

The Gut Microbiome: A Comprehensive Review of its Role in Human Health and Disease

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Abstract

Background: The gut microbiome is an extensive ecosystem of microorganisms in the human gastrointestinal tract that is critical to our health which influences physiological functions that range from metabolic processes to immune processes. This systematic review is intended to provide an overview of the gut microbiome and the members that comprise it, but will particularly overview the three dominant bacterial phyla, Firmicutes, Bacteroidetes, and Actinobacteria, whose biological functions vary in health. The gut microbiome also includes various aspects of the external environment such as nutrition, age, and geography, that can influence composition and diversity in microbial communities. The gut microbiome can be disrupted causing dysbiosis and this condition has been linked to numerous diseases which continues to reinforce our understanding of the importance of maintaining a healthy and balanced gut microbiome. Nutrition, in particular, is one of the most significant environmental influences in mediating the gut microbiome, and researchers have begun recent studies to understand the role that diet has in our gut health, specifically dietary fiber, phytochemicals, and polyphenols. Certain foods including fermented foods and whole grains are thought to positively alter microbial balance and these examples highlight that nutrition can impact microbiome health. It does not seem possible to overshadow the significance and complexity of the link between gut microbiome and human health. This manuscript reviews the importance of the microbiome in metabolic disorders, such as obesity, inflammatory bowel disease (IBD), and mental health through the gut-brain axis. The manuscript also discusses the microbiome implications in cancer and autoimmune diseases. The therapeutic potential is reviewed beyond these associations in developing fecal microbiota transplantation (FMT) strategies, psychobiotics, and prebiotics to discuss mental health. Exciting advances in novel probiotic and prebiotics formulations are also highlighted to improve health. The manuscript concludes with a summary of the main points, a list of future studies on the therapeutic potential of the microbiome, and mentions the importance of the microbiome to human health and clinical applications.

Materials and Methods: The current review article has been prepared using systematic review methodology in using reviewed published research articles, reviews articles, and scientific reports about the gut microbiome and its impact on health. Articles were obtained from accessing peer-reviewed literatures from relevant databases: PubMed, Science Direct, Google Scholar, and Wiley Online Library about the gut microbiome through the following terms (gut microbiome, dysbiosis, Firmicutes, Bacteroidetes, Actinobacteria, nutrition, pre and probiotics and gut-brain axis).

Conclusion: The gut microbiome is important for metabolism, immunity, and neurological function for human health. Dysbiosis has been associated with obesity, inflammatory bowel disease (IBD), and mental health issues. The number of studies examining potential targeted therapeutics in dysbiosis is increasing, examples include probiotics, prebiotics and fecal microbiota transplant (FMT), to help restore dysbiosis to health. Future studies will need to improve knowledge and understanding the molecular and cellular interactions of the host-microbiome, as well as support the field moving towards intentionally targeted gut health for improvement in health.

Keywords: Gut microbiome, Microbial diversity, Dysbiosis, Firmicutes, Bacteroidetes, Diet and microbiome, Prebiotics and probiotics, Gut-brain axis, Metabolic disorders, Gastrointestinal health.

1. INTRODUCTION

The human gastrointestinal tract occupies a large and complex community of microorganisms referred to as the gut microbiome including trillions of microorganisms, such as bacteria, viruses, fungi, and other microbes that are a part of the human body systems. The human gut contains hundreds of species of microorganisms in what seems to be a simple system, in which the overwhelming majority of microorganisms in the human microbiome are good types and vital to human existence. For instance, gut bacteria are essential in the digestion and fermentation of complex carbohydrates into short-chain fatty acids, and synthesizing vitamins that the human hosts need from bacteria, because i.e., the human body does not synthesize them¹. The microbiome composition of a given person can be influenced by factors, such as, but not limited to, genetics, diet, lifestyle, and environment. Therefore, an individual's gut microbiome can change in terms of the composition and the environment, which relate to health gut dysbiosis to give an example. The microbiome has been noted to be one of the determinants of overall health. The directionality of the gut microbiome process is not limited to the biological system of digestion, as it has been established that there can be impacts on immune health, inflammation, and brain function through the gut-brain axis². The microbiome can serve as a protective barrier by creating competition for resources amongst microorganisms and creating an environment that inhibits harmful bacteria from growing. In addition, studies have documented the influence of gut microbiota on your risk of development of certain chronic diseases. Gut bacteria have been linked to several acute and chronic illnesses such as obesity, diabetes, and inflammatory bowel disease. More recently, there is accumulating evidence that your gut microbiota may have an influence on mental health, with more links to mood and stress-related disorders. This indicates that maintaining a healthy microbiome will not only lead to improvement in digestion, but will also optimize healthy living, and well-being³.

2. COMPOSITION AND DIVERSITY OF THE GUT MICROBIOME

The gut microbiome is composed of a vast number of microorganisms, primarily bacteria that co-exist with the human host in a state of symbiosis. The majority of gut microbiota are classified into four main bacterial phyla: Firmicutes, Bacteroidetes, Actinobacteria and Proteobacteria. Firmicutes is the largest of the human gut microbiome phyla, containing diverse species important for breaking down dietary fiber and producing short-chain fatty acids (SCFAs) that are important for promoting gut health⁴. Genera such as *Clostridium*, *Lactobacillus*, and *Faecalibacterium* are notable members of this group. These bacteria play a significant role in maintaining the intestinal barrier and modulating the immune response. Another major phylum is Bacteroidetes, which includes the genus *Bacteroides*. Members of this phylum are particularly effective at breaking down complex carbohydrates, playing a critical role in energy metabolism⁵. Bacteroidetes and Firmicutes together make up the bulk of the gut microbiota, with their relative abundance often linked to overall gut health and metabolic function. A balanced ratio of these two phyla is associated with better metabolic outcomes, while an imbalance may be related to conditions such as obesity or inflammatory bowel disease.

