# **CORE SUPPORT**





# **CLINICAL APPLICATIONS**

- Enhances Phase II and III Detoxification Mechanisms
- Provides Key Alkalanizing Factors to Support Toxin Excretion
- · Supports Healthy Estrogen Detoxification
- · Promotes Gastrointestinal Health



# GASTROINTESTINAL HEALTH

Core Support is scientifically formulated to provide advanced support for increased Phase II liver detoxification and Phase III toxin elimination. It features clean macronutrients, key phytonutrients, fiber and alkalinizing factors to enhance the detoxification and elimination of xenobiotics and xenoestrogens from the body. Core Support provides 15 g of hypoallergenic brown rice protein and 8 g of fiber with 0 g of added sugar to effectively promote metabolic detoxification.

#### **Overview of Detoxification**

The CDC estimates that more than 100,000 chemicals are used by Americans and about 1,000 new chemicals are introduced annually.¹ Evidence suggests even low-level toxicity is related to a variety of health concerns. In addition, the standard American diet, which is high in sugar and caffeine while low in nutrient density and fiber, has been shown to slow detoxification mechanisms and reduce the elimination of toxins. Therefore, it is important to have an aggressive, daily detoxification strategy that supports the various phases of detoxification.

The liver is the body's main organ of detoxification. The detoxification process, also known as biotransformation, is a three-phase process where environmental pollutants, hormone disruptors, unhealthy estrogen metabolites, xenoestrogens (synthetic compounds that imitate estrogen), and other harmful toxins are safely processed and removed from the body.<sup>2,3</sup> These detoxification systems are very complex and require a variety of nutrients and antioxidant protection for optimal function. In Phase I, a specialized family of enzymes known as the cytochrome P450 enzymes use oxygen to form

a reactive site on toxic compounds. This process creates highly reactive compounds that are more harmful and reactive than the initial toxins. These strong free radicals can cause structural damage to lipids, proteins and DNA within the cell. This is why Phase II detoxification is a crucial step that must be ready to bind, package and neutralize these harmful intermediate metabolites so they can be safely excreted from the body. During the final step of detoxification, Phase III, neutralized, water-soluble compounds can then be excreted in the urine, bile or stool <sup>3,4</sup>

The ingredients included in Core Support were specifically chosen for their ability to support Phase II detoxification and Phase III elimination. N-acetyl cysteine, along with glycine and taurine, are well-known amino acids that enhance liver detoxification. The antioxidants included in Core Support, such as lipoic acid, green tea, ellagic acid and a unique vegetable antioxidant blend, upregulate key Phase II enzymes and protect the liver from oxidative damage. In addition, watercress, pomegranate and potassium citrate provide essential support for the kidneys, facilitating the efficient elimination of toxins through urinary excretion.

# N-Acetyl Cysteine<sup>†</sup>

N-acetyl cysteine (NAC) is a sulfhydryl-containing amino acid that is commonly used to support liver health. Though studies have shown the absorption of oral glutathione to be limited, supplementation with NAC has been shown to significantly increase circulating levels of glutathione, a primary antioxidant that protects cellular health.<sup>5-7</sup> Increasing glutathione levels increases the production of specialized antioxidant enzymes, including glutathione peroxidase, glutathione reductase and



detoxification enzymes, such as glutathione S-transferase. Through the activity of these enzymes, NAC protects the body from oxidative damage, increases Phase II biotransformation and enhances the normal breakdown of toxins and other metabolic by-products of the body.

