRT6 activity.²² Many SIRT6 activators have been found,23 including the Asian seaweed polysaccharide fucoidan.





The Role of Macrophages in Aging

Eric Verdin, MD (CEO, Buck Institute for Research on Aging, Novato, California) has been investigating the role of macrophages in aging and aging-related

Macrophages are a type of white blood cell of the immune system that engulfs and digests microbes, cellular debris, and other contaminants. The word macrophage was derived from Greek (macro=big, phage=eater) by the scientist who discovered these cells. Macrophages play an important role in the progression of inflammation-related diseases, including atherosclerosis, diabetes, and cancer.²⁴ Macrophages can be either inflammatory (M1 macrophages, which produce inflammatory responses) or anti-inflammatory (M2 macrophages, which are attracted to sites of inflammation and produce anti-inflammatory responses).25 With aging, macrophages become increasingly of the M1 type, contributing to chronic inflammation.²⁶

Aging is also characterized by an increasing number of senescent cells.27 Macrophages are attracted to senescent cells to engulf and remove them.28 But collections of macrophages and senescent cells become increasingly pro-inflammatory, causing nearby cells to become senescent, possibly also including the macrophages themselves.²⁸ The pro-inflammatory macrophages increasingly express the CD38 enzyme which degrades the important enzymatic co-factor NAD+.29

The loss of NAD+ with age results in reduced function of sirtuins and other enzymes required for DNA repair, and reduced capacity of the immune system to function optimally.30

Dr. Verdin presented evidence for the hypothesis that senescent macrophages are a substantial source of NAD+ loss and of senescent cells in aging.31,32

Immune System Rejuvenation

Gregory Fahy, PhD (Chief Scientific Officer and Co-founder, Intervene Immune, Torrance, California) described the TRIIM (Thymus Regeneration, Immunorestoring, and Insulin Mitigation) clinical trial that he designed and led, as well as the follow-up extended (TRIIM-X) trial which he is leading, and which is still seeking participants. TRIIM was a pilot study of nine male adults, but TRIIM-X has been seeking 85 adults of diverse gender and ethnicity for a study to be completed in November 2022.

The TRIIM trial was designed to regenerate the thymus gland using growth hormone, while attempting to prevent the growth hormone from causing diabetes. Metformin and dehydroepiandrosterone (DHEA) were the anti-diabetic agents.33

Dr. Fahy had been inspired to attempt the TRIIM trial by his knowledge that growth hormone has been shown to reconstitute the thymus in rats,34 dogs,35 and HIV-infected patients.36

With age, immune system function declines, leaving elderly persons vulnerable to infection. Persons 65 years or older account for 90% of influenza-related deaths in the United States.37 Immune system cells produced by the thymus gland (T-cells) decline in function with age,38 due to the fact that the thymus gland shrinks and becomes clogged with fat early in life.39 Fresh ("naive") T-cells decline by about two orders of magnitude (hundreds of times) between the ages of 20 and 70.40

Many animal experiments have shown that reduced growth hormone early in life leads to healthy longevity.⁴¹ By contrast, the TRIIM trial showed that growth hormone given late in life reversed the biological age of nine men an average of 1.5 years,33 as indicated by DNA methylation "clocks". 42-45 Not only was T-cell function improved, but there were fewer immune cells expressing the NAD+ depleting enzyme CD38. NAD+ decline with age underlies many age-related diseases, such that restoration of NAD+ could be rejuvenating.46

Neural Networks to Predict Biological Age

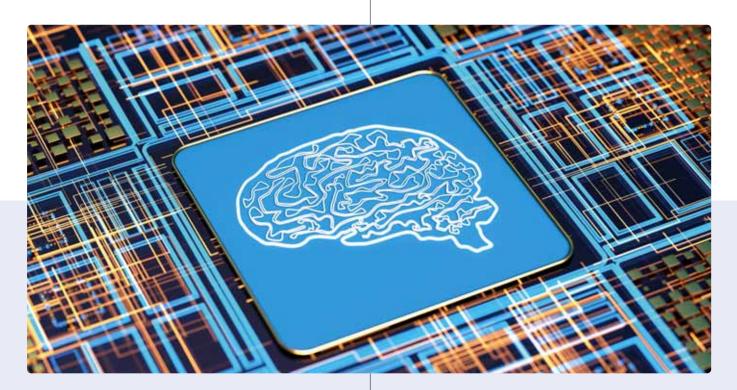
Alex Zhavoronkov, PhD (CEO, Insilico Medicine, Hong Kong Science Park, Hong Kong) is a scientist and entrepreneur who is employing the deep neural network form of artificial intelligence to understand human aging. Neural networks are computer programs that

simulate the way neurons interact to cause learning in the human brain. A neural network is deep when it contains many layers of simulated neurons. Unlike conventional computer programs which are given explicit instructions by a computer programmer, neural networks are simply "trained" by data to produce a result without the programmer knowing the details by which the simulated neurons produced that result.

