

The Science of a Healthier Life®

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August 2022

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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**.⁷ 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 <u>less</u> 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 <u>not</u> 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 urea
- 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.



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®



References

- Available at: https://www.ajmc.com/view/nafld-is-a-growing-driver-ofchronic-liver-disease-and-related-deaths. Accessed May 13, 2022.
- Kim H, Lee DS, An TH, et al. Metabolic Spectrum of Liver Failure in Type 2 Diabetes and Obesity: From NAFLD to NASH to HCC. Int J Mol Sci. 2021 Apr 26;22(9):4495.
- Kitade H, Chen G, Ni Y, et al. Nonalcoholic Fatty Liver Disease and Insulin Resistance: New Insights and Potential New Treatments. *Nutrients.* 2017 Apr 14;9(4).
- Available at: https://www.cdc.gov/mmwr/preview/mmwrhtml/ su6001a15.htm. Accessed May 9, 2022.
- Available at: https://www.cdc.gov/nchs/products/databriefs/db360. htm. Accessed May 9, 2022.
- Available at: https://www.niddk.nih.gov/health-information/liverdisease/nafld-nash/definition-facts. Accessed May 11, 2022.
- Available at: https://wwa.mayoclinic.org/diseases-conditions/ nonalcoholic-fatty-liver-disease/diagnosis-treatment/drc-20354573. Accessed May 10, 2022.
- Available at: https://www.uptodate.com/contents/cirrhosis-beyondthe-basics. Accessed May 10, 2022.
- Available at: https://www.uptodate.com/contents/pathogenesis-ofnonalcoholic-fatty-liver-disease. Accessed May 10, 2021.
- Available at: https://www.webmd.com/digestive-disorders/reversenonalcoholic-fatty-liver-disease. Accessed May 10, 2022.
- Available at: https://www.hopkinsmedicine.org/health/conditionsand-diseases/liver-anatomy-and-functions. Accessed May 10, 2022.
- Available at: https://www.mayoclinic.org/diseases-conditions/liverproblems/symptoms-causes/syc-20374502. Accessed May 10, 2022.
- Piscaglia F, Svegliati-Baroni G, Barchetti A, et al. Clinical patterns of hepatocellular carcinoma in nonalcoholic fatty liver disease: A multicenter prospective study. *Hepatology*. 2016 Mar;63(3):827-38.
- Available at: https://www.cancer.org/cancer/liver-cancer/detectiondiagnosis-staging/survival-rates.html. Accessed May 10, 2022.
- 15. Kanwal F, Kramer JR, Mapakshi S, et al. Risk of Hepatocellular Cancer in Patients With Non-Alcoholic Fatty Liver Disease. *Gastroenterology.* 2018 Dec;155(6):1828-37 e2.
- Oh RC, Hustead TR, Ali SM, et al. Mildly Elevated Liver Transaminase Levels: Causes and Evaluation. *Am Fam Physician*. 2017 Dec 1;96(11):709-15.
- 17. Available at: https://www.hopkinsmedicine.org/health/conditionsand-diseases/common-characteristics-of-liver-disease. Accessed May 10, 2022.



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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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A Brain-Specific MAGNESIUM Relieves Stress

magnesium

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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** <u>deficient</u>.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 <u>low</u> 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 <u>form</u> 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 <u>four</u> other common forms of magnesium: magnesium sulfate, oxide, citrate, and malate.⁷



WHAT YOU NEED TO KNOW

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.

References

- Available at: https://www.apa.org/news/press/releases/stress/2022/ march-2022-survival-mode. Accessed April 26, 2022.
- 2. Available at: https://www.apa.org/topics/stress/body. Accessed April 29, 2022.
- Available at: https://www.webmd.com/balance/stress-management/ stress-symptoms-effects_of-stress-on-the-body. Accessed April 29, 2022.
- Pickering G, Mazur A, Trousselard M, et al. Magnesium Status and Stress: The Vicious Circle Concept Revisited. *Nutrients.* 2020 Nov 28;12(12).
- Noah L, Pickering G, Mazur A, et al. Impact of magnesium supplementation, in combination with vitamin B6, on stress and magnesium status: secondary data from a randomized controlled trial. *Magnes Res.* 2020 Aug 1;33(3):45-57.
- Bagatela BS, Lopes IP, do Amaral Pupo IC, et al. Biochemical and hematological study of the supplementation with ATAMg, a new source of N-acetyl-based organic magnesium, on bioavailability. *Journal of Society for development in new net environment in B&H* (HealthMED). 2018;12(4):187-9.
- Uysal N, Kizildag S, Yuce Z, et al. Timeline (Bioavailability) of Magnesium Compounds in Hours: Which Magnesium Compound Works Best? *Biol Trace Elem Res.* 2019 Jan;187(1):128-36.
- Rousseau C. Efficiency of a magnesium salt (ATA MG® 770mg) on premenopausal syndrome: a clinical study report. Synapharm (Company supplied data). 2021.
- Galland L. Magnesium, stress and neuropsychiatric disorders. Magnes Trace Elem. 1991;10(2-4):287-301.
- Seelig MS. Consequences of magnesium deficiency on the enhancement of stress reactions; preventive and therapeutic implications (a review). J Am Coll Nutr. 1994 Oct;13(5):429-46.
- Cuciureanu MD, Vink R. Magnesium and stress. In: Vink R, Nechifor M, editors. *Magnesium in the Central Nervous System*. Adelaide (AU)2011.
- 12. Murck H. Magnesium and affective disorders. *Nutr Neurosci.* 2002 Dec;5(6):375-89.
- Ismail AAA, Ismail Y, Ismail AA. Chronic magnesium deficiency and human disease; time for reappraisal? *QJM*. 2018 Nov 1;111(11):759-63.
- Qu X, Jin F, Hao Y, et al. Magnesium and the risk of cardiovascular events: a meta-analysis of prospective cohort studies. *PLoS One.* 2013;8(3):e57720.

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References

1. *J Ethnopharmacol*. 2020 Oct 5;260:112996.

Nutrients. 2020 Jul 23;12(8).
Nutrients. 2018 Aug 7;10(8).

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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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References: 1. *Nutrients.* 2020 Nov 28;12(12). 2. Synapharm - Company supplied data. 2021. ATA Mg® is a registered trademark of AIDP, Inc.



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Cancer-Fighting Properties of CRUCIFEROUS VEGETABLES

BY STEPHEN RAMON

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 <u>reduced</u> risk of many cancers.^{1,2}

Two established cancer-fighting plant compounds found <u>only</u> 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.



CH₂CH₂CH₂CH₂ Sulforaphane

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}

WHAT YOU NEED TO KNOW

Reduce Cancer 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 **glucoraphanin**, 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 **glucoraphaninmyrosinase** 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 <u>inhibits</u> 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.

References

- Dinkova-Kostova AT, Fahey JW, Kostov RV, et al. KEAP1 and Done? Targeting the NRF2 Pathway with Sulforaphane. *Trends Food Sci Technol.* 2017 Nov;69(Pt B):257-69.
- Verhoeven DT, Goldbohm RA, van Poppel G, et al. Epidemiological studies on brassica vegetables and cancer risk. *Cancer Epidemiol Biomarkers Prev.* 1996 Sep;5(9):733-48.
- Favela-Gonzalez KM, Hernandez-Almanza AY, De la Fuente-Salcido NM. The value of bioactive compounds of cruciferous vegetables (Brassica) as antimicrobials and antioxidants: A review. *J Food Biochem.* 2020 Aug 3:e13414.
- Manchali S, Chidambara Murthy KN, Patil BS. Crucial facts about health benefits of popular cruciferous vegetables. *J Functional Foods*. 2012;4(1):94-106.
- 5. Higdon JV, Delage B, Williams DE, et al. Cruciferous vegetables and human cancer risk: epidemiologic evidence and mechanistic basis. *Pharmacol Res.* 2007 Mar;55(3):224-36.