# **Glycine**<sup>†</sup>

One of the six Phase II biotransformation pathways is amino acid conjugation (the attachment of amino acids to a toxin). Glycine is one of the amino acids used in this process, and it is also necessary for glutathione synthesis.8 Glycine preserves intracellular glutathione (GSH) concentration and protects cells from oxidative damage. This process is mediated by a protein called glycine transporter 1, or GLYT1.9 Research has shown that glycine treatment of human intestinal cells against an oxidative agent reduced the intracellular concentration of reactive oxygen species (ROS) when exposed to oxidative challenge.9

# Taurine<sup>†</sup>

The sulfation pathway is another important Phase II detoxification pathway. In the sulfation pathway, a sulfurcontaining molecule attaches to the toxin to produce a compound that can be excreted from the body. Studies show taurine effectively conjugates bile acids<sup>10</sup> and increases glutathione production.<sup>11</sup>

### Potassium Citrate<sup>†</sup>

Potassium is an essential mineral found in many fiber-rich whole foods and is important for various functions in the body, like electrolyte and acid-base balance. Potassium citrate is a citrate salt known to help regulate urinary pH levels, promote alkalinization in the blood and make the water-soluble toxins processed from biotransformation more easily excretable via the urine by the kidney.<sup>12</sup>

### Pomegranate Fruit Extract<sup>†</sup>

The pomegranate fruit is well known for its high antioxidant levels and ability to protect tissues from free radical damage.<sup>13,14</sup> It has been shown that pomegranates can have up to three times more antioxidants than green tea or red wine. In addition, the specialized blend of flavonoid antioxidants in pomegranate fruit extract has been shown to enhance the activity of key detoxification enzymes while supporting and protecting kidney function to increase the elimination of toxins through urinary excretion.<sup>15-18</sup>

### Watercress Extract<sup>†</sup>

Watercress is a unique cruciferous vegetable that is a member of the cabbage family Brassicaceae. It is a nutrient-dense green plant that has been shown to boost the body's detoxification mechanisms. Studies demonstrate that watercress helps reduce oxidative stress from total toxic load on the liver while

also having a protective effect on kidney function.<sup>19</sup> Watercress also promotes urinary flow to enhance the elimination of toxins via the urine.

# Lipoic Acid†

Lipoic acid is a potent antioxidant that has been shown to increase glutathione, vitamin E and vitamin C levels in the body.<sup>20</sup> Lipoic acid has been shown to support Phase II biotransformation by increasing the activity of enzymes, including NAD(P)H, quinone reductase and glutathione-Stransferase.<sup>21</sup> In a study investigating the molecular mechanisms and therapeutic potential of lipoic acid, it was observed that it significantly increased GSH in various cell types.<sup>22</sup>

### **Green Tea Leaf Extract<sup>†</sup>**

Green tea is one of the most widely consumed beverages throughout the world. One of the main polyphenols in green tea includes epigallocatechin-3-gallate (EGCG). Green tea polyphenols have been shown to increase antioxidant protection. Green tea has also been shown to enhance liver detoxification. Studies demonstrate that green tea extract increases Phase II enzymes, such as glutathione transferase, NAD(P)H, quinone reductase epoxide hydrolase and UDP-glucuronosyltransferase.<sup>23</sup> EGCG potentiates cellular defense capacity against chemical toxins, ultraviolet radiation and oxidative stress.<sup>24</sup>

# Rosemary Leaf Extract<sup>†</sup>

Rosemary includes polyphenols that are potent antioxidants.<sup>25</sup> Carnosol, an antioxidant in rosemary, induces glutathione-stransferase and other important Phase II enzymes.<sup>26</sup> Rosemary essential oil and carnosol have also been shown to increase GSH levels.<sup>27</sup>

# Vegetable Antioxidant Blend<sup>†</sup>

Core Support contains a blend of high-concentration superfood vegetables with significant antioxidant potential. The blend is high in ORAC value (oxygen radical absorbance capacity, a method of measuring antioxidant activity) and includes health-promoting compounds, like sulforaphane and glucosinolates. Cruciferous vegetables, including broccoli, kale and Brussels sprouts, increase the enzyme activity of both Phase I and Phase II biotransformation pathways. Sulforaphane induces Phase II enzymes and supports the body's response to oxidative stress to maintain normal inflammatory balance. Glucosinolates serve as precursors for biologically active metabolites, which induce Phase II enzymes via the activation of Nrf2, the master cellular switch responsible for antioxidant production.