Dr. Zhavoronkov has been inspired in his neural network efforts by the use of DNA methylation clocks to predict human age.44 Dr. Zhavoronkov trained deep neural networks on 60,000 human blood samples and thereby identified five blood components that could be used to predict human age.47 Predicting biological age can be valuable in human longevity studies because it allows a way of measuring whether a relatively shortterm therapeutic intervention successfully slows or reverses aging, rather than having to wait decades for a human lifespan to play out.

Using deep neural networks to analyze the blood samples of people of different nationalities, Dr. Zhavoronkov's team determined that South Koreans tend to have a predicted age younger than their chronological age, whereas for Eastern Europeans the predicted age is older.48

Among his efforts as an entrepreneur, in addition to founding Insilico Medicine (insilico.com, which searches for longevity drugs using artificial intelligence),



Dr. Zhavoronkov has founded Deep Longevity (deeplongevity.com, which specializes in aging clocks), and Longevity Medicine (longevity-medicine.org, which offers a Longevity Medicine Course (longevity.degree) for medical professionals.

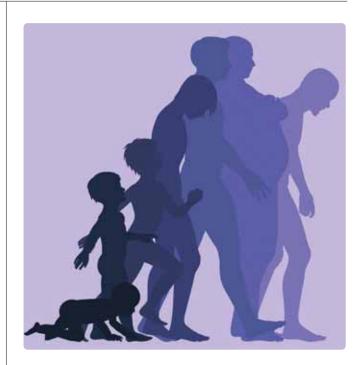
Administering \$350 million for Rejuvenation Research

Patrick Burgermeister (Partner, Kizoo Technology Ventures, Germany) spoke about the role of Kizoo Technology Ventures (www.kizoo.com) in providing seed funding, mentoring, and follow-on financing for biotechnology companies focused on rejuvenation.

Kizoo is part of the Forever Healthy Foundation (www.forever-healthy.org) founded by internet entrepreneur Michael Greve. Greve has also funded the annual **Undoing Aging** conferences in Berlin, Germany (collaborating with the rejuvenation-focused SENS Foundation). In May 2021 Greve committed an additional 300 million EUROs (about \$350 million) for support of rejuvenation biotechnology startups. Greve believes that abolishing cancer, heart attacks, and dementia is just a matter of time, money, and research.

Mr. Burgermeister said that Kizoo funding contrasts with that of typical investors looking for low-risk companies that will soon be profitable. He said that Kizoo prefers companies in the earliest stages of development (sometimes not even yet founded). Kizoo prefers to lead the way, in contrast to "lightweight investors," who look for companies which already have funding from many other investors.

Kizoo prefers companies in which the founders have a strong financial interest. Some of the companies that Kizoo has funded include Underdog Pharmaceutical (clearing non-degradable cholesterol from arteries), Revel Pharmaceutical (therapeutic enzymes to degrade age-associated molecular damage), Cellvie (mitochondrial augmentation and replacement), and



Turn Biotechnologies (epigenetically reprogramming cells to be younger), among other companies. Mr. Burgermeister serves as a director for some of the Kizoo-funded companies.

Concluding Remarks

It is encouraging to see not only the development of therapies to slow aging, but to potentially reverse aging. The advent of this potential has gratifyingly brought increasing amounts of money into aging research, along with entrepreneurial activity associated with therapies. •