- Sturm C, Wagner AE. Brassica-Derived Plant Bioactives as Modulators of Chemopreventive and Inflammatory Signaling Pathways. Int J Mol Sci. 2017 Sep 1;18(9).
- Bayat Mokhtari R, Baluch N, Homayouni TS, et al. The role of Sulforaphane in cancer chemoprevention and health benefits: a mini-review. J Cell Commun Signal. 2018 Mar;12(1):91-101.
- Tang L, Zirpoli GR, Guru K, et al. Intake of cruciferous vegetables modifies bladder cancer survival. *Cancer Epidemiol Biomarkers Prev.* 2010 Jul;19(7):1806-11.
- Wu QJ, Yang G, Zheng W, et al. Pre-diagnostic cruciferous vegetables intake and lung cancer survival among Chinese women. *Sci Rep.* 2015 May 19;5:10306.
- Elkashty OA, Tran SD. Sulforaphane as a Promising Natural Molecule for Cancer Prevention and Treatment. *Curr Med Sci.* 2021 Apr;41(2):250-69.
- Hudlikar R, Wang L, Wu R, et al. Epigenetics/Epigenomics and Prevention of Early Stages of Cancer by Isothiocyanates. *Cancer Prev Res (Phila).* 2021 Feb;14(2):151-64.
- Kaiser AE, Baniasadi M, Giansiracusa D, et al. Sulforaphane: A Broccoli Bioactive Phytocompound with Cancer Preventive Potential. *Cancers (Basel)*. 2021 Sep 25;13(19).
- Melim C, Lauro MR, Pires IM, et al. The Role of Glucosinolates from Cruciferous Vegetables (Brassicaceae) in Gastrointestinal Cancers: From Prevention to Therapeutics. *Pharmaceutics*. 2022 Jan 14;14(1).
- Sita G, Graziosi A, Hrelia P, et al. Sulforaphane Causes Cell Cycle Arrest and Apoptosis in Human Glioblastoma U87MG and U373MG Cell Lines under Hypoxic Conditions. *Int J Mol Sci.* 2021 Oct 18;22(20).
- Beaver LM, Kuintzle R, Buchanan A, et al. Long noncoding RNAs and sulforaphane: a target for chemoprevention and suppression of prostate cancer. J Nutr Biochem. 2017 Apr;42:72-83.
- Chinnakannu K, Chen D, Li Y, et al. Cell cycle-dependent effects of 3,3'-diindolylmethane on proliferation and apoptosis of prostate cancer cells. J Cell Physiol. 2009 Apr;219(1):94-9.
- Li J, Zhou Y, Yan Y, et al. Sulforaphane-cysteine downregulates CDK4 /CDK6 and inhibits tubulin polymerization contributing to cell cycle arrest and apoptosis in human glioblastoma cells. *Aging* (*Albany NY*). 2020 Aug 29;12(17):16837-51.
- Pledgie-Tracy A, Sobolewski MD, Davidson NE. Sulforaphane induces cell type-specific apoptosis in human breast cancer cell lines. *Mol Cancer Ther.* 2007 Mar;6(3):1013-21.
- Sundaram MK, R P, Haque S, et al. Dietary isothiocyanates inhibit cancer progression by modulation of epigenome. *Semin Cancer Biol.* 2021 Jan 9.
- Suppipat K, Park CS, Shen Y, et al. Sulforaphane induces cell cycle arrest and apoptosis in acute lymphoblastic leukemia cells. *PLoS One.* 2012;7(12):e51251.



- Thomson CA, Ho E, Strom MB. Chemopreventive properties of 3,3'-diindolylmethane in breast cancer: evidence from experimental and human studies. *Nutr Rev.* 2016 Jul;74(7):432-43.
- Zhang Y, Kensler TW, Cho CG, et al. Anticarcinogenic activities of sulforaphane and structurally related synthetic norbornyl isothiocyanates. *Proc Natl Acad Sci U S A*. 1994 Apr 12;91(8):3147-50.
- Yi H, Li Z, Liu X, et al. Therapeutic Mechanism of Lapatinib Combined with Sulforaphane on Gastric Cancer. *Evid Based Complement Alternat Med.* 2021;2021:9933274.
- 24. Su X, Jiang X, Meng L, et al. Anticancer Activity of Sulforaphane: The Epigenetic Mechanisms and the Nrf2 Signaling Pathway. Oxid Med Cell Longev. 2018;2018:5438179.
- Fahey JW, Holtzclaw WD, Wehage SL, et al. Sulforaphane Bioavailability from Glucoraphanin-Rich Broccoli: Control by Active Endogenous Myrosinase. *PLoS One.* 2015;10(11):e0140963.
- Davis R, Singh KP, Kurzrock R, et al. Sulforaphane inhibits angiogenesis through activation of FOXO transcription factors. *Oncol Rep.* 2009 Dec;22(6):1473-8.
- 27. Kong D, Li Y, Wang Z, et al. Inhibition of angiogenesis and invasion by 3,3'-diindolylmethane is mediated by the nuclear factor-kappaB downstream target genes MMP-9 and uPA that regulated bioavailability of vascular endothelial growth factor in prostate cancer. *Cancer Res.* 2007 Apr 1;67(7):3310-9.
- Russo M, Spagnuolo C, Russo GL, et al. Nrf2 targeting by sulforaphane: A potential therapy for cancer treatment. *Crit Rev Food Sci Nutr.* 2018 May 24;58(8):1391-405.
- Rorke EA, Adhikary G, Szmacinski H, et al. Sulforaphane covalently interacts with the transglutaminase 2 cancer maintenance protein to alter its structure and suppress its activity. *Mol Carcinog.* 2022 Jan;61(1):19-32.
- Conzatti A, Froes FC, Schweigert Perry ID, et al. Clinical and molecular evidence of the consumption of broccoli, glucoraphanin and sulforaphane in humans. *Nutr Hosp.* 2014 Nov 30;31(2):559-69.
- Tortorella SM, Royce SG, Licciardi PV, et al. Dietary Sulforaphane in Cancer Chemoprevention: The Role of Epigenetic Regulation and HDAC Inhibition. *Antioxid Redox Signal.* 2015 Jun 1;22(16):1382-424.
- Wong CP, Hsu A, Buchanan A, et al. Effects of sulforaphane and 3,3'-diindolylmethane on genome-wide promoter methylation in normal prostate epithelial cells and prostate cancer cells. *PLoS One*. 2014;9(1):e86787.
- 33. Cuadrado A, Manda G, Hassan A, et al. Transcription Factor NRF2 as a Therapeutic Target for Chronic Diseases: A Systems Medicine Approach. *Pharmacol Rev.* 2018 Apr;70(2):348-83.
- Houghton CA, Fassett RG, Coombes JS. Sulforaphane and Other Nutrigenomic Nrf2 Activators: Can the Clinician's Expectation Be Matched by the Reality? Oxid Med Cell Longev. 2016;2016:7857186.
- 35. James D, Devaraj S, Bellur P, et al. Novel concepts of broccoli sulforaphanes and disease: induction of phase II antioxidant and detoxification enzymes by enhanced-glucoraphanin broccoli. *Nutr Rev.* 2012 Nov;70(11):654-65.
- 36. Li Y, Kong D, Ahmad A, et al. Antioxidant function of isoflavone and 3,3'-diindolylmethane: are they important for cancer prevention and therapy? Antioxid Redox Signal. 2013 Jul 10;19(2):139-50.
- Heiss E, Herhaus C, Klimo K, et al. Nuclear factor kappa B is a molecular target for sulforaphane-mediated anti-inflammatory mechanisms. *J Biol Chem.* 2001 Aug 24;276(34):32008-15.
- Rahman KM, Ali S, Aboukameel A, et al. Inactivation of NF-kappaB by 3,3'-diindolylmethane contributes to increased apoptosis induced by chemotherapeutic agent in breast cancer cells. *Mol Cancer Ther.* 2007 Oct;6(10):2757-65.
- Dalessandri KM, Firestone GL, Fitch MD, et al. Pilot study: effect of 3,3'-diindolylmethane supplements on urinary hormone metabolites in postmenopausal women with a history of early-stage breast cancer. *Nutr Cancer*. 2004;50(2):161-7.
- Thomson CA, Chow HHS, Wertheim BC, et al. A randomized, placebo-controlled trial of diindolylmethane for breast cancer biomarker modulation in patients taking tamoxifen. *Breast Cancer Res Treat*. 2017 Aug;165(1):97-107.
- Smith S, Sepkovic D, Bradlow HL, et al. 3,3'-Diindolylmethane and genistein decrease the adverse effects of estrogen in LNCaP and PC-3 prostate cancer cells. *J Nutr.* 2008 Dec;138(12):2379-85.
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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**, **coro-nary 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%.