# Schisandra Berry Extract<sup>†</sup>

Schisandra is an adaptogenic botanical used medicinally to help fight off the physical and mental effects of stress. Schisandra is



also used to support liver health and neutralize the effects of toxin exposure. Schisandra enhances liver biotransformation pathways by increasing the levels of reduced glutathione in the liver as well as glutathione reductase and glutathione-S-transferase activity. In animal studies, schisandra has been shown to support Phase I metabolism and protect the liver from free radical damage induced by toxic chemical exposure following ingestion of carbon tetrachloride.<sup>31</sup>

#### Fiber Blend<sup>†</sup>

The average recommendation for fiber intake is 25-30 g per day.<sup>32</sup> Only 5% of the population meets the daily requirement.<sup>33</sup> To help close the fiber intake gap and support the elimination of toxins, Core Support provides a blend of diverse fibers, which includes resistant tapioca dextrin,<sup>34</sup> guar gum, flaxseed flour,<sup>35</sup> glucomannan,<sup>36-39</sup> fig and prune fiber, and arabinogalactan from larch trees. This fiber blend supports a higher level of microbial richness and diversity as well as intestinal barrier function.<sup>40</sup> Optimal microbiome balance plays a crucial role in elimination by influencing the metabolism of both exogenous and endogenous toxins.<sup>41-43</sup> In addition, fiber provides fuel for a healthy and diverse gut microbiome, which supports bile acid production and secretion to aid in the removal of toxins.<sup>44-46</sup>

Fiber has also been shown to have a strong positive impact on both metabolic and gut health markers. One randomized, double-blind, placebo-controlled clinical study showed resistant dextrin increased satiety and reduced feelings of hunger. This information points to the importance of fiber for those who are looking for better elimination, gut health and weight management.<sup>47</sup>

# **Directions**

Mix 2 scoops of Core Support with 8-10 ounces of the beverage of your choice to the desired thickness, 2 times daily or as recommended by your health care professional.

#### **Does Not Contain**

Gluten, yeast, artificial colors or flavors.

#### **Cautions**

Do not consume this product if you are pregnant or nursing. Consult your physician for further information.

Because Glucomannan is a bulk-forming fiber, the drink becomes viscous within 20 minutes of its preparation. Without drinking enough liquid, the product may swell in the throat, causing blockage or choking. Avoid use if you have ever had esophageal narrowing or swallowing difficulties.

# Supplement Facts

Serving Size 2 Scoops (33.8 Grams) Servings Per Container About 14

A	Amount Per Serving	% Daily Value
Calories	120	
Total Fat	1 g	1%*
Total Carbohydrate	12 g	4%*
Dietary Fiber	8 g	29%*
Total Sugars	1 g	**
Protein	15 g	30%*
Calcium	80 mg	6%
Iron	1.5 mg	8%
Magnesium	110 mg	26%
Sodium	35 mg	2%
Potassium	300 mg	6%
Rice Protein	18.9 g	**
Fiber Blend	6.7 g	
Resistant Tapioca Dextrin		**
Flaxseed Flour		**
Guar Gum Fiber (Sunfiber®)		**
Glucomannan ( <i>Amorphophallus konja</i>	c) (Root)	**
Fig (Fruit)	10) (11001)	**
Prune		**
Arabinogalactan Heartwood (from Lar	ch Tree)	**
Glycine USP	750 mg	**
Magnesium Citrate USP	750 mg	**
Potassium Citrate USP		**
	750 mg	
Vegetable Antioxidant Blend	500 mg	**
Broccoli Sprout Concentrate		**
Onion Extract (Bulb)		**
Broccoli (Herb Top)		**
Tomato (Fruit)		**
Carrot (Root)		**
Spinach (Leaf)		
Kale (Leaf)		**
Brussels Sprout (Bud)		**
Taurine USP	250 mg	**
Calcium Citrate USP	230 mg	**
Pomegranate Fruit Extract (Pomanox®)	200 mg	
L-Glutamine USP	150 mg	**
Acetyl L-Carnitine Hydrochloride (MitoCarn		**
N-Acetyl-L-Cysteine USP	125 mg	**
Watercress Extract (Aerial Parts)	100 mg	**
Alpha Lipoic Acid	50 mg	**
Green Tea Leaf Extract (Standardized to contain 45% EGCg (Ep	50 mg	** (/atellen n
		n gallate))
Rosemary Leaf Extract	50 mg	**
Schisandra Berry Extract	50 mg	**
Ellagic Acid	25 mg	**
Glucosinolates	1 mg	• • • • • • • • • • • • • • • • • • • •
* Percent Daily Values are based on a 2,00	00 calorie die	et

