

*7 Steps to Optimize Your
Microbiome - And Why You
Need To*

**A Budget-Friendly Guide to
Transforming Your Gut and Your
Health**

By

Dr Michael Klein

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Introduction to the Human Microbiome and Health

As strange as this may sound, I've had a fascination with the idea of trying to find the elixir of youth since I was a child. *Could the microbiome be it?* The human microbiome refers to the trillions of microorganisms (bacteria, viruses, fungi, etc.) living in and on our bodies, with the gut microbiome being one of the most densely populated and influential communities¹. Far from passive passengers, these microbes play integral roles in digestion, nutrient synthesis, immune system development, and even modulating our metabolism and nervous system². A well-balanced gut microbiome contributes to homeostasis and health, while disturbances (dysbiosis) have been linked to a host of diseases ranging from gastrointestinal disorders to metabolic and neuropsychiatric conditions³. In recent

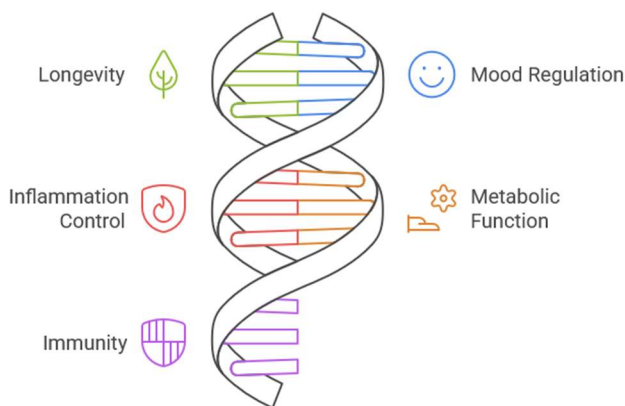
¹ Holscher HD. Dietary fiber and prebiotics and the gastrointestinal microbiota. *Gut Microbes*. 2017 Mar 4;8(2):172-184. doi: 10.1080/19490976.2017.1290756. Epub 2017 Feb 6. PMID: 28165863; PMCID: PMC5390821.

² Wu HJ, Wu E. The role of gut microbiota in immune homeostasis and autoimmunity. *Gut Microbes*. 2012 Jan-Feb;3(1):4-14. doi: 10.4161/gmic.19320. Epub 2012 Jan 1. PMID: 22356853; PMCID: PMC3337124.

³ Du Y, He C, An Y, Huang Y, Zhang H, Fu W, Wang M, Shan Z, Xie J, Yang Y, Zhao B. The Role of Short Chain Fatty Acids in Inflammation and Body Health. *Int J Mol Sci*. 2024 Jul

years, high-quality studies have increasingly shown that the microbiome is a key factor in longevity, mood regulation, inflammation control, metabolic function, and immunity.

The Microbiome's Multifaceted Role in Health and Wellness



This section synthesizes findings from the past decade (2015–2025) on how the microbiome impacts various facets of health and what interventions (dietary, lifestyle, and even herbal) can support a healthy microbiome. I am a **medical doctor** with over **20 years of experience** in pharmaceutical research across three continents—Africa, Asia, and Australia. Over the years, I have served on the African, Asian, and

5;25(13):7379. doi: 10.3390/ijms25137379. PMID: 39000498; PMCID: PMC11242198.

European boards of three multinational pharmaceutical companies and spent much of my career analyzing vast amounts of data to extract **high-value information** from the constant flood of new clinical trials. I have seen firsthand the benefits and limitations of modern medicine. This book is part of a series aimed at the curious as well as at cash- and time-strapped families looking for effective and science-backed solutions for common health problems. My goal is to simplify the overwhelming amount of peer-reviewed scientific information and present it in a way that is both easy-to-read and practical.

Microbiome and Longevity

Emerging research suggests that certain microbiome characteristics are associated with healthy aging and longevity. For example, studies have found that **centenarians (people aged 100+) often harbor a unique and diverse gut microbiota** compared to younger adults⁴. Here are just four ways in which the

⁴ Wang J, Qie J, Zhu D, Zhang X, Zhang Q, Xu Y, Wang Y, Mi K, Pei Y, Liu Y, Ji G, Liu X. The landscape in the gut microbiome of long-lived families reveals new insights on longevity and aging - relevant neural and immune function. *Gut Microbes*. 2022 Jan-Dec;14(1):2107288. doi: 10.1080/19490976.2022.2107288. PMID: 35939616; PMCID: PMC9361766.

gut bugs of Healthy centenarians could be the key to their long lives.