Actinobacteria, though present in smaller amounts, also play an essential role in gut health. The genus *Bifidobacterium* is a well-known member of this phylum and is often found in probiotic supplements due to its beneficial effects on gut health, particularly in early childhood development and in maintaining a healthy immune response. The phylum *Proteobacteria*, while less dominant, contains both beneficial and pathogenic species. An overgrowth of *Proteobacteria* is often considered a marker for gut dysbiosis, which may contribute to inflammation and disease. Members of this phylum, such as *Escherichia coli*, are common in the gut, but an increase in pathogenic strains can lead to infections or chronic inflammation⁶.

In addition to the predominant bacterial phyla, the gut microbiome also consists of lower abundances of archaea, fungi, and viruses. The archaea of interest, *Methanobrevibacter*, are a group of archaea that contribute to digestive processes by eliminating gas through methanogenesis. The composition of the gut microbiome will change from person to person based upon several factors, including a person's diet, genetics, environment, and lifestyle, but its essential roles are maintained. A multiplicity of the microbiome allows for improved absorption of nutrients, a balanced immune system and resistance to disease, while a reduced diversity is associated with specific diseases including obesity, diabetes and gastrointestinal diseases. Therefore, diversity and the composition of the microbiome is one of the most significant markers of overall gut health⁷.

2.1. Role of Firmicutes, Bacteroidetes, and Actinobacteria

The gut microbiome is primarily composed of several important bacterial phyla, among them Firmicutes, Bacteroidetes, and Actinobacteria that are critical for supporting the gut's health and functioning. These phyla enhance nutrient metabolism, immune function, and pathogen protection, each providing specific benefits to the host.

2.2. Firmicutes

Firmicutes represent a prolific phylum of microorganisms identified within the human gastrointestinal tract, and are primarily involved in carbohydrate catabolism. Organisms within this phylum comprise, *Lactobacillus*, *Clostridium* and *Faecalibacterium*, exhibit an impressive capability to ferment complex carbohydrates/ dietary fibers into bioactive compounds or short-chain fatty acids (SCFAs) including; butyrate, acetate and propionate. SCFAs exhibit myriad beneficial roles, including; acting as the principal energy source of colonic epithelial cells (colonocytes), to promoting cellular ion transport and improving gut barrier integrity, as well as dampening negative inflammation signals to promote lower levels of inflammation through modulation of the immune system. Firmicutes are also known for their role in stabilising a healthy microbial community, whereby; *Faecalibacterium prausnitzii* is identified within this group and is known for its anti-inflammation properties, thus a diminished abundance of this species is often seen in diseases such as inflammatory bowel disease (IBD). The ratio of Firmicutes-to-Bacteroidetes is also commonly studied in the context of metabolic health, with one possible conclusion being that a higher ratio is associated with obesity, however, more research is needed⁸.

2.3. Bacteroidetes

Bacteroidetes is an additional predominant phylum in the gastrointestinal (GI) tract; it is critical in decomposing intricate polysaccharide structures, such as resistant starches and various plant-based fibers. Its phylogenetic composition includes species like *Bacteroides* and *Prevotella*, which are incredibly specialized in fermenting carbohydrates that humans are not equipped to digest. This fermentation process is important because it helps to utilize the maximum energy from food, and it produces SCFAs that contribute to gut health via the nourishment of colon cell structures and immune regulation. Significantly, Bacteroidetes have been shown to assist in regulating fat storage and metabolism. Several studies have suggested that an abundance of Bacteroidetes (relative to Firmicutes) appears to correlate with leanness while a lesser amount seems to correlate with obesity. Members of Bacteroidetes also detoxify harmful chemicals and synthesize nutrients like certain vitamins, emphasizing their importance in gut and overall health⁹.

2.4. Actinobacteria

Actinobacteria, while present in smaller quantities compared to Firmicutes and Bacteroidetes, still play a significant role in the gut ecosystem. The most well-known genus within this phylum is *Bifidobacterium*, which is often found in high concentrations in infants and plays a critical role in shaping the early immune system. These bacteria are essential for breaking down oligosaccharides, particularly in breast milk, which aids in the healthy development of the infant gut microbiome. In adults, *Bifidobacterium* species continue to contribute to gut health by supporting the digestion of complex carbohydrates, producing vitamins like vitamin B12, and inhibiting the growth of harmful bacteria through competitive exclusion and the production of antimicrobial compounds. Their presence is often associated with beneficial effects on the immune system, helping to regulate inflammation and prevent conditions like irritable bowel syndrome (IBS) and allergies. Actinobacteria are also frequently included in probiotic formulations due to their role in promoting a healthy gut environment¹⁰.

3. FACTORS INFLUENCING GUT MICROBIOME DIVERSITY

The diversity of the gut microbiome is responsive to a myriad of influences, providing a basis for its composition and function. Three of the strongest influences are age, diet, and geography. Each of these variables plays an important role in determining the richness and diversity of microbes in the gastrointestinal tract, ultimately contributing to health and disease. Age is one of the most important determinants of gut microbiome diversity and several early life influences influence its establishment and longer-term stability. Microbiome development starts at birth due in large part to maternal influences; the baby's first microbial colonization derived, largely, from the mother's vaginal, gut, and skin microbiota during delivery, as well as through breastfeeding. In the case of maternal death or

maternal illness, the infant does not receive a critical source of microbial transfer and immune protection, and microbiome development can be delayed or altered. The microbiome alterations are often characterized by lower numbers of beneficial taxa (e.g., *Bifidobacterium* and *Lactobacillus*) and more opportunistic pathogens¹¹.

In an analogous circumstance, infants with inadequate nutrition or formula feeding experience a lack of the normal microbiota maturation and diversity, which may reflect fewer numbers of *Bifidobacteria* and butyrate-producing bacteria; these factors may be later risk factors for metabolic and immune disorders even later in life. As the child continues to grow, the microbiota develops new microbes and the diversity increases, which is dependent on diet, environment, and health status, until it reaches an adult-like microbiota, generally by age 3. The adult microbiota state is generally stable and diverse, however as adults age, their microbiota is less diverse and has decreased resilience, shows increased inflammation and has lower resistance to infection¹².