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



DEKEIZER



FAHY



BURGERMEISTER



GORBUNOVA



VERDIN

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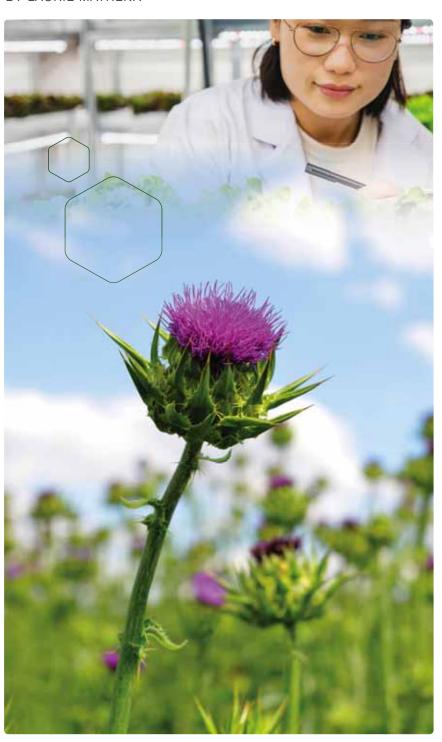
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What Is Milk Thistle?

BY LAURIE MATHENA



Deaths from liver cirrhosis jumped 65% in the U.S. between 1999 and 2016.3,4

Key factors contributing to liver disease are obesity, diet, and alcohol consumption.^{1,2}

Milk thistle extract has demonstrated protective effects against an array of liver disorders.5-8

Nonalcoholic **Fatty Liver Disease**

In the past, viral hepatitis and excess alcohol ingestion were considered the greatest threats to the liver.2

Today, the surge in diabetes and obesity has resulted in an epidemic of nonalcoholic fatty liver disease (NAFLD),9 which is characterized by the accumulation of fatty compounds in the liver in the absence of chronic alcohol use.

It can progress over time to cause liver *fibrosis* (tissue scarring) that can lead to liver cancer, cirrhosis, and/or liver failure. 10-13

Extracts of the herb milk thistle, containing the compound silvma**rin**, have long been used to protect liver function in patients with liver disease.

Several clinical trials found that milk thistle, alone or in combination with vitamin E, and phosphatidylcholine reduces liver fat, fibrosis, and enzyme levels in patients with NAFLD.14-17

A meta-analysis of eight randomized, controlled trials found that milk thistle may help improve NAFLD.¹⁷

Milk thistle has been shown to improve blood markers of liver damage and may also help reduce fasting glucose and LDL cholesterol elevations that often accompany NAFLD.17,18

Measuring liver enzyme blood levels is an important tool for identifying liver injury and helping track response to treatment.

Liver function tests are included in many Life Extension® blood panels.

Liver Toxins

The liver is susceptible to toxins known as **hepatotoxins**, which include the commonly used pain medication acetaminophen, alcohol, and others.

In rats, milk thistle extract helped prevent liver damage when given before or during the exposure to a liver toxin.19

In one study, a group of patients with alcohol-induced liver disease who were treated with milk thistle extract showed improvements in liver enzymes and liver pathology.8

Cirrhosis

Cirrhosis is the end stage of chronic liver injury in both alcoholic and nonalcoholic liver disease. It is generally considered irreversible, but scientists have observed promising improvements with milk thistle extract in clinical trials.

In one study, giving 420 mg of milk thistle extract to patients with cirrhosis was associated with an overall higher four-year survival rate.5

For optimal absorption, a standardized extract of milk thistle is used by readers of this publication that has been made more bioavailable via a phospholipid delivery system.20 •

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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Probiotics Target Liver Health

One in four adults in the U.S. has nonalcoholic fatty liver disease (NAFLD).21-24

Despite over four decades of research, there are still no medications approved by the U.S. Food and Drug Administration (FDA) to treat fatty liver disease.

However, research has found a link between the gut microbiota and liver health. This led researchers to create a blend of probiotics and a prebiotic to target liver health.

In two clinical trials of people with NAFLD, a carefully designed blend of seven probiotics and a prebiotic decreased a marker of liver damage and reduced levels of fibrosis (scarring) from moderate or almost severe to normal. 25,26

These findings suggest that the **probiotic-prebiotic** blend stopped the progression of liver disease and reversed liver damage that was already present.

Other Nutrients That Promote Liver Health

Some nutrients have also shown promise as a way to help control liver disease.

In human trials, vitamin E tocotrienols improved markers of liver health seen on an ultrasound, while reducing liver enzymes, C-reactive protein, and signs of oxidative stress.27-29

Phosphatidylcholine is an essential phospholipid which is a vital part of cellular membranes. Phospholipids have been used safely for years to protect liver function in patients with various liver diseases.30 In a number of human trials, phosphatidylcholine intake alone or with other nutrients improved NAFLD, reducing liver enzyme levels and improving ultrasound findings. 30-32 A more bioavailable form of phosphatidylcholine known as polyenylphosphatidylcholine or PCC is the preferred choice for liver support as it specifically targets hepatocytes.