1] The microbiome of those over 100 years of age tend to have a **higher abundance of *beneficial short-chain fatty acid (SCFA)*-producing bacteria**, which **provide energy** to colon cells and have **anti-inflammatory** effects⁵. These SCFAs (like butyrate) may help reduce chronic inflammation and **improve immune function** in the elderly.



2] In one large-scale analysis of over 9,000 individuals, researchers observed that “healthy aging” was marked by an increasingly unique microbiome composition in older adults, particularly a **decline in common core bacteria such as *Bacteroides***⁶. Notably, seniors who

⁵ Wang J, Qie J, Zhu D, Zhang X, Zhang Q, Xu Y, Wang Y, Mi K, Pei Y, Liu Y, Ji G, Liu X. The landscape in the gut microbiome of long-lived families reveals new insights on longevity and aging - relevant neural and immune function. Gut Microbes. 2022 Jan-Dec;14(1):2107288. doi: 10.1080/19490976.2022.2107288. PMID: 35939616; PMCID: PMC9361766.

⁶ Wilmski T, Diener C, Rappaport N, Patwardhan S, Wiedrick J, Lapidus J, Earls JC, Zimmer A, Glusman G, Robinson M, Yurkovich JT, Kado DM, Cauley JA, Zmuda J, Lane NE, Magis AT, Lovejoy JC, Hood L, Gibbons SM, Orwoll ES, Price ND. Gut microbiome pattern reflects healthy ageing and predicts survival in humans. Nat Metab. 2021 Feb;3(2):274-286. doi: 10.1038/s42255-021-00348-0. Epub 2021 Feb 18. Erratum in:

retained a high abundance of *Bacteroides* (i.e. a less unique, more average microbiome) had **lower 4-year survival rates**, suggesting that a shift toward a distinctive, diverse microbiome in late life correlates with longevity. To put it simply, *as people age, their gut bacteria become more unique compared to others*. Those who maintain a more “average” mix of gut microbes—similar to younger adults—tend to have **shorter lifespans**. On the other hand, **seniors with a more diverse and individualized gut microbiome** seem to **live longer**, suggesting that **making your gut bugs more diverse** will likely be one of the keys to living longer.

3] Centenarians also exhibit microbes that *produce specialized metabolites* thought to protect health. A striking finding from Japan showed people over 100 had higher levels of gut bacteria **that generate secondary bile acids**, compounds known to fend off pathogens and regulate immunity⁷. One such bile acid

Nat Metab. 2021 Apr;3(4):586. doi: 10.1038/s42255-021-00377-9. PMID: 33619379; PMCID: PMC8169080.

⁷ Sato Y, Atarashi K, Plichta DR, Arai Y, Sasajima S, Kearney SM, Suda W, Takeshita K, Sasaki T, Okamoto S, Skelly AN, Okamura Y, Vlamakis H, Li Y, Tanoue T, Takei H, Nittono H, Narushima S, Irie J, Itoh H, Moriya K, Sugiura Y, Suematsu M, Moritoki N, Shibata S, Littman DR, Fischbach MA, Uwamino Y, Inoue T, Honda A, Hattori M, Murai T, Xavier RJ, Hirose N, Honda K. Novel bile acid biosynthetic pathways are enriched in the microbiome of centenarians. Nature. 2021

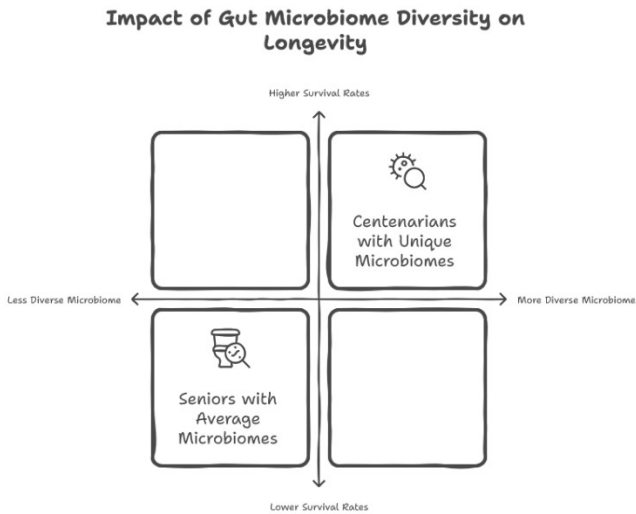
molecule, isoallolithocholic acid, was isolated from centenarian gut bacteria and found to strongly inhibit *Clostridioides difficile*, a dangerous antibiotic-resistant bacterium that causes severe gut infections. In experiments, this centenarian-derived molecule could kill or suppress multiple harmful microbes, indicating that the gut microbiome of the extremely aged can produce chemicals that help **keep infections at bay and maintain microbial balance**⁸. As a result, researchers believe a unique microbiome might be one reason centenarians are relatively resilient to infections and age-related illnesses.