Aging, dietary, medication or antibiotic use, and immunosuppression contribute to altered changes in microbial diversity. Puberty is a critical developmental window during which the gut microbiome composition can change drastically, due to revolutions in hormones, the diet, and metabolism. The significant increase in hormones of sex, such as estrogen and testosterone, parallel changes in microbial communities that are distinct across sexes, including differences in taxa diversity and degrees of abundance during puberty. Microbial composition is also recently suggested to relate to timing of pubertal onset, as dysbiosis is suggested to be related to early or precocious puberty with diminished short chain fatty acids. In the wake of COVID-19 pandemic, there has been an increase in reports of precocious puberty, possibly associated with diet, stress, and lifestyle, changes that impact gut microbiota balance, however there are still speculative intergenerational changes. Overall, puberty is a critical transition period where evolution of the gut microbiome and host physiology are co-driving the alterations in metabolism and reproductive physiology, which may have implications on long-term metabolic and reproductive health¹³.

Alterations related to advanced age with regard to dietary consumption, pharmacologic utilization (including antibiotic administration), and a declining immune system all influence microbial diversity in the GI, as presented in chapters about age in this review. Puberty is an important developmental window that begins a period of gut microbiome transition due to influences from hormonal shifts (e.g., estrogen or testosterone changes), dietary impacts, and metabolic changes. The rise in sex hormones, such as estrogen and testosterone, promotes sex-specific microbiome profiles, with evidence demonstrating differences in taxa and microbial diversity by sex during adolescence. Recent research has also linked gut microbiota to the initiation of puberty and that microbiome dysbiosis and lower profolocal, short-chain fatty acids (SCFA), were associated with early or precocious puberty onset. The pandemic that began in 2020 and subsequent lockdowns has manifested in increased reports of precocious puberty. This phenomenon may be associated with impacts on diet, stress, and changes in lifestyle, which are all factors driven by environmental exposures. It is unclear to what degree intergenerational exposure has or demonstrates gut microbial impact. Hence, in summary, puberty represents a period of significant transition whereby microbial dysbiosis or alterations in microbial diversity influences changes in host physiology, which impacts long-term consequences on metabolic health and reproductive status¹⁴.

Diet is arguably the most direct and modifiable factor affecting the diversity and composition of the gut microbiome. Different dietary patterns can significantly alter the balance of microbial species in the gut. A diet rich in plant-based foods, such as fruits, vegetables, whole grains, and legumes, tends to promote greater microbial diversity. These foods provide complex carbohydrates and fibers that serve as prebiotics, feeding beneficial bacteria like *Bifidobacterium* and *Lactobacillus*. The fermentation of these fibers by gut bacteria results in the production of short-chain fatty acids (SCFAs), which support gut health and reduce inflammation. In contrast, a diet high in processed foods, sugars, and unhealthy fats can reduce gut microbiome diversity. Diets low in fiber may lead to a decline in beneficial bacteria and an overgrowth of opportunistic or pathogenic microbes. Long-term consumption of such diets has been linked to conditions like obesity, diabetes, and gastrointestinal disorders. Moreover, the inclusion of fermented foods such as yogurt, kefir, kimchi, and sauerkraut in the diet introduces probiotics, which can enhance microbial diversity by directly adding beneficial bacteria to the gut. On the other hand, excessive alcohol consumption and certain medications, like antibiotics, can drastically reduce diversity by killing off large portions of the microbiome¹⁵.

Dietary components such as fiber, phytochemicals, and polyphenols play a fundamental role in determining the composition and health of the gut microbiome. These substances not only promote the growth of beneficial bacteria but also contribute to vital metabolic processes that enhance gut function and overall well-being. Dietary fiber, an essential element of plant-based foods including fruits, vegetables, whole grains, and legumes, acts as a substrate for microbial fermentation, resulting in the production of short-chain fatty acids (SCFAs) such as butyrate, propionate, and acetate¹⁶. These SCFAs are crucial for maintaining gut integrity, providing energy to colon cells, and reducing inflammation. Fiber can be classified into soluble and insoluble types soluble fiber found in oats, legumes, and fruits supports the growth of beneficial bacteria like *Bifidobacterium* and *Lactobacillus*, while insoluble fiber from whole grains and vegetables promotes bowel regularity and increases stool bulk¹⁷. Similarly, phytochemicals, which are bioactive compounds found in plant foods such as flavonoids, carotenoids, and glucosinolates, play multifaceted roles in gut health by stimulating beneficial bacterial growth, inhibiting pathogenic species, and improving nutrient absorption. Diets rich in phytochemicals, particularly those resembling the Mediterranean diet, are associated with higher microbial diversity and reduced risk of chronic conditions including obesity, diabetes, and cancer¹⁸. Polyphenols, a major group of phytochemicals abundant in foods like tea, coffee, berries, and red wine, further influence gut microbiota composition by acting as prebiotics, enhancing populations of beneficial bacteria, and generating anti-inflammatory metabolites during fermentation. Collectively, these dietary components fiber, phytochemicals, and polyphenols work synergistically to enhance microbial diversity, maintain gut homeostasis, and promote long-term health and wellness¹⁹.

Geography and ethnicity play a significant role in shaping the gut microbiome, reflecting variations in lifestyle, environmental exposures, and dietary habits across regions and populations. Individuals living in rural, non-industrialized societies generally possess a more diverse gut microbiome compared to those in urban or industrialized settings, largely due to higher intake of fiber-rich, plant-based diets and greater exposure to environmental microbes. In contrast, urbanized populations consuming processed foods with lower fiber intake often show reduced microbial diversity, influenced further by antibiotic use and decreased contact with diverse environmental microorganisms²⁰. Moreover, comparative studies among ethnic groups have revealed distinct microbial compositions that may explain variations in metabolic health outcomes. For instance, East Asians (EAs) experience poorer metabolic health at lower body mass indices compared to other ethnic groups, and recent multi-omic studies have demonstrated marked differences in gut microbiota between East Asian and White individuals living in the same region. White participants were enriched with *Akkermansia muciniphila*, a mucin-degrading bacterium linked to metabolic benefits, whereas East Asian subjects showed higher abundance of multiple fermentative bacterial phyla and elevated short-chain fatty acid end-products such as acetate, propionate, and isobutyrate. These differences persisted even after accounting for dietary intake and were more pronounced in lean individuals, suggesting that host ethnicity and geography exert a strong influence on gut microbiota composition. Collectively, these findings highlight that geographical location, lifestyle, and ethnicity interact to shape the gut microbiome, potentially contributing to population-specific metabolic and health disparities²¹.