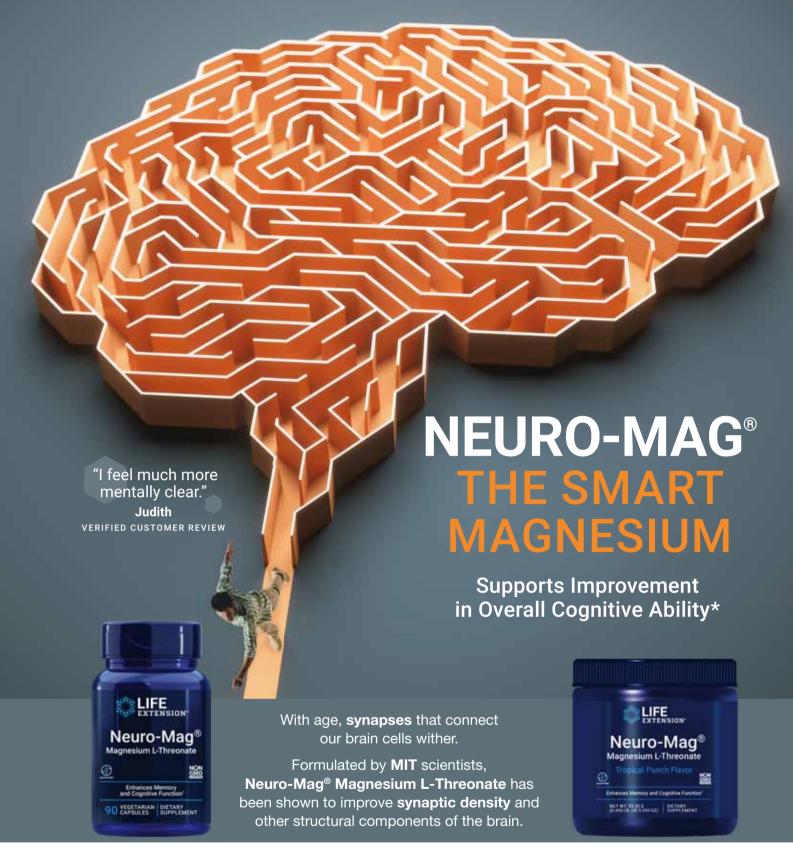
N-acetyl-L-cysteine (NAC), a versatile sulfur-rich compound prevents liver damage following acetaminophen poisoning.33 NAC rapidly restores depleted glutathione levels, sparing liver cells from the effects of oxidative damage.34-36