Percent Daily Values are based on a 2,000 calorie diet
 Daily Value not established.

Other Ingredients: Natural Flavors, Rebaudioside M and Ascorbyl Palmitate.

ID# 680001 (Core Support Vanilla) NET WT. 1 lb 0.69 oz (16.69 oz) (473.2 g)



# Supplement Facts

Serving Size 2 Scoops (35.3 Grams) Servings Per Container About 14

	Amount Per Serving	% Daily Value
Calories	130	
Total Fat	1.5 g	2%*
Saturated Fat	0.5 g	3%*
Total Carbohydrate	12 g	4%*
Dietary Fiber	8 g	29%*
Total Sugars	1 g	**
Protein	15 g	30%*
Calcium	70 mg	5%
Iron	4 mg	22%
Magnesium	110 mg	26%
Sodium	10 mg	<1%
Potassium	600 mg	13%
Rice Protein	17.4 g	**
Fiber Blend	6.7 g	
Resistant Tapioca Dextrin		**
Flaxseed Flour		**
Guar Gum Fiber (Sunfiber®)		**
Glucomannan (Amorphophallus konja	ac) (Root)	**
Fig (Fruit)		**
Prune		**
Arabinogalactan Heartwood (from La	rch Tree)	**
Glycine USP	750 mg	**
Magnesium Citrate USP	750 mg	**
Potassium Citrate USP	750 mg	**
Vegetable Antioxidant Blend	500 mg	
Broccoli Sprout Concentrate		**
Onion Extract (Bulb)		**
Broccoli (Herb Top)		**
Tomato (Fruit)		**
Carrot (Root)		**
Spinach (Leaf)		**
Kale (Leaf)		**
Brussels Sprout (Bud)		**
Taurine USP	250 mg	**
Calcium Citrate USP	230 mg	**
Pomegranate Fruit Extract (Pomanox®)	200 mg	**
L-Glutamine USP	150 mg	**
Acetyl L-Carnitine Hydrochloride (MitoCarn	n®) 125 mg	**
N-Acetyl-L-Cysteine USP	125 mg	**
Watercress Extract (Aerial Parts)	100 mg	**
Alpha Lipoic Acid	50 mg	**
Green Tea Leaf Extract	50 mg	**
(Standardized to contain 45% EGCg (Ep		n gallate)) **
Rosemary Leaf Extract	50 mg	**
Schisandra Berry Extract	50 mg	**
Ellagic Acid	25 mg	**
Glucosinolates	1 mg	
* Percent Daily Values are based on a 2,0 ** Daily Value not established.	00 calorie die	et

\*\* Daily Value not established.

Other Ingredients: Natural Flavors, Rebaudioside M and

ID# 681001 (Core Support Chocolate) NET WT. 1 lb 1.43 oz (17.43 oz) (494.2 g)

Ascorbyl Palmitate.