Dysbiosis is a term that describes an imbalance in the gut microbiome, characterized by a decrease in microbial diversity and a shift in the composition of microbial communities. This condition can arise from a variety of factors, including poor dietary choices, chronic stress, medication use (particularly antibiotics), and lifestyle changes. In a healthy gut microbiome, a diverse range of beneficial bacteria coexists, playing crucial roles in digestion, immune function, and overall health. However, dysbiosis often results in a decline of these beneficial microorganisms, leading to an overgrowth of pathogenic or opportunistic bacteria, which can have profound implications for health. The consequences of dysbiosis are particularly evident in gastrointestinal disorders. Conditions such as irritable bowel syndrome (IBS) and inflammatory bowel disease (IBD) have been associated with a disrupted microbiome. In these cases, individuals may experience symptoms like abdominal pain, bloating, and irregular bowel habits. The imbalance can lead to increased intestinal permeability, commonly referred to as “leaky gut” which allows harmful substances to enter the bloodstream, triggering inflammation and exacerbating symptoms²².

Dysbiosis also plays a significant role in metabolic disorders. Research has shown that imbalances in gut bacteria can influence weight gain and insulin sensitivity. For example, certain bacterial species associated with obesity can enhance energy extraction from dietary sources, leading to increased fat

storage. Conversely, a diverse gut microbiome is often linked to better metabolic health, aiding in glucose regulation and fat metabolism. This relationship suggests that dysbiosis may contribute to the rising prevalence of obesity and type 2 diabetes, conditions that have become increasingly common in modern societies. Furthermore, the gut microbiome is intimately connected to the immune system. A healthy microbiome helps train and regulate immune responses, but dysbiosis can lead to immune dysfunction. This may result in an overactive immune response, contributing to the development of autoimmune diseases, such as rheumatoid arthritis and multiple sclerosis. Individuals with dysbiosis are also more susceptible to infections due to a compromised gut barrier function, highlighting the protective role of a balanced microbiome²³.

Dysbiosis, characterized by an imbalance in the gut microbiome, is often associated with chronic low-grade inflammation, a common feature of obesity. Certain gut bacteria can produce pro-inflammatory compounds, such as lipopolysaccharides (LPS), which enter the bloodstream and provoke systemic inflammation. This inflammatory response can interfere with the body's ability to regulate glucose and fat metabolism, increasing the risk of developing type 2 diabetes, cardiovascular diseases, and other obesity-related conditions. Furthermore, the presence of inflammatory cytokines in the bloodstream can lead to insulin resistance, compounding metabolic dysfunction and promoting weight gain. The gut microbiome also plays a role in the secretion of hormones involved in appetite regulation. Beneficial gut bacteria can stimulate the production of hormones like GLP-1 (glucagon-like peptide-1) and PYY (peptide YY), which are involved in promoting satiety and reducing food intake. GLP-1 is released in response to food intake and helps to enhance insulin secretion while inhibiting glucagon release, which can lower blood sugar levels. PYY is released after eating and signals fullness to the brain, reducing appetite²⁴. Conversely, an imbalance in gut bacteria may lead to lower levels of these satiety hormones, resulting in increased hunger and overeating, further contributing to obesity. The gut microbiome communicates with the central nervous system via the gut-brain axis, influencing behaviours related to food intake and energy expenditure. Certain gut bacteria can produce neurotransmitters, such as serotonin and GABA (gamma-aminobutyric acid), which play roles in mood and appetite regulation. For instance, approximately 90% of the body's serotonin is produced in the gut, and an optimal gut microbiome can support balanced levels of this neurotransmitter. Disruptions in the gut microbiome may lead to altered neurotransmitter levels, potentially resulting in changes in mood and eating behaviours. This interplay between the gut microbiome and brain underscores the significance of gut health in mental well-being and eating patterns²⁵.

The relationship between the gut microbiome and mental health has garnered significant attention in recent years, emphasizing the importance of the gut-brain axis, a bidirectional communication network linking the gut and the central nervous system. This connection highlights how gut bacteria can influence mood, cognition, and behavior through the production of neurotransmitters and signaling molecules such as serotonin, dopamine, and gamma-aminobutyric acid (GABA). Dysbiosis, or imbalance in gut microbial composition, has been associated with a range of neuropsychiatric and immunological disorders, including anxiety, depression, autism spectrum disorder, and even neurodegenerative diseases. Notably, dysbiosis during early life has been linked to an increased risk of developing allergies and asthma, indicating that early microbial disturbances can have long-term effects on immune and neural health. Recent therapeutic strategies for depression and other mood disorders increasingly focus on restoring the gut-brain axis through the use of probiotics, prebiotics, dietary modulation, and even fecal microbiota transplantation (FMT), collectively referred to as "psychobiotics." This emerging field of research offers promising potential for microbiome-based interventions that target both mental and physical well-being²⁶.

A diverse gut microbiome is essential for proper immune development, and disruptions in microbial exposure can hinder this process, leading to heightened sensitivities and inflammatory responses. This emphasizes the importance of microbial diversity in early life for shaping a resilient immune system. Addressing dysbiosis typically involves a multifaceted approach that includes dietary modifications, such as increasing fiber intake and reducing processed foods, alongside the incorporation of probiotics and prebiotics to support beneficial bacteria. Lifestyle changes, such as managing stress and promoting regular physical activity, are also critical in restoring microbial balance. Overall, understanding dysbiosis and its implications highlights the importance of maintaining a healthy gut microbiome for promoting optimal health and preventing disease²⁷.