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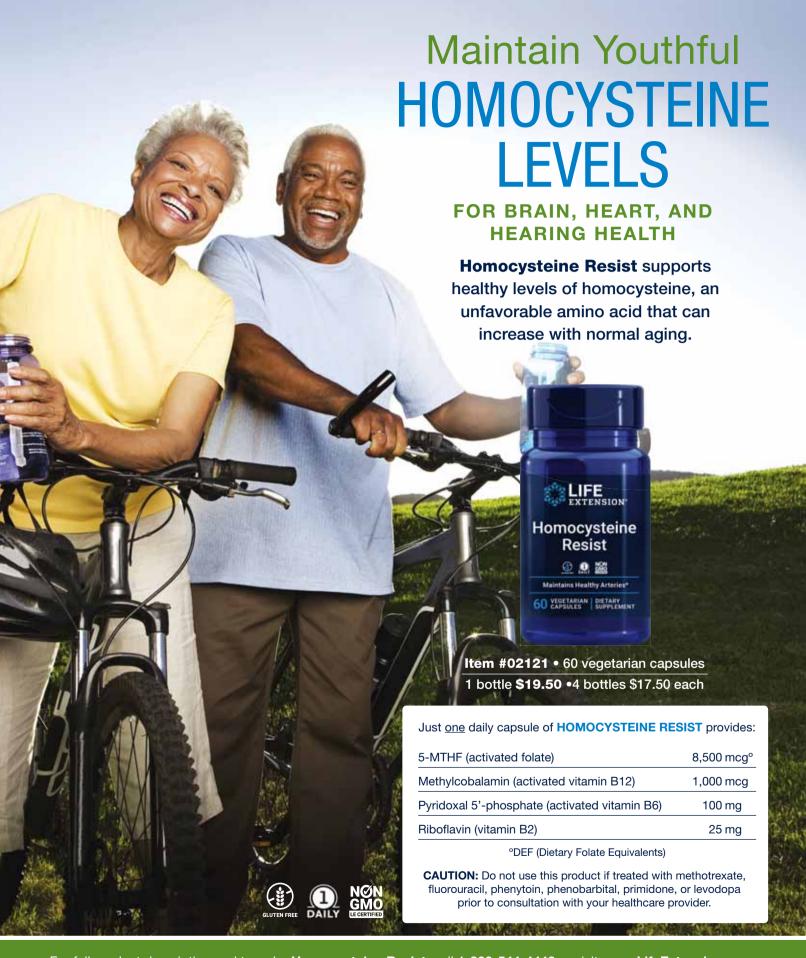
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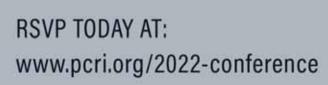
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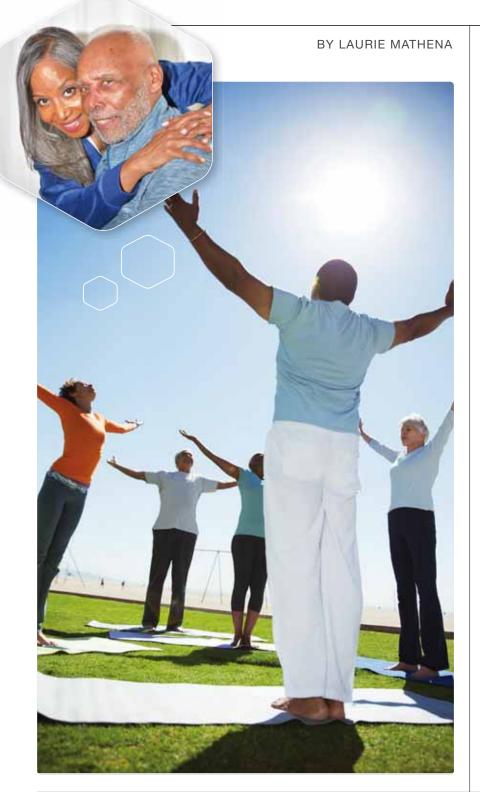




[KEYNOTE TOPICS]

- All Prostate Cancer Treatments
- Newly Diagnosed
- Diet & Exercise
- · Sexual Dysfunction
- Active Surveillance
- Treatment Side Effects
- Prostate Imaging
- Benign Prostate Hyperplasia (BPH)
- Advanced Treatment Strategies

Robin Downes: "Yoga Saved My Father's Life"



Robin Downes is an Emmy awardwinning journalist who worked on high-profile television shows such as the ABC News show 2020 and The Cosby Show, and as the production coordinator for Spike Lee films.

But after experiencing a personal tragedy, Downes left behind her prestigious career in the entertainment industry to pursue something she believed had much more meaning: yoga.

Over the next 25 years, Downes sold millions of yoga instructional videos and became the "yoga instructor to the stars," teaching private yoga sessions to celebrities and sports stars like Brandy, Vanessa Williams, and NBA star John Sally.

Now, her most important client yet is someone she affectionately calls "Sergeant Major"—her father.

How Yoga "Resurrected" **Her Father**

Downes admittedly had a fairytale career in the entertainment industry. So, at age 41, when she was faced with marriage problems and her mother's breast cancer diagnosis, she needed tools to help her cope with stress.

That's when she turned to yoga.

"In 1995, I walked into my first yoga class, and I knew that this was what I needed," said Downes. "When stress and reality hit, it was yoga that saved me."

Now, at age 61, she is using those same tools to help save her elderly father.

Downes' father was malnourished, depressed, and drinking too much—plus dealing with health problems like congestive heart failure, high blood pressure, and dementia.

He needed an intervention.

"I was able to draw on the wonderful experience I had gathered from treating my VIP yoga clients," said Downes. "I thought maybe I could use yoga to help him out in his final days of life, but that was four years ago. Instead, yoga was able to resurrect my dad."

Downes started with simple, seated poses designed to help her father breathe better.