KNOW

Safe, Quick **Constipation Relief**

- Scientists have identified a probiotic strain, Bifidobacterium lactis HN019, that relieves ongoing constipationwithout 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 <u>not</u> 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 slowerthan-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 shortchain 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 *l*ess 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 <u>no</u> 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.

Prescriptions for Constipation

In difficult cases of patients with **chronic idiopathic constipation**, an ongoing condition that does not seem to have a cause, doctors may prescribe either Trulance[®] or Linzess[®]. These drugs are **prosecretory agents** which release water into the intestines and reduce bowel transit time. Without insurance, these drugs can cost over **\$500** for a thirty-day supply.

References

- Available at: https://www.uptodate.com/contents/etiology-andevaluation-of-chronic-constipation-in-adults. Accessed May 9, 2022.
- 2. Sumida K, Molnar MZ, Potukuchi PK, et al. Constipation and risk of death and cardiovascular events. *Atherosclerosis.* 2019 Feb;281:114-20.
- Ishiyama Y, Hoshide S, Mizuno H, et al. Constipation-induced pressor effects as triggers for cardiovascular events. *J Clin Hypertens* (*Greenwich*). 2019 Mar;21(3):421-5.
- Waller PA, Gopal PK, Leyer GJ, et al. Dose-response effect of Bifidobacterium lactis HN019 on whole gut transit time and functional gastrointestinal symptoms in adults. *Scand J Gastroenterol.* 2011 Sep;46(9):1057-64.
- 5. Sundboll J, Thygesen SK, Veres K, et al. Risk of cancer in patients with constipation. *Clin Epidemiol.* 2019;11:299-310.
- Kubota Y, Iso H, Tamakoshi A. Bowel Movement Frequency, Laxative Use, and Mortality From Coronary Heart Disease and Stroke Among Japanese Men and Women: The Japan Collaborative Cohort (JACC) Study. J Epidemiol. 2016 May 5;26(5):242-8.
- Prasad J, Gill H, Smart J, et al. Selection and Characterisation of Lactobacillus and Bifidobacterium Strains for Use as Probiotics. International Dairy Journal. 1998 1998/12/01/;8(12):993-1002.
- Salminen S, Salminen E. Lactulose, lactic acid bacteria, intestinal microecology and mucosal protection. *Scand J Gastroenterol Suppl.* 1997;222:45-8.
- Reigstad CS, Salmonson CE, Rainey JF, 3rd, et al. Gut microbes promote colonic serotonin production through an effect of shortchain fatty acids on enterochromaffin cells. *FASEB J.* 2015 Apr;29(4):1395-403.

- Essien BE, Grasberger H, Romain RD, et al. ZBP-89 regulates expression of tryptophan hydroxylase I and mucosal defense against Salmonella typhimurium in mice. *Gastroenterology.* 2013 Jun;144(7):1466-77, 77 e1-9.
- Spohn SN, Mawe GM. Non-conventional features of peripheral serotonin signalling - the gut and beyond. *Nat Rev Gastroenterol Hepatol.* 2017 Jul;14(7):412-20.
- Ahmed M, Prasad J, Gill H, et al. Impact of consumption of different levels of Bifidobacterium lactis HN019 on the intestinal microflora of elderly human subjects. *J Nutr Health Aging.* 2007 Jan-Feb;11(1):26-31.
- Emmanuel A, Cools M, Vandeplassche L, et al. Prucalopride improves bowel function and colonic transit time in patients with chronic constipation: an integrated analysis. *Am J Gastroenterol.* 2014 Jun;109(6):887-94.
- 14. Available at: https://www.mayoclinic.org/drugs-supplements/prucalopride-oral-route/side-effects/drg-20452343?p=1. Accessed May 5, 2022.
- Available at: https://medlineplus.gov/ency/article/002282.htm. Accessed May 11, 2022.



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FISETIN Promotes Brain Health

BY RICHARD REYNOLDS

Nutrients derived from **plants** have been shown to protect against changes in the **brain** that lead to neurological disorders.

In **2021**, two review articles identified **fisetin** as one of the most promising.^{1,2}

Preclinical studies show the potential of **fisetin** to reduce the impact of **stroke**, mild **cognitive impairment**, and **dementia**.^{1,2}

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 **hyper-phosphorylated tau**. In preclinical models, fisetin *pre-vents* 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 free-radical 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.^{10,11} 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.

References

- 1. Maher P. Preventing and Treating Neurological Disorders with the Flavonol Fisetin. *Brain Plast*. 2021 Feb 9;6(2):155-66.
- Ravula AR, Teegala SB, Kalakotla S, et al. Fisetin, potential flavonoid with multifarious targets for treating neurological disorders: An updated review. *Eur J Pharmacol.* 2021 Nov 5;910:174492.
- 3. Grynkiewicz G, Demchuk OM. New Perspectives for Fisetin. *Front Chem.* 2019;7:697.
- 4. Akay. A cross over pilot pharmacokinetic study of fisetin 1000 mg and formulated fisetin 200 mg administered in a single dose to healthy volunteers. *Manufacturer's study (in press for future publication)*. 2020.
- Wang L, Cao D, Wu H, et al. Fisetin Prolongs Therapy Window of Brain Ischemic Stroke Using Tissue Plasminogen Activator: A Double-Blind Randomized Placebo-Controlled Clinical Trial. *Clin Appl Thromb Hemost.* 2019 Jan-Dec;25:1076029619871359.
- Ahmad A, Ali T, Park HY, et al. Neuroprotective Effect of Fisetin Against Amyloid-Beta-Induced Cognitive/Synaptic Dysfunction, Neuroinflammation, and Neurodegeneration in Adult Mice. *Mol Neurobiol.* 2017 Apr;54(3):2269-85.
- Xiao S, Lu Y, Wu Q, et al. Fisetin inhibits tau aggregation by interacting with the protein and preventing the formation of beta-strands. *Int J Biol Macromol.* 2021 May 1;178:381-93.



- Kim S, Choi KJ, Cho SJ, et al. Fisetin stimulates autophagic degradation of phosphorylated tau via the activation of TFEB and Nrf2 transcription factors. *Sci Rep.* 2016 Apr 26;6:24933.
- Maher P. How fisetin reduces the impact of age and disease on CNS function. *Front Biosci (Schol Ed)*. 2015 Jun 1;7(1):58-82.
- Molagoda IMN, Jayasingha J, Choi YH, et al. Fisetin inhibits lipopolysaccharide-induced inflammatory response by activating beta-catenin, leading to a decrease in endotoxic shock. *Sci Rep.* 2021 Apr 16;11(1):8377.
- Xu M-X, Ge C-X, Li Q, et al. Fisetin nanoparticles protect against PM2.5 exposure-induced neuroinflammation by down-regulation of astrocytes activation related NF-B signaling pathway. *Journal of Functional Foods*. 2020 2020/02/01/;65:103716.
- Currais A, Prior M, Dargusch R, et al. Modulation of p25 and inflammatory pathways by fisetin maintains cognitive function in Alzheimer's disease transgenic mice. *Aging Cell.* 2014 Apr;13(2):379-90.
- Maher P, Akaishi T, Abe K. Flavonoid fisetin promotes ERK-dependent long-term potentiation and enhances memory. *Proc Natl Acad Sci U S A*. 2006 Oct 31;103(44):16568-73.
- He WB, Abe K, Akaishi T. Oral administration of fisetin promotes the induction of hippocampal long-term potentiation in vivo. *J Pharmacol Sci.* 2018 Jan;136(1):42-5.
- Maher P. Modulation of multiple pathways involved in the maintenance of neuronal function during aging by fisetin. *Genes Nutr.* 2009 Dec;4(4):297-307.
- Chrysanthou M, Miro Estruch I, Rietjens I, et al. In Vitro Methodologies to Study the Role of Advanced Glycation End Products (AGEs) in Neurodegeneration. *Nutrients*. 2022 Jan 15;14(2).
- 17. Chen C, Li XH, Tu Y, et al. Abeta-AGE aggravates cognitive deficit in rats via RAGE pathway. *Neuroscience*. 2014 Jan 17;257:1-10.
- Thangthaeng N, Poulose SM, Miller MG, et al. Preserving Brain Function in Aging: The Anti-glycative Potential of Berry Fruit. *Neuromolecular Med.* 2016 Sep;18(3):465-73.
- Kritsilis M, S VR, Koutsoudaki PN, et al. Ageing, Cellular Senescence and Neurodegenerative Disease. *Int J Mol Sci.* 2018 Sep 27;19(10):2937.
- 20. Kirkland JL, Tchkonia T. Cellular Senescence: A Translational Perspective. *EBioMedicine*. 2017 Jul;21:21-8.
- Kirkland JL, Tchkonia T, Zhu Y, et al. The Clinical Potential of Senolytic Drugs. J Am Geriatr Soc. 2017 Oct;65(10):2297-301.
- Zhu Y, Tchkonia T, Pirtskhalava T, et al. The Achilles' heel of senescent cells: from transcriptome to senolytic drugs. *Aging Cell*. 2015 Aug;14(4):644-58.
- 23. Yousefzadeh MJ, Zhu Y, McGowan SJ, et al. Fisetin is a senotherapeutic that extends health and lifespan. *EBioMedicine*. 2018 Oct;36:18-28.