#### References

- Agency for Toxic Substances and Disease Registry. Chemicals, Cancer, and You. Division of Health Assessment and Consultation. 2011. Accessed January 9, 2024. https:// stacks.cdc.gov/view/cdc/45777
- 2. Raftogianis R, Creveling C, Weinshilboum R, Weisz J. Estrogen metabolism by conjugation. *J Natl Cancer Inst Monogr.* 2000;(27):113-24. doi: 10.1093/oxfordjournals. jncimonographs.a024234. PMID: 10963623.
- 3. Parkinson A, Ogilvie BW. Chapter 6. Biotransformation of Xenobiotics. In: Klaassen CD, Watkins JB, III. eds. Casarett & Doull's Essentials of Toxicology, 2e. McGraw Hill; 2010. Accessed January 04, 2024. https://accesspharmacy.mhmedical.com/content.aspx?bookid=449&sectionid=39910772
- 4. Almazroo OA, Miah MK, Venkataramanan R. Drug Metabolism in the Liver. *Clin Liver Dis.* 2017 Feb;21(1):1-20. doi: 10.1016/j.cld.2016.08.001. Epub 2016 Oct 15. PMID: 27842765.
- 5. Witschi A, et al. The systemic availability of oral glutathione. *Eur J Clin Pharmacol.* 1992;43:667-9.
- 6. De Rosa SC, et al. N-acetylcysteine replenishes glutathione in HIV infection. *Eur J Clin Invest*. 2000 Oct;30(10):841-2.
- 7. Atkuri KR, Mantovani JJ, Herzenberg LA, et al. N-Acetylcysteine—a safe antidote for cysteine/glutathione deficiency. *Curr Opin Pharmacol.* 2007 Aug;7(4):355-9. Review.
- 8. McCarty MF, O'Keefe JH, DiNicolantonio JJ. Dietary Glycine Is Rate-Limiting for Glutathione Synthesis and May Have Broad Potential for Health Protection. *Ochsner J.* 2018 Spring;18(1):81-87. PMID: 29559876; PMCID: PMC5855430.
- 9. Howard A, Tahir I, et al. Glycine transporter GLYT1 is essential for glycine-mediated protection of human intestinal epithelial cells against oxidative damage. *J Physiol.* 2010; 588(Pt 6):995-1009.
- 10. Birdsall T C. Therapeutic applications of taurine. *Altern Med Rev.* 1998; 3(2):128-136.
- 11. Das J, Ghosh J, et al. Protective role of taurine against arsenic-induced mitochondria-dependent hepatic apoptosis via the inhibition of PKCdelta-JNK pathway. *PLoS One.* 2010; 5(9):e12602.
- 12. Tanner GA. Potassium citrate/citric acid intake improves renal function in rats with polycystic kidney disease. *J Am Soc Nephrol.* 1998 Jul;9(7):1242-8. doi: 10.1681/ASN. V971242. PMID: 9644634.



- 13. Avila-Carrasco L, Garcia-Mayorga EA, Diaz-Avila DL, Garza- Veloz I, Martinez-Fierro ML, Gonzalez-Mateo GT. Potential Therapeutic Effects of Natural Plant Compounds in Kidney Disease. *Molecules*. 2021 Oct 9;26(20):6096. doi: 10.3390/molecules26206096. PMID: 34684678; PMCID: PMC8541433.
- 14. Hodges RE, Minich DM. Modulation of Metabolic Detoxification Pathways Using Foods and Food-Derived Components: A Scientific Review with Clinical Application. *J Nutr Metab.* 2015;2015:760689. doi: 10.1155/2015/760689. Epub 2015 Jun 16. PMID: 26167297; PMCID: PMC4488002.
- 15. Bakır S, Yazgan UC, İbiloğlu İ, Elbey B, Kızıl M, Kelle M. The protective effect of pomegranate extract against cisplatin toxicity in rat liver and kidney tissue. *Arch Physiol Biochem*. 2015;121(4):152-6. doi: 10.3109/13813455.2015.1068336. Epub 2015 Aug 6. PMID: 26247305.
- 16. Soliman NA, Mansour SW, Ammar MA, Hassan NA, Mohamed RHA. Possible role of pomegranate fruit in reversing renal damage in rats exposed to Phenylhydrazine. *Open Vet J.* 2023 Oct;13(10):1268-1276. doi: 10.5455/OVJ.2023. v13.i10.5. Epub 2023 Oct 31. PMID: 38027401; PMCID: PMC10658015.
- 17. Barch DH, Rundhaugen LM, Pillay NS. Ellagic acid induces transcription of the rat glutathione S-transferase-Ya gene. *Carcinogenesis*. 1995 Mar;16(3):665-8. doi: 10.1093/carcin/16.3.665. PMID: 7697830.
- 18. Barch DH, Rundhaugen LM, Stoner GD, Pillay NS, Rosche WA. Structure-function relationships of the dietary anticarcinogen ellagic acid. *Carcinogenesis*. 1996 Feb;17(2):265-9. doi: 10.1093/carcin/17.2.265. PMID: 8625448.
- Panahi Kokhdan E, Khodabandehloo H, Ghahremani H, Doustimotlagh AH. A Narrative Review on Therapeutic Potentials of Watercress in Human Disorders. Evid Based Complement *Alternat Med.* 2021 May 7;2021:5516450. doi: 10.1155/2021/5516450. PMID: 34055006; PMCID: PMC8123986.
- 20. Smith A R, Shenvi S V, et al. Lipoic acid as a potential therapy for chronic diseases associated with oxidative stress. *Curr Med Chem.* 2004; 11(9):1135-1146.
- 21. Flier J, Van Muiswinkel F L, et al. The neuroprotective antioxidant alpha-lipoic acid induces detoxication enzymes in cultured astro glial cells. *Free Radic Res.* 2002; 36(6):695-699.
- 22. Shay KP, Moreau RF, et al. Alpha-lipoic acid as a dietary supplement: molecular mechanisms and therapeutic potential. *Biochem Biophys Acta*. 2009; 1790(10):1149-1160.