"As we get older, we start to walk hunched over. My father had begun to walk with a cane. All of this diminishes your capacity to breathe," said Downes.

It also impacts balance and flexibility.

To help combat this, Downes likes to do a pose called Tadasana Mountain Pose with her father. It encourages participants to ground their feet, lift their chest, push their shoulders back, and sit tall—and then take deep, cleansing breaths.

Poses like these—all of which can be done seated in a chair—can help older individuals gain flexibility, strength, and balance, while also providing what Downes calls a "sense of ease."

Studies have shown the beneficial impact this can have on dementia patients.

In one study, regular yoga sessions were found to improve respiratory function, improve balance control in Alzheimer's patients, and help calm agitated patients.

And in an exciting study of people with mild cognitive impairment, 12 weeks of yoga led to short- and long-term improvements in executive functioning and produced beneficial effects on depressed mood and resilience.

Yoga can also reduce stress for both the patient and the caregiver, which is why Downes participates in a program called Caregivers Embracing Elder Care.

Through this program, Downes guides caregivers through meditation and yoga moves, and a geriatric care manager provides practical tips for caregivers of elderly patients.

"Yoga provides a greater sense of ease, greater mobility, and ultimately greater longevity," said Downes.

Yoga Flava®

Downes knew her father might not be receptive to traditional yoga. Fortunately, her yoga style is anything but traditional.

Her yoga company, called Yoga Flava®, is unique because it combines ancient yoga practice with urban culture and modern music.

"It uses ancient moves to contemporary grooves," said Downes.

In fact, when Downes pioneered Yoga Flava® in 1995, she produced an instructional video and became the first African American female to have an internationally distributed yoga video.

Working with her father showed her that this unique take on yoga transcended generations as well as cultures. "Yoga Flava® has achieved such remarkable success because it uses music that's familiar," said Downes. "For me, that means Erykah Badu or Mary J. Blige. For my father, that means doing yoga to salsa music or Nat King Cole."

Choosing Your Best Life

Downes knows that in order to care for her father, she needs to care for herself as well. That's why, in addition to her yoga practice, she has started boxing.

"It's another way of relieving stress, and it's a great cross-training program for me at this point," said Downes. "I know I need to be doing everything I can to help with caregiving. In addition to managing my stress level, I need to maintain my strength."

She also eats lots of fresh fruit and vegetables and takes numerous supplements, including vitamins A, C, and D, biotin, iodine, zinc, selenium, collagen, turmeric, and fish oil.

She believes it's all part of a holistic lifestyle that contributes to longevity, both for her and for her father.

"Every day we can pick and choose how we want to live our best life," said Downes. "For me, it's about perspective. I get to enjoy my father, and I get to create a lifestyle that can bring both of us joy."