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References

- 1. Anticancer Agents Med Chem. 2013 Sep;13(7):967-70.
- 2. Nutr J. 2016;15:6. 3. Obesity (Silver Spring). 2014;22(1):63-71.



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

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- Targets longevity pathways²⁻⁶
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- Suppresses excess mTOR activation⁸

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- References
 1. Available at: https://www.mayo.edu/
 research/clinical-trials/cls-20438802.
 Accessed June 22, 2020.
 2. Life Sci. 2018 Jan 15;193:171-9.
 3. Mini Rev Med Chem. 2018;18(13):1151-7.
 4. Nutr Res Pract. 2017 Oct;11(5):430-4.
 5. Biochem Biophys Res Commun. 2015
 Nov 27;467(4):638-44.
 6. Int Immunopharmacol. 2017 Apr;45:135-47.
 7. EBioMedicine. 2018 Oct;35:18-28.
 J. Ukut Biochem. 2013 Aug;24(8):154-754.
- J Nutr Biochem. 2013 Aug;24(8):1547-54. Manufacturer's study (in press for future publication). 2020.

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GLUTEN ERFE

2021 AGING CONFERENCES





BY BEN BEST

Research has increasingly been finding ways to reverse aging in laboratory animals, and much of this research could also benefit humans. Some age-reversal therapies are currently being used by self-experimenters.

During the last year or two, scientists have continued to work and think about solving 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 diseases.

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.



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FAHY









GORBUNOVA

VERDIN

References

- Amaya-Montoya M, Perez-Londono A, Guatibonza-Garcia V, et al. Cellular Senescence as a Therapeutic Target for Age-Related Diseases: A Review. Adv Ther. 2020 Apr;37(4):1407-24.
- Suda M, Shimizu I, Katsuumi G, et al. Senolytic vaccination improves normal and pathological age-related phenotypes and increases lifespan in progeroid mice. *Nature Aging.* 2021 2021/12/01;1(12):1117-26.
- Xu M, Pirtskhalava T, Farr JN, et al. Senolytics improve physical function and increase lifespan in old age. *Nat Med.* 2018 Aug;24(8):1246-56.
- Martinez-Zamudio RI, Dewald HK, Vasilopoulos T, et al. Senescence-associated beta-galactosidase reveals the abundance of senescent CD8+ T cells in aging humans. *Aging Cell.* 2021 May;20(5):e13344.
- Lehmann J, Baar MP, de Keizer PLJ. Senescent Cells Drive Frailty through Systemic Signals. *Trends Mol Med.* 2018 Nov;24(11):917-8.
- De Blander H, Morel AP, Senaratne AP, et al. Cellular Plasticity: A Route to Senescence Exit and Tumorigenesis. *Cancers (Basel).* 2021 Sep 11;13(18).
- Baskar R, Lee KA, Yeo R, et al. Cancer and radiation therapy: current advances and future directions. Int J Med Sci. 2012;9(3):193-9.
- 8. Liu B, Ezeogu L, Zellmer L, et al. Protecting the normal in order to better kill the cancer. *Cancer Med.* 2015 Sep;4(9):1394-403.
- Simay Demir YD, Ozdemir A, Sucularli C, et al. The implication of ROCK 2 as a potential senotherapeutic target via the suppression of the harmful effects of the SASP: Do senescent cancer cells really engulf the other cells? *Cell Signal*. 2021 Aug;84:110007.
- Liu XL, Ding J, Meng LH. Oncogene-induced senescence: a double edged sword in cancer. Acta Pharmacol Sin. 2018 Oct;39(10): 1553-8.
- Baar MP, Brandt RMC, Putavet DA, et al. Targeted Apoptosis of Senescent Cells Restores Tissue Homeostasis in Response to Chemotoxicity and Aging. *Cell.* 2017 Mar 23;169(1):132-47 e16.
- van Willigenburg H, de Keizer PLJ, de Bruin RWF. Cellular senescence as a therapeutic target to improve renal transplantation outcome. *Pharmacol Res.* 2018 Apr;130:322-30.
- Gorbunova V, Seluanov A, Mita P, et al. The role of retrotransposable elements in ageing and age-associated diseases. *Nature*. 2021 Aug;596(7870):43-53.
- 14. Schumacher B, Pothof J, Vijg J, et al. The central role of DNA damage in the ageing process. *Nature.* 2021 Apr;592(7856):695-703.
- Simon M, Van Meter M, Ablaeva J, et al. LINE1 Derepression in Aged Wild-Type and SIRT6-Deficient Mice Drives Inflammation. *Cell Metab.* 2019 Apr 2;29(4):871-85 e5.
- Wu X, Sun M, Yang Z, et al. The Roles of CCR9/CCL25 in Inflammation and Inflammation-Associated Diseases. *Front Cell Dev Biol.* 2021;9:686548.
- Guerrero A, De Strooper B, Arancibia-Carcamo IL. Cellular senescence at the crossroads of inflammation and Alzheimer's disease. *Trends Neurosci.* 2021 Sep;44(9):714-27.
- Serrano-Lopez J, Martin-Antonio B. Inflammaging, an Imbalanced Immune Response That Needs to Be Restored for Cancer Prevention and Treatment in the Elderly. *Cells.* 2021 Sep 28;10(10).
- Grabowska W, Sikora E, Bielak-Zmijewska A. Sirtuins, a promising target in slowing down the ageing process. *Biogerontology.* 2017 Aug;18(4):447-76.
- 20. Tian X, Firsanov D, Zhang Z, et al. SIRT6 Is Responsible for More Efficient DNA Double-Strand Break Repair in Long-Lived Species. *Cell.* 2019 Apr 18;177(3):622-38 e22.
- Roichman A, Elhanati S, Aon MA, et al. Restoration of energy homeostasis by SIRT6 extends healthy lifespan. *Nat Commun.* 2021 May 28;12(1):3208.
- Van Meter M, Gorbunova V, Seluanov A. SIRT6: a promising target for cancer prevention and therapy. *Adv Exp Med Biol.* 2014;818:181-96.
- 23. Akter R, Afrose A, Rahman MR, et al. A Comprehensive Analysis into the Therapeutic Application of Natural Products as SIRT6 Modulators in Alzheimer's Disease, Aging, Cancer, Inflammation, and Diabetes. *Int J Mol Sci.* 2021 Apr 17;22(8).