4] There is also evidence that certain gut microbes in centenarians may counteract the **chronic low-grade inflammation often seen in aging** (“inflammaging”). For instance, *Bacteroides fragilis* was enriched in one centenarian study and is thought to promote longevity

Nov;599(7885):458-464. doi: 10.1038/s41586-021-03832-5. Epub 2021 Jul 29. PMID: 34325466.

⁸ McMillan AS, Theriot CM. Bile acids impact the microbiota, host, and *C. difficile* dynamics providing insight into mechanisms of efficacy of FMTs and microbiota-focused therapeutics. Gut Microbes. 2024 Jan-Dec;16(1):2393766. doi: 10.1080/19490976.2024.2393766. Epub 2024 Sep 3. Erratum in: Gut Microbes. 2024 Jan-Dec;16(1):2411134. doi: 10.1080/19490976.2024.2411134. PMID: 39224076; PMCID: PMC11376424.

by inducing the anti-inflammatory cytokine IL-10, helping to balance immune responses⁹.



Taken together, these findings suggest that nurturing a diverse, robust gut microbiome—rich in beneficial, anti-inflammatory and pathogen-fighting organisms—

⁹ Wang J, Qie J, Zhu D, Zhang X, Zhang Q, Xu Y, Wang Y, Mi K, Pei Y, Liu Y, Ji G, Liu X. The landscape in the gut microbiome of long-lived families reveals new insights on longevity and aging - relevant neural and immune function. *Gut Microbes*. 2022 Jan-Dec;14(1):2107288. doi: 10.1080/19490976.2022.2107288. PMID: 35939616; PMCID: PMC9361766.

could be a key to healthier aging and increased longevity.

Not only observational links, but **causal evidence from animal studies**



supports the microbiome's role in aging: transferring gut microbes from young mice into older mice has

been shown to *delay age-related decline* in *physical* and *cognitive* functions¹⁰. While more research (especially in humans) is needed, the current data point to the microbiome as an important determinant of lifespan and a potential target for promoting healthy aging¹¹.

¹⁰ Zhang N, Zhang Y, Wang Z, Pan F, Ren R, Li Z, Zhao H, Luo X, Li Z, Wang L, Mo R, Sun G, Peng L, Ni M, Yang Y. Regular fecal microbiota transplantation to Senescence Accelerated Mouse-Prone 8 (SAMP8) mice delayed the aging of locomotor and exploration ability by rejuvenating the gut microbiota. *Front Aging Neurosci.* 2022 Oct 3;14:991157. doi: 10.3389/fnagi.2022.991157. PMID: 36262889; PMCID: PMC9574184.

¹¹ Wilmanski T, Diener C, Rappaport N, Patwardhan S, Wiedrick J, Lapidus J, Earls JC, Zimmer A, Glusman G, Robinson M, Yurkovich JT, Kado DM, Cauley JA, Zmuda J, Lane NE, Magis AT, Lovejoy JC, Hood L, Gibbons SM, Orwoll ES, Price ND. Gut microbiome pattern reflects healthy ageing and predicts survival in humans. *Nat Metab.* 2021 Feb;3(2):274-286. doi: 10.1038/s42255-021-00348-0. Epub 2021 Feb 18. Erratum in: *Nat Metab.* 2021 Apr;3(4):586. doi:

Gut Bacteria, Mood Regulation, and the Brain

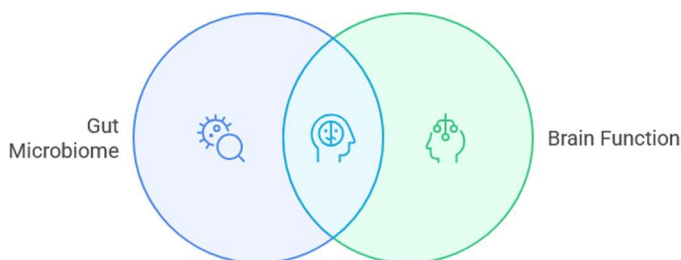
The gut-brain axis is a bidirectional communication network through which the gut microbiome can influence brain chemistry, mood, and behavior. A large body of evidence now indicates that gut bacteria play a surprising role in **neurological and psychological health**. One major mechanism is through **neurotransmitters**: it's estimated that about *90% of the body's serotonin* – a neurotransmitter crucial for mood regulation – is produced in the **gastrointestinal tract** with the help of gut microbes¹². Certain intestinal bacteria stimulate the gut's enterochromaffin cells to synthesize serotonin; in a

10.1038/s42255-021-00377-9. PMID: 33619379; PMCID: PMC8169080.