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02270 DNA Protection Formula	02357 Life Extension Mix™ Tablets with Extra Niacin				
02431 Essential Youth - L-Ergothioneine	02365 Life Extension Mix™ Tablets without Copper				
02119 GEROPROTECT® Ageless Cell™	02292 Once-Daily Health Booster • 30 softgels				
02415 GEROPROTECT® Languity A L™	02291 Once-Daily Health Booster • 60 softgels				
02133 GEROPROTECT® Longevity A.l.™ 02401 GEROPROTECT® Stem Cell	02313 One-Per-Day Tablets 02428 Plant-Based Multivitamin				
02211 Grapeseed Extract	02317 Two-Per-Day Capsules • 60 capsules				
00954 Mega Green Tea Extract (decaffeinated)	02314 Two-Per-Day Capsules • 00 Capsules				
00953 Mega Green Tea Extract (lightly caffeinated)	02316 Two-Per-Day Capsules • 60 tablets				
01513 Optimized Fucoidan with Maritech® 926	02315 Two-Per-Day Tablets • 120 tablets				
02230 Optimized Resveratrol	NERVE & COMFORT SUPPORT				
01637 Pycnogenol® French Maritime Pine Bark Extract					
02210 Resveratrol	02202 ComfortMAX™				
00070 RNA (Ribonucleic Acid)	02303 Discomfort Relief				
02301 Senolytic Activator®	PERSONAL CARE				
01208 Super R-Lipoic Acid	01006 Biosil™•5 mg, 30 veg capsules				
01919 X-R Shield	01007 Biosil™•1 fl oz				
MEN'S HEALTH	00321 Dr. Proctor's Advanced Hair Formula				
02209 Male Vascular Sexual Support	00320 Dr. Proctor's Shampoo				
00455 Mega Lycopene Extract	02322 Hair, Skin & Nails Collagen Plus Formula				
02306 Men's Bladder Control	01278 Life Extension Toothpaste				
01789 PalmettoGuard® Saw Palmetto and Beta-Sitosterol	00408 Venotone				
01790 PalmettoGuard® Saw Palmetto/Nettle Root Formula	00409 Xyliwhite Mouthwash 02304 Youthful Collagen				
and Beta-Sitosterol	02252 Youthful Legs				
01837 Pomi-T®	PET CARE				
01373 Prelox® Enhanced Sex for Men					
01940 Super MiraForte with Standardized Lignans 02500 Testosterone Elite	01932 Cat Mix				
02300 lestosterone Line	01931 Dog Mix				
01909 Triple Strength ProstaPollen™					
01909 Triple Strength ProstaPollen™ 02029 Ultra Prostate Formula	PROBIOTICS				
02029 Ultra Prostate Formula	PROBIOTICS 01622 Bifido GI Balance				
02029 Ultra Prostate Formula MINERALS	PROBIOTICS 01622 Bifido GI Balance 01825 FLORASSIST® Balance				
02029 Ultra Prostate Formula MINERALS 01661 Boron	PROBIOTICS 01622 Bifido GI Balance 01825 FLORASSIST® Balance 02421 FLORASSIST® Daily Bowel Regularity				
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02029 Ultra Prostate Formula MINERALS 01661 Boron 02107 Extend-Release Magnesium 01677 Iron Protein Plus 02403 Lithium	PROBIOTICS 01622 Bifido GI Balance 01825 FLORASSIST® Balance 02421 FLORASSIST® Daily Bowel Regularity 02125 FLORASSIST® GI with Phage Technology 01821 FLORASSIST® Heart Health 02250 FLORASSIST® Mood Improve				
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80179	Brightening Peptide Serum	01786	Melatonin 6 Hour Timed Release 3 mg, 60 veg tablets	
80176	Collagen Boosting Peptide Cream	01721	Optimized Tryptophan Plus	
80156	Collagen Boosting Peptide Serum	01444		
02408	Collagen Peptides for Skin & Joints	01445	-	
80180	CoQ10 and Stem Cell Rejuvenation Cream	02502	Rest & Renew	
80169	Cucumber Hydra Peptide Eye Cream	VITAMINS		
02423	Daily Skin Defense	01533	Ascorbyl Palmitate	
80141	DNA Support Cream	00920	Benfotiamine with Thiamine	
80163	Eye Lift Cream	00664		
80123	Face Rejuvenating Anti-Oxidant Cream	01945		
80109	Hyaluronic Facial Moisturizer	00102	Biotin	
80110	Hyaluronic Oil-Free Facial Moisturizer	00084	Buffered Vitamin C Powder	
80138	Hydrating Anti-Oxidant Facial Mist	02229	Fast-C® and Bio-Quercetin Phytosome	
00661 55495	Hydroderm	02075	Gamma E Mixed Tocopherol Enhanced with	
80103	Instensive Moisturizing Cream Lifting & Tightening Complex		Sesame Lignans	
80168	Melatonin Advanced Peptide Cream	02070		
80114	Mild Facial Cleanser	01913	5 / 1	
80172	Multi Stem Cell Hydration Cream	01674	•	
80159	Multi Stem Cell Skin Tightening Complex	02244	1, , , , , , , , , , , , , , , , , , ,	
80122	Neck Rejuvenating Anti-Oxidant Cream	02232	1 , , , , ,	
80174	Purifying Facial Mask	01936		
80150	Renewing Eye Cream	00065		
80142	Resveratrol Anti-Oxidant Serum	00373	No Flush Niacin	
	Shade Factor™	01939		
	Skin Care Collection Anti-Aging Serum		Pyridoxal 5'-Phosphate Caps	
02130	Skin Care Collection Day Cream	01400	· ·	
02131	Skin Care Collection Night Cream	02334	Super K Super Vitamin E	
80166	Skin Firming Complex		Vegan Vitamin D3	
02096	Skin Restoring Ceramides	02422	-	
80130	Skin Stem Cell Serum		Vitamin B6	
80164	Skin Tone Equalizer	00361		
80143	Stem Cell Cream with Alpine Rose		Vitamin B12 Methylcobalamin • 1 mg, 60 veg lozenges	
80148	Tightening & Firming Neck Cream	01537		
80161	Triple-Action Vitamin C Cream		Vitamin C and Bio-Quercetin Phytosome • 60 veg tablets	
80162	Ultimate MicroDermabrasion		Vitamin C and Bio-Quercetin Phytosome • 250 veg tablets	
80173	Ultimate Peptide Serum		Vitamin D3 • 25 mcg (1000 IU), 90 softgels	
80178	Ultimate Telomere Cream	01751		
80160	Ultra Eyelash Booster	01713	Vitamin D3 • 125 mcg (5000 IU), 60 softgels	
80101	Ultra Wrinkle Relaxer	01718	Vitamin D3 • 175 mcg (7000 IU), 60 softgels	
80113 80104	Under Eye Refining Serum	01758	Vitamin D3 with Sea-lodine™	
80104	Under Eye Rescue Cream Vitamin C Lip Rejuvenator	02040	Vitamins D and K with Sea-lodine™	
80129	Vitamin C Serum	WEIG	GHT MANAGEMENT & BODY COMPOSITION	
80136	Vitamin D Lotion	02479	7-Keto® DHEA Metabolite • 100 mg, 60 veg capsules	
	Vitamin K Cream	01807	· · · · · · · · · · · · · · · · · · ·	
SLEE	11 11 11 11 11 11 11 11 11 11 11 11 11	02207	· · · · · · · · · · · · · · · · · · ·	
		02504	Body Trim and Appetite Control	
01512	Bioactive Milk Peptides	02478	DHEA Complete	
02300	Circadian Sleep	01738		
01551	Enhanced Sleep with Melatonin	01908	Mediterranean Trim with Sinetrol™ -XPur	
01511	Enhanced Sleep without Melatonin	01432	Optimized Saffron	
02234	Fast-Acting Liquid Melatonin	00818	Super CLA Blend with Sesame Lignans	
01669 02308	Glycine Herbal Sleep PM	02509	Waistline Control™	
02306	L-Tryptophan	WON	MEN'S HEALTH	
01722	Melatonin • 300 mcg, 100 veg capsules	01942	Breast Health Formula	
01083	Melatonin • 500 mcg, 200 veg capsules	01626		
00329	Melatonin • 1 mg, 60 capsules	01894		
02503	Melatonin • 3 mg, 60 gummies	01054	-	
00330	Melatonin • 3 mg, 60 veg capsules	02204	Menopause 731™	
00331	Melatonin • 10 mg, 60 veg capsules	02319		
00331	Melatonin • 3 mg, 60 veg lozenges	01441	Progesta-Care®	
02201	Melatonin IR/XR	01649	9	
01787	Melatonin 6 Hour Timed Release	02507		
	300 mcg, 100 veg tablets		•	
01788	Melatonin 6 Hour Timed Release			