- 24. Shapouri-Moghaddam A, Mohammadian S, Vazini H, et al. Macrophage plasticity, polarization, and function in health and disease. *J Cell Physiol.* 2018 Sep;233(9):6425-40.
- 25. Yunna C, Mengru H, Lei W, et al. Macrophage M1/M2 polarization. *Eur J Pharmacol.* 2020 Jun 15;877:173090.
- 26. Yarbro JR, Emmons RS, Pence BD. Macrophage Immunometabolism and Inflammaging: Roles of Mitochondrial Dysfunction, Cellular Senescence, CD38, and NAD. *Immunometabolism.* 2020;2(3):e200026.
- Childs BG, Gluscevic M, Baker DJ, et al. Senescent cells: an emerging target for diseases of ageing. *Nat Rev Drug Discov.* 2017 Oct;16(10):718-35.
- Campbell RA, Docherty MH, Ferenbach DA, et al. The Role of Ageing and Parenchymal Senescence on Macrophage Function and Fibrosis. *Front Immunol.* 2021;12:700790.
- Covarrubias AJ, Kale A, Perrone R, et al. Senescent cells promote tissue NAD(+) decline during ageing via the activation of CD38(+) macrophages. *Nat Metab.* 2020 Nov;2(11):1265-83.
- Hogan KA, Chini CCS, Chini EN. The Multi-faceted Ecto-enzyme CD38: Roles in Immunomodulation, Cancer, Aging, and Metabolic Diseases. *Front Immunol.* 2019;10:1187.
- 31. Behmoaras J, Gil J. Similarities and interplay between senescent cells and macrophages. *J Cell Biol.* 2021 Feb 1;220(2).
- 32. Sharma R. Perspectives on the dynamic implications of cellular senescence and immunosenescence on macrophage aging biology. *Biogerontology.* 2021 Dec;22(6):571-87.
- 33. Fahy GM, Brooke RT, Watson JP, et al. Reversal of epigenetic aging and immunosenescent trends in humans. *Aging Cell.* 2019 Dec;18(6):e13028.
- 34. French RA, Broussard SR, Meier WA, et al. Age-associated loss of bone marrow hematopoietic cells is reversed by GH and accompanies thymic reconstitution. *Endocrinology*. 2002 Feb;143(2):690-9.
- Goff BL, Roth JA, Arp LH, et al. Growth hormone treatment stimulates thymulin production in aged dogs. *Clin Exp Immunol.* 1987 Jun;68(3):580-7.
- Napolitano LA, Schmidt D, Gotway MB, et al. Growth hormone enhances thymic function in HIV-1-infected adults. *J Clin Invest.* 2008 Mar;118(3):1085-98.
- Mannick JB, Del Giudice G, Lattanzi M, et al. mTOR inhibition improves immune function in the elderly. *Sci Transl Med.* 2014 Dec 24;6(268):268ra179.
- Naylor K, Li G, Vallejo AN, et al. The influence of age on T cell generation and TCR diversity. J Immunol. 2005 Jun 1;174(11):7446-52.
- 39. Abbott A. First hint that body's 'biological age' can be reversed. *Nature.* 2019 Sep;573(7773):173.
- 40. Fulop T, Larbi A, Pawelec G, et al. Immunology of Aging: the Birth of Inflammaging. *Clin Rev Allergy Immunol.* 2021 Sep 18:1-14.
- Sadagurski M, Landeryou T, Cady G, et al. Growth hormone modulates hypothalamic inflammation in long-lived pituitary dwarf mice. *Aging Cell.* 2015 Dec;14(6):1045-54.
- Lu AT, Quach A, Wilson JG, et al. DNA methylation GrimAge strongly predicts lifespan and healthspan. *Aging (Albany NY)*. 2019 Jan 21;11(2):303-27.
- Hannum G, Guinney J, Zhao L, et al. Genome-wide methylation profiles reveal quantitative views of human aging rates. *Mol Cell.* 2013 Jan 24;49(2):359-67.
- 44. Horvath S. DNA methylation age of human tissues and cell types. *Genome Biol.* 2013;14(10):R115.
- Levine ME, Lu AT, Quach A, et al. An epigenetic biomarker of aging for lifespan and healthspan. *Aging (Albany NY).* 2018 Apr 18;10(4):573-91.
- Fang EF, Lautrup S, Hou Y, et al. NAD(+) in Aging: Molecular Mechanisms and Translational Implications. *Trends Mol Med.* 2017 Oct;23(10):899-916.
- Putin E, Mamoshina P, Aliper A, et al. Deep biomarkers of human aging: Application of deep neural networks to biomarker development. *Aging (Albany NY)*. 2016 May;8(5):1021-33.
- Mamoshina P, Kochetov K, Putin E, et al. Population Specific Biomarkers of Human Aging: A Big Data Study Using South Korean, Canadian, and Eastern European Patient Populations. J Gerontol A Biol Sci Med Sci. 2018 Oct 8;73(11):1482-90.

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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 **silymarin**, 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 <u>reduce</u> 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.

References

- 1. Available at: https://www.nytimes. com/2018/07/18/health/cirrhosis-liver-cancer. html. Accessed March 28, 2022.
- Tapper EB, Parikh ND. Mortality due to cirrhosis and liver cancer in the United States, 1999-2016: observational study. *BMJ*. 2018;362:k2817.
- Marchesini G, Petta S, Dalle Grave R. Diet, weight loss, and liver health in nonalcoholic fatty liver disease: Pathophysiology, evidence, and practice. *Hepatology*. 2016 Jun;63(6):2032-43.



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

- Cheemerla S, Balakrishnan M. Global Epidemiology of Chronic Liver Disease. *Clin Liver Dis* (Hoboken). 2021 May;17(5):365-70.
- Ferenci P, Dragosics B, Dittrich H, et al. Randomized controlled trial of silymarin treatment in patients with cirrhosis of the liver. *J Hepatol.* 1989 Jul;9(1):105-13.
- Federico A, Dallio M, Loguercio C. Silymarin/ Silybin and Chronic Liver Disease: A Marriage of Many Years. *Molecules*. 2017 Jan 24;22(2).
- Vailati A, Aristia L, Sozze E, et al. Randomized open study of the dose-effect relationship of a short course of IdB 1016 in patients with viral or alcoholic hepatitis. *Fitoterapia*. 1993;64(3):219-28.
- Salmi HA, Sarna S. Effect of silymarin on chemical, functional, and morphological alterations of the liver. A double-blind controlled study. *Scand J Gastroenterol*. 1982 Jun;17(4):517-21.
- Le MH, Devaki P, Ha NB, et al. Prevalence of non-alcoholic fatty liver disease and risk factors for advanced fibrosis and mortality in the United States. *PLoS One*. 2017;12(3):e0173499.
- Available at: https://www.uptodate.com/contents/cirrhosis-beyond-the-basics. Accessed May 10, 2022.
- 11. Calzadilla Bertot L, Adams LA. The Natural Course of Non-Alcoholic Fatty Liver Disease. Int J Mol Sci. 2016 May 20;17(5):774.
- Musso G, Gambino R, Cassader M, et al. Metaanalysis: natural history of non-alcoholic fatty liver disease (NAFLD) and diagnostic accuracy of non-invasive tests for liver disease severity. *Ann Med.* 2011 Dec;43(8):617-49.
- Noureddin M, Vipani A, Bresee C, et al. NASH Leading Cause of Liver Transplant in Women: Updated Analysis of Indications For Liver Transplant and Ethnic and Gender Variances. Am J Gastroenterol. 2018 Nov;113(11):1649-59.
- Cacciapuoti F, Scognamiglio A, Palumbo R, et al. Silymarin in non alcoholic fatty liver disease. World J Hepatol. 2013 Mar 27;5(3):109-13.
- Federico A, Trappoliere M, Tuccillo C, et al. A new silybin-vitamin E-phospholipid complex improves insulin resistance and liver damage in patients with non-alcoholic fatty liver disease: preliminary observations. *Gut.* 2006 Jun;55(6):901-2.

- Loguercio C, Andreone P, Brisc C, et al. Silybin combined with phosphatidylcholine and vitamin E in patients with nonalcoholic fatty liver disease: a randomized controlled trial. *Free Radic Biol Med.* 2012 May 1;52(9):1658-65.
- Zhong S, Fan Y, Yan Q, et al. The therapeutic effect of silymarin in the treatment of nonalcoholic fatty disease: A meta-analysis (PRISMA) of randomized control trials. *Medicine (Baltimore)*. 2017 Dec;96(49):e9061.
- Huseini HF, Larijani B, Heshmat R, et al. The efficacy of Silybum marianum (L.) Gaertn. (silymarin) in the treatment of type II diabetes: a randomized, double-blind, placebocontrolled, clinical trial. *Phytother Res.* 2006 Dec;20(12):1036-9.
- Clichici S, Olteanu D, Nagy AL, et al. Silymarin inhibits the progression of fibrosis in the early stages of liver injury in CCl(4)-treated rats. J Med Food. 2015 Mar;18(3):290-8.
- Kidd P, Head K. A review of the bioavailability and clinical efficacy of milk thistle phytosome: a silybin-phosphatidylcholine complex (Siliphos). *Altern Med Rev.* 2005 Sep;10(3):193-203.
- Younossi Z, Anstee QM, Marietti M, et al. Global burden of NAFLD and NASH: trends, predictions, risk factors and prevention. *Nat Rev Gastroenterol Hepatol*. 2018 Jan;15(1):11-20.
- Available at: https://www.niddk.nih.gov/healthinformation/liver-disease/nafld-nash/definitionfacts. Accessed June 8, 2022.
- Estes C, Razavi H, Loomba R, et al. Modeling the epidemic of nonalcoholic fatty liver disease demonstrates an exponential increase in burden of disease. *Hepatology*. 2018 Jan;67(1):123-33.
- Alkhouri N, Scott A. An Update on the Pharmacological Treatment of Nonalcoholic Fatty Liver Disease: Beyond Lifestyle Modifications. *Clin Liver Dis (Hoboken)*. 2018 Apr;11(4):82-6.
- Mofidi F, Poustchi H, Yari Z, et al. Synbiotic supplementation in lean patients with non-alcoholic fatty liver disease: a pilot, randomised, double-blind, placebo-controlled, clinical trial. *Br J Nutr.* 2017 Mar;117(5):662-8.