¹² Fung TC, Vuong HE, Luna CDG, Pronovost GN, Aleksandrova AA, Riley NG, Vavilina A, McGinn J, Rendon T, Forrest LR, Hsiao EY. Intestinal serotonin and fluoxetine exposure modulate bacterial colonization in the gut. *Nat Microbiol*. 2019 Dec;4(12):2064-2073. doi: 10.1038/s41564-019-0540-4. Epub 2019 Sep 2. PMID: 31477894; PMCID: PMC6879823.

landmark mice study, eliminating specific bacteria led to **more than a 50% drop in gut serotonin levels**, which normalized when those microbes were reintroduced¹³. This demonstrates how profoundly

Interplay of Gut and Brain



microbes can affect serotonin production, which in turn may influence **mood, appetite, and gastrointestinal motility**.

Beyond serotonin, gut microbes also play a role in producing and modifying other brain chemicals (neurotransmitters) that affect mental and emotional well-being. For instance, certain strains of *Lactobacillus* and *Bifidobacterium*—which are commonly found in probiotic foods like yogurt—can produce **GABA** (gamma-aminobutyric acid). GABA is an inhibitory neurotransmitter, meaning **it helps calm the nervous system, reduce anxiety, and promote relaxation**.

¹³ Katherine Z. Sanidad et al. , Gut bacteria-derived serotonin promotes immune tolerance in early life. Sci. Immunol.9,eadj4775(2024).DOI:10.1126/sciimmunol.adj4775

Additionally, other gut bacteria can influence the metabolism of *dopamine* and *noradrenaline*, two key neurotransmitters involved in motivation, focus, and the body's response to stress. **Dopamine** is often called the "pleasure chemical" because it plays a central role in feelings of reward and motivation, while **noradrenaline** (also called norepinephrine) helps regulate alertness and the fight-or-flight response.

The gut microbiome communicates with the central nervous system (the brain and spinal cord) using two main pathways¹⁴:

- 1) **Neural pathway** (via the vagus nerve): The vagus nerve acts as a direct communication highway between the gut and brain, carrying signals that can influence mood, stress responses, and even behavior.
- 2) **Endocrine pathway** (hormonal signaling): Gut bacteria help regulate hormones and neurotransmitters, affecting brain function and emotional balance.

In simple terms, this means that your gut bacteria don't just help with digestion—*they actively shape your brain chemistry and mental health*. A balanced

¹⁴ Appleton J. The Gut-Brain Axis: Influence of Microbiota on Mood and Mental Health. *Integr Med* (Encinitas). 2018 Aug;17(4):28-32. PMID: 31043907; PMCID: PMC6469458.

microbiome could promote better mood, reduced stress, and overall cognitive well-being, while an imbalanced microbiome may contribute to anxiety, depression, and other mental health issues¹⁵.

Research in germ-free animals (animals raised in completely sterile conditions without any gut bacteria) illustrates the importance of communication between the gut and the brain. Studies on germ-free mice (mice raised without any microbiome) have shown that **these animals experience abnormal brain development, including heightened stress responses** (meaning they react more strongly to stress compared to normal mice). Additionally, they exhibit lower levels of **brain-derived neurotrophic factor (BDNF)** in key brain regions like the hippocampus, which is critical for *memory formation, learning, and emotional regulation*.

Gut bacteria play a crucial role in regulating **stress and anxiety**. They help "fine-tune" the hypothalamic-pituitary-adrenal (HPA) axis, the body's central stress-response system, during early life¹⁶. The



¹⁵ Appleton J. The Gut-Brain Axis: Influence of Microbiota on Mood and Mental Health. *Integr Med (Encinitas)*. 2018 Aug;17(4):28-32. PMID: 31043907; PMCID: PMC6469458.

¹⁶ Appleton J. The Gut-Brain Axis: Influence of Microbiota on Mood and Mental Health. *Integr Med (Encinitas)*. 2018 Aug;17(4):28-32. PMID: 31043907; PMCID: PMC6469458.

HPA axis controls how the body reacts to stress by managing hormones like cortisol (the stress hormone). If this system is poorly regulated, a person may be more prone to anxiety and stress-related disorders later in life.