Prebiotics and probiotics are fundamental dietary components that significantly impact gut health by influencing the gut microbiome. Prebiotics are non-digestible food elements, primarily soluble fibers, that specifically stimulate the growth and activity of beneficial bacteria in the gastrointestinal tract. Unlike probiotics, which are live microorganisms, prebiotics serve as food for these microorganisms, encouraging their proliferation and activity. This selective stimulation helps to cultivate a balanced and diverse gut microbiome, which is essential for optimal digestive health and overall well-being. Common sources include garlic, onions, leeks, asparagus, bananas, and whole grains. These foods contain oligosaccharides that resist digestion in the upper gastrointestinal tract, allowing them to reach the colon, where fermentation occurs. This fermentation results in the production of short-chain fatty acids (SCFAs), such as butyrate, acetate, and propionate, which provide numerous health benefits. SCFAs serve as an energy source for colonocytes, enhance gut barrier function, and exhibit anti-inflammatory properties. Additionally, a diet rich in prebiotics can improve metabolic health by enhancing insulin sensitivity, regulating appetite, and lowering the risk of obesity and type 2 diabetes. By promoting beneficial bacteria such as *Bifidobacterium* and *Lactobacillus*, prebiotics help counteract dysbiosis, a condition characterized by an imbalance of gut microorganisms. Maintaining this balance is crucial for a healthy immune system and may also play a role in preventing various gastrointestinal disorders, allergies, and metabolic diseases²⁸.

Probiotics are live microorganisms that, when consumed in adequate amounts, confer health benefits to the host. They primarily consist of specific strains of bacteria and yeasts known for their positive effects on gut health. The most common probiotic strains include *Lactobacillus*, *Bifidobacterium* and *Saccharomyces boulardii*. Probiotics are found in various fermented foods and supplements, including yogurt, kefir, sauerkraut, kimchi, and dietary supplements. Probiotics contribute to gut health in several ways. They help increase the diversity of the gut microbiome, which is associated with better health outcomes, and inhibit harmful bacteria by competing for nutrients and producing antimicrobial substances, reducing the risk of infections and gastrointestinal disorders. Regular intake of probiotics is associated with improved digestion, enhanced immune function, and reduced symptoms of gastrointestinal disorders such as irritable bowel syndrome (IBS) and inflammatory bowel disease (IBD). Additionally, probiotics may positively influence mental health through the gut-brain axis, suggesting that the gut microbiome's state can affect mood and cognitive function²⁹.

In contrast, the Western diet, characterized by high levels of processed foods, added sugars, unhealthy fats, and low fiber intake, negatively impacts the gut microbiome. This diet often lacks essential components like whole grains, fruits, and vegetables, leading to a reduction in microbial diversity. Low-fiber diets provide fewer substrates for beneficial gut bacteria, resulting in a decline in their populations and an increase in potentially harmful species. This shift can contribute to dysbiosis, which is linked to various health problems, including obesity, metabolic syndrome, inflammatory bowel diseases, and autoimmune disorders. Moreover, the high intake of sugars and unhealthy fats associated with the Western diet can promote inflammation and disrupt gut barrier function, leading to increased intestinal permeability, often referred to as "leaky gut." This condition allows toxins and undigested food particles to enter the bloodstream, triggering systemic inflammation and contributing to chronic diseases³⁰.

The Western diet is also linked to increased production of harmful metabolites, such as trimethylamine N-oxide (TMAO), associated with cardiovascular disease. Furthermore, the imbalance created by this dietary pattern may impair the gut/brain axis, negatively influencing mood and cognitive function. Overall, prebiotics and probiotics are vital for promoting gut health and maintaining a balanced microbiome, while the Western diet can have adverse effects, leading to dysbiosis and various health problems. Adopting a more balanced diet rich in fiber, whole foods, and fermented products can help mitigate these negative effects and support a healthier gut microbiome. By prioritizing dietary choices that nourish beneficial gut bacteria, individuals can significantly enhance their overall health and well-being³¹.

4. GUT MICROBIOME AND HUMAN HEALTH

The gut microbiome plays a significant role in the development and management of obesity and metabolic disorders, influencing various physiological processes that affect body weight and metabolic health. Research has demonstrated that the composition and diversity of gut microorganisms can significantly impact energy balance, fat storage, and overall metabolic function.

4.1. Relationship Between Gut Microbiome and Obesity

Studies have found that individuals with obesity often exhibit distinct gut microbiome compositions compared to those of normal weight. A key observation is the increased ratio of Firmicutes to Bacteroidetes in obese individuals. Firmicutes, a phylum of bacteria, are more efficient at extracting energy from the diet, leading to increased caloric absorption and potential weight gain. This heightened efficiency may result from the ability of Firmicutes to ferment dietary fibers more effectively than Bacteroidetes. In contrast, individuals with a healthier weight and more diverse microbiomes tend to have a higher proportion of Bacteroidetes, which may contribute to lower energy extraction and reduced body weight. The balance of these two phyla highlights the importance of gut microbiome composition in energy metabolism³². The gut microbiome influences metabolic functions through the fermentation of dietary fibers, resulting in the production of short-chain fatty acids (SCFAs). SCFAs, including acetate, propionate, and butyrate, are critical to energy metabolism and appetite regulation. For example, butyrate serves as an energy source for colonocytes (the cells lining the colon) and has been shown to enhance insulin sensitivity, thereby reducing the risk of developing insulin resistance, which is a precursor to type 2 diabetes. In contrast, an imbalance in gut bacteria can lead to decreased SCFA production, contributing to insulin resistance and increased fat accumulation. Research has shown that individuals with obesity often exhibit lower SCFA levels, highlighting the need for a healthy gut microbiome to support optimal metabolic health³³.

Understanding the connection between the gut microbiome and obesity presents potential therapeutic avenues for managing weight and metabolic health. Strategies to modulate the gut microbiome include dietary interventions that increase fiber intake, along with the consumption of prebiotics and probiotics. Prebiotics serve as food for beneficial gut bacteria, while probiotics introduce live beneficial bacteria into the gut. Both strategies can promote a more balanced and diverse microbiome, which is essential for optimal metabolic function. Additionally, targeted microbiome therapies, such as faecal microbiota transplantation, have shown promise in clinical settings for restoring microbial balance and improving metabolic outcomes in obese individuals. Lifestyle changes that emphasize a diet rich in fiber, whole foods, and fermented products can also play a crucial role in promoting a healthy gut microbiome and supporting weight management³⁴. The multifaceted relationship between the gut microbiome and obesity underscores the importance of understanding microbial health in the context of metabolic disorders. Ongoing research continues to unravel the complex interactions within the gut microbiome and their implications for obesity, offering valuable insights into new approaches for combating obesity and promoting metabolic health.