- Eslamparast T, Poustchi H, Zamani F, et al. Synbiotic supplementation in nonalcoholic fatty liver disease: a randomized, double-blind, placebo-controlled pilot study. *Am J Clin Nutr.* 2014 Mar;99(3):535-42.
- Magosso E, Ansari MA, Gopalan Y, et al. Tocotrienols for normalisation of hepatic echogenic response in nonalcoholic fatty liver: a randomised placebo-controlled clinical trial. *Nutr J.* 2013 Dec 27;12(1):166.
- Pervez MA, Khan DA, Ijaz A, et al. Effects of Delta-tocotrienol Supplementation on Liver Enzymes, Inflammation, Oxidative stress and Hepatic Steatosis in Patients with Nonalcoholic Fatty Liver Disease. *Turk J Gastroenterol.* 2018 Mar;29(2):170-6.
- Pervez MA, Khan DA, Slehria AUR, et al. Delta-tocotrienol supplementation improves biochemical markers of hepatocellular injury and steatosis in patients with nonalcoholic fatty liver disease: A randomized, placebocontrolled trial. *Complement Ther Med*. 2020 Aug;52:102494.
- Dajani Al, Abuhammour A. Agents for the treatment of fatty liver disease: focus on essential phospholipids. *Drugs & Therapy Perspectives*. 2021;37(6):249-64.
- Maev IV, Samsonov AA, Palgova LK, et al. Effectiveness of phosphatidylcholine as adjunctive therapy in improving liver function tests in patients with non-alcoholic fatty liver disease and metabolic comorbidities: real-life observational study from Russia. *BMJ Open Gastroenterol.* 2020;7(1):e000368.
- Maev IV, Samsonov AA, Palgova LK, et al. Effectiveness of phosphatidylcholine in alleviating steatosis in patients with non-alcoholic fatty liver disease and cardiometabolic comorbidities (MANPOWER study). *BMJ Open Gastroenterol.* 2020;7(1):e000341.
- Millea PJ. N-acetylcysteine: multiple clinical applications. *Am Fam Physician*. 2009 Aug 1;80(3):265-9.
- Bajt ML, Knight TR, Lemasters JJ, et al. Acetaminophen-induced oxidant stress and cell injury in cultured mouse hepatocytes: protection by N-acetyl cysteine. *Toxicol Sci.* 2004 Aug;80(2):343-9.
- de Oliveira CP, Simplicio FI, de Lima VM, et al. Oral administration of S-nitroso-N-acetylcysteine prevents the onset of non alcoholic fatty liver disease in rats. *World J Gastroenterol.* 2006 Mar 28;12(12):1905-11.
- Mehta K, Van Thiel DH, Shah N, et al. Nonalcoholic fatty liver disease: pathogenesis and the role of antioxidants. *Nutr Rev.* 2002 Sep;60(9):289-93.

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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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Sesame Lignans, Olive Extract, Krill & Astaxanthin

Super Omega-3 Plus EPA/DHA Fish Oil,

- Super Omega-3 EPA/DHA Fish Oil, Sesame Lignans & 01985 Olive Extract • 60 enteric coated softgels
- 01984 Super Omega-3 EPA/DHA Fish Oil, Sesame Lignans & Olive Extract • 120 enteric coated softgels
- 01986 Super Omega-3 EPA/DHA Fish Oil, Sesame Lignans & Olive Extract • 240 softgels
- 01812 Provinal[®] Purified Omega-7 Vegetarian DHA

01640

FOOD

- 02008 California Estate Extra Virgin Olive Oil
- Rainforest Blend Decaf Ground Coffee 02170
- **Rainforest Blend Ground Coffee** 02169
- 02171 Rainforest Blend Whole Bean Coffee
- 00438 Stevia[™] Organic Liquid Sweetner Stevia[™] Sweetener 00432
- **GLUCOSE MANAGEMENT**
- CinSulin[®] with InSea^{2®} and Crominex[®] 3+ 01503
- CoffeeGenic® Green Coffee Extract 01620
- Glycemic Guard[™] 02122
- 00925 Mega Benfotiamine
- 01803 Tri Sugar Shield®

HEART HEALTH

- 01066 Aspirin (Enteric Coated)
- 01842 BioActive Folate & Vitamin B12 Caps
- 01700 Cardio Peak™
- 02121 Homocysteine Resist
- 02018 Optimized Carnitine
- Super-Absorbable CoQ10 Ubiquinone with 01949 d-Limonene • 50 mg, 60 softgels
- 01951 Super-Absorbable CoQ10 Ubiquinone with d-Limonene • 100 mg, 60 softgels
- 01929 Super Ubiquinol CoQ10
- 01427 Super Ubiquinol CoQ10 with Enh Mitochondrial Support[™] • 50 mg, 30 softgels
- 01425 Super Ubiquinol CoQ10 with Enh Mitochondrial Support[™] • 50 mg, 100 softgels
- 01437 Super Ubiguinol CoQ10 with Enh Mitochondrial Support[™] • 100 mg, 30 softgels
- 01426 Super Ubiguinol CoQ10 with Enh Mitochondrial Support[™] • 100 mg, 60 softgels
- 01431 Super Ubiquinol CoQ10 with Enh Mitochondrial Support[™] • 200 mg, 30 softgels
- 01733 Super Ubiquinol CoQ10 with PQQ
- 01859 TMG Liquid Capsules
- 00349 TMG Powder

HORMONE BALANCE

- 00454 DHEA 15 mg, 100 capsules
- 00335 DHEA • 25 mg, 100 capsules
- 00882 DHEA • 50 mg, 60 capsules
- DHEA 25 mg, 100 vegetarian dissolve in mouth tablets 00607
- 01689 DHEA • 100 mg, 60 veg capsules
- 02368 Optimized Broccoli and Cruciferous Blend
- 00302 Pregnenolone • 50 mg, 100 capsules
- 00700 Pregnenolone • 100 mg, 100 capsules
- 01468 Triple Action Cruciferous Vegetable Extract
- 01469 Triple Action Cruciferous Vegetable Extract and Resveratrol

IMMUNE SUPPORT

- 02411 5 Day Elderberry Immune
- 00681 **AHCC®**
- 02302 Bio-Quercetin®
- 02410 Black Elderberry + Vitamin C
- Enhanced Zinc Lozenges 01961
- 01704 Immune Modulator with Tinofend®
- 02425 Immune Packs with Vitamin C & D, Zinc and Probiotic

- 02005 Immune Senescence Protection Formula™
- 00316 Kyolic[®] Garlic Formula 102
- 00789 Kyolic[®] Reserve
- 01681 Lactoferrin (Apolactoferrin) Caps
- 02426 Mushroom Immune with Beta Glucans
- NK Cell Activator™ 01903
- 01394 **Optimized Garlic**
- 01309 **Optimized Quercetin** 01811
- Peony Immune 00525
- ProBoost Thymic Protein A 01708 Reishi Extract Mushroom Complex
- 01906
- Standardized Cistanche
- 13685 Ten Mushroom Formula® 01097
- Ultra Soy Extract 01561 Zinc Lozenges