- 23. Yu R, Jiao JJ, et al. Activation of mitogen-activated protein kinases by green tea polyphenols: potential signaling pathways in the regulation of antioxidantresponsive element-mediated phase II enzyme gene expression. *Carcinogenesis*. 1997; 18(2):451-456.
- 24. Na HK, Surh YJ. Modulation of Nrf2-mediated antioxidant and detoxifying enzyme induction by the green tea polyphenol EGCG. *Food Chem Toxicol.* 2008; 46(4):1271-1278.
- 25. Offord EA, Mace K, et al. Mechanisms involved in the chemoprotective effects of rosemary extract studied in human liver and bronchial cells. *Cancer Lett.* 1997; 114 (1-2):275-281.
- 26. Singletary KW. Rosemary extract and carnosol stimulate rat liver glutathione-S-transferase and quinone reductase activities. *Cancer Lett.* 1996; 100(1-2):139-144.
- 27. Chen CC, Chen HL, et al. Upregulation of NF-E2-related factor-2-dependent glutathione by carnosol provokes a cytoprotective response and enhances cell survival. *Acta Pharmacol Sin.* 2011; 32(1):62-69.
- 28. Nestle M. Broccoli sprouts as inducers of carcinogendetoxifying enzyme systems: clinical, dietary, and policy implications. *Proc Natl Acad Sci U S A.* 1997; 94(21):11149-11151.
- 29. Kim HJ, Barajas B, et al. Nrf2 activation by sulforaphane restores the age-related decrease of T(H)1 immunity: role of dendritic cells. *J Allergy Clin Immunol*. 2008; 121(5):1255-1261.
- 30. Haack M, Lowinger M, et al. Breakdown products of neoglucobrassicin inhibit activation of Nrf2 target genes mediated by myrosinase-derived glucoraphanin hydrolysis products. *Biol Chem.* 2010; 391(11):1281-1293.
- 31. Zhu M, Yeung RY, Lin KF, Li RC. Improvement of phase I drug metabolism with Schisandra chinensis against CCl4 hepatotoxicity in a rat model. *Planta Med.* 2000 Aug;66(6):521-5.
- 32. Barber TM, Kabisch S, Pfeiffer AFH, Weickert MO. The Health Benefits of Dietary Fibre. *Nutrients*. 2020 Oct 21;12(10):3209. doi: 10.3390/nu12103209. PMID: 33096647; PMCID: PMC7589116.
- 33. Quagliani D, Felt-Gunderson P. Closing America's Fiber Intake Gap: Communication Strategies From a Food and Fiber Summit. *Am J Lifestyle Med.* 2016 Jul 7;11(1):80-85. doi: 10.1177/1559827615588079. PMID: 30202317; PMCID: PMC6124841.