4.2. Inflammatory bowel disease (IBD) and Dysbiosis

Inflammatory bowel disease (IBD), which includes conditions such as Crohn's disease and ulcerative colitis, is marked by chronic inflammation of the gastrointestinal tract. The etiology of IBD is multifactorial, involving genetic predispositions, environmental triggers, and immune system dysregulation. A critical aspect of IBD is the role of the gut microbiome and its relationship with dysbiosis, an imbalance in the composition and function of gut microorganisms. In individuals with IBD, significant changes occur in the composition of the gut microbiota. Studies indicate that these patients often exhibit reduced microbial diversity compared to healthy individuals. This dysbiotic state is characterized by a decrease in beneficial bacteria, such as *Faecalibacterium prausnitzii* and specific strains of *Bifidobacterium* and *Lactobacillus*. These beneficial microbes are essential for maintaining gut health and modulating the immune response. Conversely, IBD patients frequently have an increased abundance of potentially harmful bacteria, such as certain *Escherichia coli* strains and other Enterobacteriaceae. This imbalance can worsen the inflammatory processes associated with IBD, as the pathogenic bacteria may release toxins or other harmful metabolites that further disrupt gut health³⁵.

The gut microbiome plays a vital role in shaping the immune system. A healthy microbiome fosters an environment of tolerance to non-pathogenic antigens and maintains intestinal homeostasis. Dysbiosis disrupts this balance, leading to inappropriate immune activation. In IBD, the presence of pathogenic bacteria can trigger heightened immune responses, resulting in excessive inflammation. This chronic inflammatory state can lead to tissue damage and contribute to the symptoms of IBD, such as abdominal pain, diarrhoea, and rectal bleeding. The immune response may also produce inflammatory cytokines and other mediators that perpetuate the cycle of inflammation and damage to the intestinal lining. The intestinal epithelium acts as a critical barrier that prevents the entry of harmful pathogens and substances

into the bloodstream. A healthy gut microbiome contributes to the maintenance of this barrier. However, dysbiosis can compromise epithelial integrity. Certain harmful bacteria produce factors that can damage the epithelial cells, resulting in increased intestinal permeability, a condition often referred to as “leaky gut”. This permeability allows bacteria and their by-products to enter the bloodstream, triggering systemic inflammatory responses that further exacerbate IBD. The disruption of the epithelial barrier is a significant contributor to the pathogenesis of IBD, as it promotes ongoing inflammation and the potential for infections³⁶.

The gut microbiome is also crucial for metabolizing dietary components, particularly fibers, into short-chain fatty acids (SCFAs) such as butyrate, acetate, and propionate. SCFAs play important roles in maintaining gut health; they serve as an energy source for colonocytes (the cells lining the colon) and help regulate immune responses. In the context of IBD, dysbiosis often results in decreased SCFA production. The reduction in SCFAs can compromise the health of the intestinal lining and exacerbate inflammation. Butyrate, in particular, is known for its anti-inflammatory properties and its ability to enhance epithelial barrier function. A deficiency in SCFAs can lead to worsened symptoms and an increased risk of flare-ups in patients with IBD³⁷.

The strong association between dysbiosis and IBD highlights the potential for targeting the gut microbiome as a therapeutic strategy. Interventions such as probiotics and prebiotics have garnered attention for their potential to restore microbial balance and alleviate symptoms of IBD. Probiotics, which are live microorganisms, can help replenish beneficial bacteria and inhibit the growth of pathogenic species. Certain probiotic strains have shown promise in clinical studies, demonstrating the ability to reduce inflammation and improve quality of life for IBD patients. Prebiotics, non-digestible fibers that nourish beneficial gut bacteria, can also support a healthier microbiome by promoting the growth of favourable microbial populations. Additionally, dietary interventions that emphasize high-fiber foods and fermented products can enhance microbial diversity and support gut health. Thus, dysbiosis plays a pivotal role in the pathophysiology of inflammatory bowel disease. The altered composition and functionality of the gut microbiome can significantly impact immune responses, epithelial barrier integrity, and metabolite production, all of which are critical factors in the onset and progression of IBD. Understanding these connections can inform the development of targeted therapies aimed at restoring microbial balance and improving patient outcomes in IBD³⁸.

4.3. Mental health and the gut-brain axis

The gut-brain axis represents a complex bidirectional communication network linking the gastrointestinal system to the central nervous system, underscoring the significant influence of the gut microbiome on mental health. At the core of this interaction is the gut microbiome, which consists of trillions of microorganisms that produce various metabolites, including neurotransmitters, short-chain fatty acids (SCFAs), and other bioactive compounds. These metabolites play critical roles in brain function; for instance, the majority of the body’s serotonin, a key neurotransmitter involved in mood regulation, is synthesized in the gut³⁹. This highlights the profound connection between gut health and mental well-being, with alterations in microbial composition potentially affecting neurotransmitter levels and thereby influencing mood and behavior. Moreover, the gut microbiome is integral to modulating the immune system. Dysbiosis, characterized by an imbalance in gut bacteria, can lead to increased intestinal permeability, often referred to as “leaky gut”.

This condition allows pro-inflammatory substances to enter the bloodstream, triggering systemic inflammation that can adversely impact brain function. Such inflammation is recognized as a contributing factor in various mental health disorders, including depression and anxiety. A balanced microbiome is thus essential for both maintaining gut health and promoting mental well-being, as it plays a pivotal role in regulating inflammatory responses. The gut microbiome also exerts influence over the hypothalamic-pituitary-adrenal (HPA) axis, a central component of the body’s stress response system. Dysregulation of the HPA axis has been associated with stress-related disorders, including anxiety and depression. The gut microbiota can modulate stress responses through its influence on hormone release, such as cortisol, which is vital for mood regulation. Research suggests that specific microbial compositions can lead to altered stress responses, demonstrating the interconnectedness of gut health and mental health⁴⁰.