INFLAMMATION MANAGEMENT

- 01639 5-LOX Inhibitor with AprèsFlex®
- 02324 Advanced Curcumin Elite™
- **Turmeric Extract, Ginger & Turmerones** 01709 Black Cumin Seed Oil
- 02310 Black Cumin Seed Oil and Curcumin Elite™
- 00202 Boswella
- 02467 Curcumin Elite[™] Turmeric Extract • 30 veg capsules
- Curcumin Elite[™] Turmeric Extract 60 veg capsules 02407
- 01804 Cytokine Suppress[®] with EGCG
- 02223 **Pro-Resolving Mediators**
- 00318 Serraflazyme
- 01203 Specially-Coated Bromelain
- 00407 Super Bio-Curcumin® Turmeric Extract
- 01254 Zyflamend[™] Whole Body

JOINT SUPPORT

- 02404 Arthro-Immune Joint Support
- 02238 ArthroMax[®] Advanced NT2 Collagen[™] & AprèsFlex[®]
- 01617 ArthroMax[®] with Theaflavins & AprèsFlex[®]
- 02138 ArthroMax® Elite
- 00965 Fast-Acting Joint Formula
- 02430 Fast Acting Relief
- 00522 Glucosamine/Chondroitin Capsules
- 02420 Glucosamine Sulfate
- 02424 Joint Mobility
- 01600 Krill Healthy Joint Formula
- 01050 Krill Oil

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01651

01571

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02402

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01393

01608

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00342

01080

01884

02361

- 00451 MSM (Methylsulfonylmethane)
- 02231 NT2 Collagen™

Uric Acid Control

- **KIDNEY & BLADDER SUPPORT**
- 00862 Cran-Max[®] Cranberry Whole Fruit Concentrate

Water-Soluble Pumpkin Seed Extract

Advanced Milk Thistle • 60 softgels

Advanced Milk Thistle • 120 softgels

PectaSol-C® Modified Citrus Pectin Powder

PectaSol-C® Modified Citrus Pectin Capsules

01424 Optimized Cran-Max[®] with Ellirose[™]

LIVER HEALTH & DETOXIFICATION

Milk Thistle • 60 veg capsules

FLORASSIST[®] Liver Restore[™]

Glutathione, Cysteine & C

Liver Efficiency Formula

N-Acetyl-L-Cysteine

Anti-Alcohol Complex

Calcium D-Glucarate

Chlorophyllin

HepatoPro

Silymarin

SOD Booster

LONGEVITY & WELLNESS

- 00457 Alpha-Lipoic Acid
- 01625 AppleWise
- 02414 **Bio-Fisetin**
- 01214 **Blueberry Extract**
- 01438 Blueberry Extract and Pomegranate
- 02270 DNA Protection Formula
- 02431 Essential Youth L-Ergothioneine
- 02119 GEROPROTECT® Ageless Cell™ 02415 GEROPROTECT® Autophagy Renew
- 02133 GEROPROTECT[®] Longevity A.I.™
- 02401 GEROPROTECT® Stem Cell
- 02211 Grapeseed Extract
- 00954 Mega Green Tea Extract (decaffeinated)
- 00953 Mega Green Tea Extract (lightly caffeinated)
- 01513 Optimized Fucoidan with Maritech® 926
- 02230 Optimized Resveratrol
- 01637 Pycnogenol® French Maritime Pine Bark Extract
- 02210 Resveratrol
- 00070 RNA (Ribonucleic Acid)
- 02301 Senolytic Activator®
- 01208 Super R-Lipoic Acid
- 01919 X-R Shield

MEN'S HEALTH

- 02209 Male Vascular Sexual Support
- 00455 Mega Lycopene Extract
- 02306 Men's Bladder Control
- 01789 PalmettoGuard[®] Saw Palmetto and Beta-Sitosterol
- 01790 PalmettoGuard[®] Saw Palmetto/Nettle Root Formula and Beta-Sitosterol
- 01837 Pomi-T®
- Prelox[®] Enhanced Sex for Men 01373
- 01940 Super MiraForte with Standardized Lignans
- 02500 **Testosterone Elite**
- 01909 Triple Strength ProstaPollen™
- 02029 Ultra Prostate Formula

MINERALS

- 01661 Boron
- 02107 **Extend-Release Magnesium**
- 01677 Iron Protein Plus
- 02403 Lithium
- 01459 Magnesium Caps
- 01682 Magnesium (Citrate)
- **Only Trace Minerals** 01328
- Optimized Chromium with Crominex® 3+ 01504
- Potassium with Extend-Release Magnesium 02309
- Sea-lodine[™] 01740
- 01879 Se-Methyl L-Selenocysteine
- 01778 Super Selenium Complex
- 00213 Vanadyl Sulfate
- 01813 Zinc Caps

MISCELLANEOUS

- 00577 Potassium lodide
- 00657 Solarshield[®] Sunglasses

MOOD & STRESS MANAGEMENT

- 02434 Calm-Mag
- 02312 **Cortisol-Stress Balance**
- 00987 **Enhanced Stress Relief** 5 HTP 01074
- 01683
- L-Theanine 02175
- SAMe (S-Adenosyl-Methionine) 200 mg, 30 enteric coated vegetarian tablets 02176 SAMe (S-Adenosyl-Methionine)
- 400 mg, 30 enteric coated vegetarian tablets 02174 SAMe (S-Adenosyl-Methionine)
- 400 mg, 60 enteric coated vegetarian tablets
- 02429 Theanine XR[™] Stress Relief

MULTIVITAMINS

- 02199 Children's Formula Life Extension Mix™
- 02354 Life Extension Mix[™] Capsules
- 02364 Life Extension Mix[™] Capsules without Copper
- 02356 Life Extension Mix[™] Powder
- 02355 Life Extension Mix[™] Tablets
- 02357 Life Extension Mix[™] Tablets with Extra Niacin
- 02365 Life Extension Mix[™] Tablets without Copper
- 02292 Once-Daily Health Booster 30 softgels
- 02291 Once-Daily Health Booster 60 softgels
- 02313 One-Per-Day Tablets
- 02428 Plant-Based Multivitamin
- 02317 Two-Per-Day Capsules 60 capsules
- 02314 Two-Per-Day Capsules 120 capsules
- 02316 Two-Per-Day Tablets 60 tablets
- 02315 Two-Per-Day Tablets 120 tablets

NERVE & COMFORT SUPPORT

- 02202 ComfortMAX[™]
- 02303 Discomfort Relief

PERSONAL CARE

- 01006 Biosil[™] 5 mg, 30 veg capsules
- Biosil[™] 1 fl oz 01007
- 00321 Dr. Proctor's Advanced Hair Formula
- 00320 Dr. Proctor's Shampoo
- 02322 Hair, Skin & Nails Collagen Plus Formula
- 01278 Life Extension Toothpaste
- 00408 Venotone
- 00409 Xyliwhite Mouthwash
- 02304 Youthful Collagen
- 02252 Youthful Legs

PET CARE

- 01932 Cat Mix
- 01931 Dog Mix

02125

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02208

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02203

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80165

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80151

SKIN CARE

PROBIOTICS

- 01622 **Bifido GI Balance**
- 01825 FLORASSIST® Balance 02421 FLORASSIST® Daily Bowel Regularity