Interestingly, disrupting the gut microbiome later in life—such as by taking antibiotics (which kill both harmful and beneficial bacteria)—can also cause *temporary changes in behavior and brain function*. Studies show that such disruptions can lead to increased anxiety-like behavior and changes in BDNF levels, reinforcing the idea that the gut microbiome continues to influence the brain even in adulthood. In humans, numerous studies have linked gut microbiome composition to mental health conditions such as anxiety and depression. A 2019 population study published in *Nature Microbiology* found that **people with depression consistently lacked two key genera of bacteria, *Coprococcus* and *Dialister***, even after accounting for antidepressant use¹⁷. Intriguingly, *Coprococcus* was also correlated with higher quality of life indicators, suggesting these bacteria may have

¹⁷ Valles-Colomer M, Falony G, Darzi Y, Tigchelaar EF, Wang J, Tito RY, Schiweck C, Kurilshikov A, Joossens M, Wijmenga C, Claes S, Van Oudenhove L, Zhernakova A, Vieira-Silva S, Raes J. The neuroactive potential of the human gut microbiota in quality of life and depression. *Nat Microbiol*. 2019 Apr;4(4):623-632. doi: 10.1038/s41564-018-0337-x. Epub 2019 Feb 4. PMID: 30718848.

neuroactive properties beneficial to mental well-being¹⁸. The absence of certain butyrate-producing bugs in depression could contribute to neuroinflammation or altered neurotransmitter levels. On the flip side, some gut bacteria are associated with positive mood and might produce metabolites that have antidepressant or anxiolytic effects. These findings have given rise to the concept of “psychobiotics” – probiotics or interventions that can improve mental health by modulating the gut microbiome¹⁹. Early clinical trials are testing whether consuming specific probiotic strains can alleviate depression or anxiety. For instance, one trial showed a multi-strain probiotic reduced rumination and aggressive thoughts in people with moderate depression²⁰, though more research is needed for definitive recommendations.

¹⁸ Valles-Colomer M, Falony G, Darzi Y, Tigchelaar EF, Wang J, Tito RY, Schiweck C, Kurilshikov A, Joossens M, Wijmenga C, Claes S, Van Oudenhove L, Zhernakova A, Vieira-Silva S, Raes J. The neuroactive potential of the human gut microbiota in quality of life and depression. *Nat Microbiol*. 2019 Apr;4(4):623-632. doi: 10.1038/s41564-018-0337-x. Epub 2019 Feb 4. PMID: 30718848.

¹⁹ Xiong RG, Li J, Cheng J, Zhou DD, Wu SX, Huang SY, Saimaiti A, Yang ZJ, Gan RY, Li HB. The Role of Gut Microbiota in Anxiety, Depression, and Other Mental Disorders as Well as the Protective Effects of Dietary Components. *Nutrients*. 2023 Jul 23;15(14):3258. doi: 10.3390/nu15143258. PMID: 37513676; PMCID: PMC10384867.

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Gut microbiome links have also emerged in neurological disorders. In Parkinson's disease (PD), for example, patients often exhibit an altered gut microbiome, and constipation can precede motor symptoms by years. Fascinating experiments in 2016 provided causal evidence: mice engineered to overproduce human α -synuclein (a model of Parkinson's) **did not develop the usual motor dysfunction unless they had a gut microbiome present**²¹. Germ-free PD-model mice had much milder symptoms, but when colonized with gut bacteria from Parkinson's patients, they developed worse motor deficits and brain inflammation than mice colonized with healthy human microbiota²². This indicates that microbial metabolites and products can trigger or exacerbate PD pathology in the host.

²¹ Sampson TR, Debelius JW, Thron T, Janssen S, Shastri GG, Ilhan ZE, Challis C, Schretter CE, Rocha S, Gradinaru V, Chesselet MF, Keshavarzian A, Shannon KM, Krajmalnik-Brown R, Wittung-Stafshede P, Knight R, Mazmanian SK. Gut Microbiota Regulate Motor Deficits and Neuroinflammation in a Model of Parkinson's Disease. *Cell*. 2016 Dec 1;167(6):1469-1480.e12. doi: 10.1016/j.cell.2016.11.018. PMID: 27912057; PMCID: PMC5718049.

²² Sampson TR, Debelius JW, Thron T, Janssen S, Shastri GG, Ilhan ZE, Challis C, Schretter CE, Rocha S, Gradinaru V, Chesselet MF, Keshavarzian A, Shannon KM, Krajmalnik-Brown R, Wittung-Stafshede P, Knight R, Mazmanian SK. Gut Microbiota Regulate Motor Deficits and Neuroinflammation in a Model of Parkinson's Disease. *Cell*. 2016 Dec 1;167(6):1469-1480.e12. doi: 10.1016/j.cell.2016.11.018. PMID: 27912057; PMCID: PMC5718049.

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