Furthermore, emerging studies indicate that the gut microbiome may affect neuroplasticity, the brain’s capacity to adapt and reorganize. Beneficial gut bacteria produce SCFAs that confer neuroprotective

effects, supporting the growth of neurons and enhancing synaptic plasticity, essential for learning, memory, and emotional regulation. A healthy microbiome may bolster cognitive functions and contribute to resilience against stress and anxiety, emphasizing the importance of gut health in mental health outcomes. Clinical implications of these findings are profound, as they suggest that interventions targeting the gut microbiome, such as probiotics and dietary modifications, could alleviate symptoms of anxiety and depression. Research has identified specific probiotic strains associated with reduced anxiety and improved mood in clinical trials, highlighting the potential of microbiome-targeted therapies. Diets rich in fiber and fermented foods can support a healthy microbiome, enhancing mental health outcomes by fostering microbial diversity and resilience⁴¹.

The concept of “psychobiotics” has emerged as a key area of interest within this framework, referring to probiotics that may improve mental health by modulating the gut-brain axis. Research is ongoing to identify specific strains and their effects on various mental health disorders, providing hope for innovative therapeutic approaches that leverage the gut-brain connection to enhance mental health and well-being. Understanding these intricate relationships opens new avenues for interventions aimed at improving mental health through the promotion of a balanced and healthy gut microbiome⁴².

4.4. Other diseases (Cancer, Autoimmune disorders)

The gut microbiome plays a pivotal role in the pathogenesis of various diseases, including cancer and autoimmune disorders, through its intricate interactions with the host’s immune system and metabolic processes.

4.5. Gut microbiome and Cancer

The connection between the gut microbiome and cancer is an area of significant research interest, revealing that microbial composition can impact cancer development, progression, and response to therapy. The gut microbiome consists of trillions of microorganisms, including bacteria, viruses, fungi, and archaea, each contributing to a unique ecosystem that can influence health. Certain bacteria, such as *Lactobacillus* and *Bifidobacterium*, are known for their potential protective effects against cancer. They can enhance the immune response by stimulating the production of immune cells and promoting the release of anti-inflammatory cytokines. Additionally, these beneficial bacteria produce short-chain fatty acids (SCFAs) during the fermentation of dietary fibers, which have been shown to exert anti-cancer effects. Butyrate, a prominent SCFA, can inhibit cancer cell proliferation and promote apoptosis in colon cancer cells. It also plays a role in regulating gene expression related to cell growth and differentiation, further underscoring its importance in cancer prevention⁴³. On the other hand, dysbiosis is a state of microbial imbalance and contributes to cancer development. For example, an increase in pathogenic bacteria, such as *Fusobacterium nucleatum*, has been associated with colorectal cancer. This bacterium can promote tumor growth by enhancing inflammation and modulating the immune response in a way that favours tumor progression. In addition to direct effects on tumor cells, dysbiosis can lead to increased intestinal permeability, allowing toxins and microbial products to translocate into the bloodstream.

This condition, often referred to as “leaky gut,” may trigger systemic inflammation, which is recognized as a key factor in cancer development across various tissues. The gut microbiome also influences the efficacy of cancer treatments, particularly immunotherapy. Research indicates that a diverse microbiome is associated with better responses to immune checkpoint inhibitors, therapies designed to enhance the body’s immune response against tumors. For example, certain bacterial species, such as *Akkermansia muciniphila*, have been linked to improved treatment outcomes in patients receiving these therapies. These findings suggest that modifying the gut microbiome through dietary interventions, probiotics, or other means could enhance the effectiveness of cancer treatments and help mitigate adverse effects⁴⁴.

4.6. Gut Microbiome and Autoimmune Disorders

Autoimmune disorders arise when the immune system mistakenly attacks the body’s own cells, leading to chronic inflammation and tissue damage. The gut microbiome is intricately involved in immune regulation and tolerance, with research showing that its composition can significantly influence the development and progression of these diseases. A healthy microbiome promotes the differentiation of regulatory T cells (Tregs), which are essential for maintaining immune homeostasis and preventing autoimmunity⁴⁵. In contrast, dysbiosis can lead to an imbalance in immune cell populations and an

increased risk of autoimmune reactions. In conditions like Rheumatoid Arthritis (RA), specific alterations in gut microbiota composition have been identified. Studies have reported a reduction in microbial diversity and an increase in pro-inflammatory bacteria in individuals with RA. For instance, bacteria from the *Prevotella* genus have been associated with RA, suggesting that these organisms may contribute to inflammation and joint damage. Similarly, Multiple Sclerosis (MS) has been linked to specific microbial profiles, with some studies indicating that certain gut bacteria may influence the severity of neuroinflammation and demyelination associated with the disease. The mechanisms by which the gut microbiome contributes to autoimmune disorders are complex and multifaceted⁴⁶. Dysbiosis can lead to the production of autoantibodies, antibodies that mistakenly target the body's own tissues thereby exacerbating autoimmune processes. Additionally, changes in gut permeability associated with dysbiosis can allow bacterial antigens to enter the bloodstream, further provoking an immune response. In some cases, molecular mimicry, where microbial components resemble host tissues, may trigger autoimmune responses, leading to conditions such as lupus or type 1 diabetes. Emerging research is exploring the potential for microbiome-targeted therapies as adjunctive treatments for autoimmune disorders. Probiotics, prebiotics, and dietary interventions are being investigated for their ability to restore microbial balance, enhance immune regulation, and alleviate symptoms. For instance, specific probiotic strains have shown promise in modulating the immune response in individuals with inflammatory bowel disease (IBD), a condition characterized by chronic inflammation of the gastrointestinal tract that shares similarities with autoimmune disorders⁴⁷. The gut-brain axis also plays a role in autoimmune diseases, as psychological stress can influence gut health and exacerbate immune dysregulation. Stress can alter gut microbiota composition, potentially leading to increased inflammation and worsening of autoimmune symptoms. Understanding these intricate relationships provides insights into how the gut microbiome can affect not only physical health but also the mental well-being of individuals with autoimmune conditions⁴⁸.