FLORASSIST® Heart Health

FLORASSIST® Oral Hygiene

FLORASSIST® Throat Health

Jarro-Dophilus® for Women

FLORASSIST[®] Prebiotic

Theralac® Probiotics

TruFlora[®] Probiotics

FLORASSIST® Mood Improve

FLORASSIST® GI with Phage Technology

FLORASSIST® Immune & Nasal Defense

FLORASSIST® Winter Immune Support

Jarro-Dophilus EPS® • 60 veg capsules

Jarro-Dophilus EPS® • 120 veg capsules

Advanced Anti-Glycation Peptide Serum

Advanced Probiotic-Fermented Eye Serum

Advanced Under Eye Serum with Stem Cells

Advanced Growth Factor Serum

Advanced Lightening Cream

Advanced Retinol Serum

80152 Advanced Triple Peptide Serum

Anti-Aging Mask

Advanced Hyaluronic Acid Serum

Advanced Peptide Hand Therapy

All-Purpose Soothing Relief Cream

Anti-Aging Rejuvenating Face Cream

Amber Self MicroDermAbrasion

80153 Anti-Aging Rejuvenating Scalp Serum

- Brightening Peptide Serum 80179 Collagen Boosting Peptide Cream 80176 80156 Collagen Boosting Peptide Serum 02408 Collagen Peptides for Skin & Joints 80180 CoQ10 and Stem Cell Rejuvenation Cream 80169 Cucumber Hydra Peptide Eye Cream 02423 Daily Skin Defense 80141 **DNA Support Cream** 80163 Eye Lift Cream 80123 Face Rejuvenating Anti-Oxidant Cream 80109 Hyaluronic Facial Moisturizer Hyaluronic Oil-Free Facial Moisturizer 80110 80138 Hydrating Anti-Oxidant Facial Mist 00661 Hydroderm 55495 Instensive Moisturizing Cream 80103 Lifting & Tightening Complex 80168 Melatonin Advanced Peptide Cream 80114 Mild Facial Cleanser 80172 Multi Stem Cell Hydration Cream Multi Stem Cell Skin Tightening Complex 80159 80122 Neck Rejuvenating Anti-Oxidant Cream Purifying Facial Mask 80174 80150 **Renewing Eye Cream Resveratrol Anti-Oxidant Serum** 80142 01938 Shade Factor™ 02129 Skin Care Collection Anti-Aging Serum 02130 Skin Care Collection Day Cream 02131 Skin Care Collection Night Cream 80166 Skin Firming Complex 02096 Skin Restoring Ceramides 80130 Skin Stem Cell Serum 80164 Skin Tone Equalizer 80143 Stem Cell Cream with Alpine Rose 80148 Tightening & Firming Neck Cream 80161 Triple-Action Vitamin C Cream 80162 Ultimate MicroDermabrasion 80173 Ultimate Peptide Serum 80178 Ultimate Telomere Cream Ultra Eyelash Booster 80160 80101 Ultra Wrinkle Relaxer 80113 Under Eye Refining Serum 80104 Under Eye Rescue Cream 80171 Vitamin C Lip Rejuvenator 80129 Vitamin C Serum 80136 Vitamin D Lotion 80102 Vitamin K Cream **SLEEP**
- 01512 Bioactive Milk Peptides
- 02300 Circadian Sleep
- 01551 Enhanced Sleep with Melatonin
- 01511 Enhanced Sleep without Melatonin
- 02234 Fast-Acting Liquid Melatonin
- 01669 Glycine
- 02308 Herbal Sleep PM
- 01722 L-Tryptophan
- 01668 Melatonin 300 mcg, 100 veg capsules
- 01083 Melatonin 500 mcg, 200 veg capsules
- 00329 Melatonin 1 mg, 60 capsules
- 02503 Melatonin 3 mg, 60 gummies
- 00330 Melatonin 3 mg, 60 veg capsules 00331 Melatonin • 10 mg, 60 veg capsules
- 00331 Melatonin 10 mg, 60 veg capsules 00332 Melatonin • 3 mg, 60 veg lozenges
- 02201 Melatonin IR/XR
- 01787 Melatonin 6 Hour Timed Release 300 mcg, 100 veg tablets
- 01788 Melatonin 6 Hour Timed Release 750 mcg, 60 veg tablets

- 01786 Melatonin 6 Hour Timed Release 3 mg, 60 veg tablets
- 01721 Optimized Tryptophan Plus 01444 Quiet Sleep
- 01445 Quiet Sleep Melatonin
- 02502 Rest & Renew

VITAMINS

- 01533 Ascorbyl Palmitate 00920 Benfotiamine with Thiamine
- 00664 Beta-Carotene
- 01945 BioActive Complete B-Complex
- 00102 Biotin
- 00084 Buffered Vitamin C Powder
- 02229 Fast-C[®] and Bio-Quercetin Phytosome 02075 Gamma E Mixed Tocopherol Enhanced with Sesame Lignans
- 02070 Gamma E Mixed Tocopherol & Tocotrienols
- 01913 High Potency Optimized Folate
- 01674 Inositol Caps
- 02244 Liquid Vitamin D3 50 mcg (2000 IU)
- 02232 Liquid Vitamin D3 (Mint) 50 mcg (2000 IU)
- 01936 Low-Dose Vitamin K2
- 00065 MK-7
- 00373 No Flush Niacin
- 01939 Optimized Folate (L-Methylfolate)
- 01217 Pyridoxal 5'-Phosphate Caps
- 01400 Super Absorbable Tocotrienols
- 02334 Super K
- 01863 Super Vitamin E
- 02422 Vegan Vitamin D3
- 02028 Vitamin B5 (Pantothenic Acid)
- 01535 Vitamin B6
- 00361 Vitamin B12 Methylcobalamin
- 01536 Vitamin B12 Methylcobalamin 1 mg, 60 veg lozenges
- 01537 Vitamin B12 Methylcobalamin 5 mg, 60 veg lozenges
- 02228 Vitamin C and Bio-Quercetin Phytosome 60 veg tablets
- 02227 Vitamin C and Bio-Quercetin Phytosome 250 veg tablets
- 01753 Vitamin D3 25 mcg (1000 IU), 90 softgels
- 01751 Vitamin D3 25 mcg (1000 IU), 250 softgels
- 01713 Vitamin D3 125 mcg (5000 IU), 60 softgels
- 01718 Vitamin D3 175 mcg (7000 IU), 60 softgels
- 01758 Vitamin D3 with Sea-Iodine™
- 02040 Vitamins D and K with Sea-Iodine™

WEIGHT MANAGEMENT & BODY COMPOSITION

- 02479 7-Keto® DHEA Metabolite 100 mg, 60 veg capsules
- 01807 Advanced Appetite Suppress
- 02207 AMPK Metabolic Activator
- 02504 Body Trim and Appetite Control
- 02478 DHEA Complete
- 01738 Garcinia HCA
- 01908 Mediterranean Trim with Sinetrol[™] -XPur
- 01432 Optimized Saffron
- 00818 Super CLA Blend with Sesame Lignans
- 02509 Waistline Control™

WOMEN'S HEALTH

- 01942 Breast Health Formula
- 01626 Enhanced Sex for Women 50+
- 01894 Estrogen for Women
- 01064 Femmenessence MacaPause®
- 02204 Menopause 731[™]
- 02319 Prenatal Advantage
- 01441 Progesta-Care®
- 01649 Super-Absorbable Soy Isoflavones
- 02507 Youthful Woman 40+ with B-Complex



EXTENSION

Super Selenium Complex 200 mcg & Vitamin E

Providing the Most Advanced Form of Selenium



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

Super Selenium Complex combines <u>three</u> complementary forms of selenium with vitamin E for additional antioxidant protection. Item #01778 • 100 vegetarian capsules 1 bottle \$10.50 • 4 bottles \$9 each

Each bottle provides a supply that lasts more than three months.

For full product description and to order **Super Selenium Complex**, call 1-800-544-4440 or visit www.LifeExtension.com

THREE WAYS TO BUILD STRONGER BONES

CUSTOMIZE YOUR BONE-HEALTH PROGRAM

- 1. Bone Restore helps maintain healthy bone density with three different forms of **calcium** plus vitamin D3 and magnesium, boron, zinc, manganese, and silicon.
- 2. Bone Restore with Vitamin K2, same formula as Bone Restore plus 200 mcg of vitamin K2.

OR

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

LIFE

Bone Restore

Elite

Super Potent K2

Clinical Strength K2 for Bone Density

120 CAPSULES SUPPLEMENT

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Bone Restore Item #01726 • 120 capsules 1 bottle \$16.50 • 4 bottles \$14.25 each

LIFE

Bone

Restore'

Superior Calcium Formula

120 CAPEULES SUPPLEMENT

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Bone Restore with Vitamin K2* Item #01727 • 120 capsules 1 bottle \$18 • 4 bottles \$16.50 each

EXTENSION

Bone

Restore

Vitamin K2

Superior Calcium Formula

120 CAPSULES SUPPLEMENT

NON

Bone Restore Elite with Super Potent K2* Item #02416 • 120 capsules 1 bottle \$34.50 • 4 bottles \$31.50 each

For full product description and to order all **BONE RESTORE** products, call **1-800-544-4440** or visit **www.LifeExtension.com**

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

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These statements have not been evaluated by the Food and Drug Administration. These products are not intended to diagnose, treat, cure, or prevent any disease.



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