- 34. Wen JJ, Li MZ, Nie SP. Dietary supplementation with resistant starch contributes to intestinal health. *Curr Opin Clin Nutr Metab Care*. 2023;26(4):334-340.
- 35. Hall C 3rd, Tulbek MC, Xu Y. Flaxseed. *Adv Food Nutr Res*. 2006;51:1-97.
- 36. Wu WT, Cheng HC, Chen HL. Ameliorative effects of konjac glucomannan on human faecal β-glucuronidase activity, secondary bile acid levels and faecal water toxicity towards Caco-2 cells. *Br J Nutr.* 2011;105(4):593-600.
- 37. Zalewski BM, Szajewska H. Effect of glucomannan supplementation on body weight in overweight and obese children: protocol of a randomised controlled trial. *BMJ Open.* 2015;5(4):e007244. Published 2015 Apr 13.
- 38. Vuksan V, Sievenpiper JL, Owen R, et al. Beneficial effects of viscous dietary fiber from Konjac-mannan in subjects with the insulin resistance syndrome: results of a controlled metabolic trial. *Diabetes Care*. 2000;23(1):9-14.
- 39. Sood N, Baker WL, Coleman CI. Effect of glucomannan on plasma lipid and glucose concentrations, body weight, and blood pressure: systematic review and meta-analysis. *Am J Clin Nutr.* 2008;88(4):1167-1175.
- 40. Kocot AM, Jarocka-Cyrta E, Drabińska N. Overview of the Importance of Biotics in Gut Barrier Integrity. *Int J Mol Sci.* 2022;23(5):2896. Published 2022 Mar 7.
- Jeong HG, Kang MJ, Kim HG, Oh DG, Kim JS, Lee SK, Jeong TC. Role of intestinal microflora in xenobioticinduced toxicity. *Mol Nutr Food Res.* 2013 Jan;57(1):84-99. doi: 10.1002/ mnfr.201200461. Epub 2012 Nov 20. PMID: 23166009.
- 42. Strand J. Distinctive Detoxification: The Case for Including the Microbiome in Detox Strategy. *Integr Med (Encinitas)*. 2022 Sep;21(4):26-30. PMID: 36644597; PMCID: PMC9542931.
- Zhang YL, Li ZJ, Gou HZ, Song XJ, Zhang L. The gut microbiota-bile acid axis: A potential therapeutic target for liver fibrosis. *Front Cell Infect Microbiol*. 2022 Sep 15;12:945368. doi: 10.3389/fcimb.2022.945368. PMID: 36189347; PMCID: PMC9519863.
- 44. Strand J. Distinctive Detoxification: The Case for Including the Microbiome in Detox Strategy. *Integr Med (Encinitas)*. 2022 Sep;21(4):26-30. PMID: 36644597; PMCID: PMC9542931.

- 45. Zhang YL, Li ZJ, Gou HZ, Song XJ, Zhang L. The gut microbiota-bile acid axis: A potential therapeutic target for liver fibrosis. *Front Cell Infect Microbiol*. 2022 Sep 15;12:945368. doi: 10.3389/fcimb.2022.945368. PMID: 36189347; PMCID: PMC9519863.
- 46. Tilg H, Adolph TE, Trauner M. Gut-liver axis: Pathophysiological concepts and clinical implications. *Cell Metab.* 2022 Nov 1;34(11):1700-1718. doi: 10.1016/j. cmet.2022.09.017. Epub 2022 Oct 7. PMID: 36208625.
- 47. Guérin-Deremaux L, Pochat M, Reifer C, Wils D, Cho S, Miller LE. The soluble fiber NUTRIOSE induces a dose-dependent beneficial impact on satiety over time in humans. *Nutr Res.* 2011 Sep;31(9):665-72. doi: 10.1016/j.nutres.2011.09.004. PMID: 22024490.

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