750 mcg, 60 veg tablets



Selenium promotes the body's production of **glutathione**, a potent cellular antioxidant. It also encourages healthy cell division, thyroid health, and immune function.

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Super Selenium Complex, call 1-800-544-4440

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OR

2. Bone Restore with Vitamin K2, same formula as Bone Restore plus 200 mcg of vitamin K2.



 Bone Restore Elite, same formula as Bone Restore plus
 45,000 mcg of vitamin K2.





Bone Restore
Item #01726 • 120 capsules

Bone

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Superior Calcium Formula

120 CAPSULES SUPPLEMENT

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Bone Restore with Vitamin K2*
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1 bottle **\$18** • 4 bottles \$16.50 each

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1 bottle \$34.50 • 4 bottles \$31.50 each

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* CAUTION: If you are taking a vitamin K antagonist (e.g. warfarin), consult your healthcare practitioner before taking this product.

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IN THIS EDITION OF LIFE EXTENSION MAGAZINE®





7 SILENT DISORDER GOES EPIDEMIC

Nonalcoholic **fatty liver disease** (NAFLD) afflicts up to **75%** of **overweight** adults and **type II diabetics**. Learn how to control or reverse this life-threatening disorder.

20 CALMING EFFECTS OF MAGNESIUM

Low magnesium levels can worsen chronic stress. A new magnesium enables a calming effect.





28 CANCER-FIGHTING PATHWAYS

Compounds found in **cruciferous vegetables** target **six pathways** to reduce **cancer** risk.



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