5. THERAPEUTIC POTENTIAL OF THE GUT MICROBIOME

5.1. Fecal microbiota transplantation (FMT)

Fecal microbiota transplantation (FMT) is a therapeutic procedure aimed at restoring the balance of the gut microbiome by transferring fecal material from a healthy donor to a recipient. This approach has gained attention for its effectiveness in treating various gastrointestinal disorders, particularly those associated with dysbiosis, including recurrent *Clostridium difficile* infection (CDI), inflammatory bowel disease (IBD), and other conditions related to metabolic and immune functions. The fundamental concept of FMT is based on the understanding that a healthy gut microbiome is vital for maintaining gut health and overall well-being. Dysbiosis, characterized by an imbalance in microbial diversity with an overgrowth of harmful bacteria, can lead to significant health issues. FMT seeks to rectify this imbalance by introducing a healthy microbiome from a donor. The fecal matter from the donor contains a diverse array of beneficial microorganisms that can restore the microbiome's ecological balance. The FMT procedure begins with the careful selection of a donor, who undergoes extensive screening to ensure they are free from infectious diseases and gastrointestinal disorders. The donor's fecal matter is then processed to eliminate potential pathogens while retaining beneficial microbes. The prepared stool is introduced into the recipient's gastrointestinal tract using various methods, including colonoscopy, enema, or encapsulated oral delivery. Once in the gut, the healthy microorganisms begin to proliferate, outcompete pathogenic bacteria, and help restore the natural microbial community⁴⁹.

FMT is most widely recognized for its efficacy in treating recurrent CDI, a condition often resistant to standard antibiotic therapies. Clinical studies have shown that FMT can achieve cure rates of over 80-90% in patients experiencing recurrent infections. By reintroducing a healthy microbiome, FMT helps inhibit the growth of *C. difficile*, restoring normal gut function and microbial diversity. In addition to CDI, FMT is being explored as a treatment for other gastrointestinal disorders, particularly IBD, which includes conditions such as ulcerative colitis and Crohn's disease. Preliminary research suggests that FMT may help induce remission in some IBD patients by promoting a more diverse microbial population and reducing inflammation within the gut. However, more extensive studies are needed to establish its long-term safety and effectiveness in these conditions. Beyond gastrointestinal issues, researchers are investigating FMT's potential applications in other health areas. Studies have begun to examine the role of gut microbiota in metabolic conditions like obesity and type 2 diabetes, as well as neurological disorders such as autism spectrum disorders and depression. The concept of the gut-brain axis, the bidirectional communication pathway between the gut and the brain, highlights the possibility

that gut microbes can influence mental health and metabolic processes. Some research has suggested that alterations in the gut microbiome may impact mood and behavior, opening the door for FMT as a potential treatment modality in these areas⁵⁰.

Despite its promising potential, FMT raises important safety concerns, particularly regarding the transmission of infectious agents through fecal material. To minimize these risks, thorough donor screening is essential. The screening process evaluates the donor's medical history, dietary habits, and lifestyle choices to ensure they are healthy and free from communicable diseases. The processing of fecal samples is also designed to eliminate pathogens while preserving the beneficial microbial population. There have been rare but serious adverse effects reported following FMT, such as gastrointestinal discomfort and, in some instances, severe infections. Continuous monitoring and research are crucial for evaluating the long-term safety and efficacy of FMT across diverse patient populations. The regulatory status of FMT varies by region and country. In some areas, FMT is classified as an investigational therapy and is subject to stringent regulations. In others, particularly for the treatment of recurrent CDI, it is offered as a standard treatment option through specialized medical centres. As scientific understanding of the gut microbiome evolves, regulatory agencies are continually assessing guidelines for FMT practices to ensure patient safety while enabling access to this innovative therapeutic approach. FMT represents a significant advancement in the treatment of conditions linked to gut dysbiosis, illustrating the potential of the microbiome in therapeutic applications. By effectively restoring a balanced microbial community, FMT has the capacity to improve health outcomes across a variety of disorders⁵¹.

5.2. Psychobiotics and prebiotics in mental health

Psychobiotics and prebiotics represent two significant avenues in the exploration of how gut health influences mental well-being. Psychobiotics are a specific subset of probiotics that are believed to exert positive effects on mood, anxiety, and cognitive function by acting through the gut-brain axis. This axis is a complex communication network linking the gastrointestinal tract and the brain, suggesting that the gut microbiome plays a crucial role in neurological processes. Psychobiotics, particularly certain strains of *Lactobacillus* and *Bifidobacterium*, have been identified for their potential to alleviate symptoms associated with anxiety and depression⁵². These beneficial bacteria can produce various neuroactive compounds, including Gammaaminobutyric acid (GABA), serotonin, and other neurotransmitters that influence mood and behavior. In addition to neurotransmitter production, psychobiotics are known to generate Short-chain fatty acids (SCFAs) during the fermentation of dietary fibers. SCFAs possess anti-inflammatory properties and contribute to overall brain health. Emerging research suggests that these psychobiotics can provide a viable therapeutic strategy for managing mood disorders, with clinical trials showing promising results in reducing anxiety and depressive symptoms among participants who incorporated specific probiotic strains into their diets⁵³.

Prebiotics, in contrast, are non-digestible food components, primarily fibers, that serve as substrates for beneficial gut bacteria. They nourish and promote the growth of these microorganisms, which can enhance the overall diversity and functionality of the gut microbiome. Foods rich in prebiotics include garlic, onions, asparagus, bananas, and whole grains. When consumed, prebiotics resist digestion in the upper gastrointestinal tract and reach the colon, where they undergo fermentation by gut bacteria. This fermentation process leads to the production of SCFAs, which play an important role in maintaining gut health and have been linked to various metabolic and neurological benefits. The relationship between prebiotic and mental health lies in their capacity to support beneficial gut bacteria that produce neurotransmitters and other neuroactive substances. A diet high in prebiotic fibers may positively influence mood and cognitive function, potentially alleviating symptoms of anxiety and depression⁵⁴.

Recent studies have indicated that prebiotic supplementation can enhance the effects of psychobiotics, creating a synergistic relationship that benefits mental health. For example, prebiotics can help improve the survival and efficacy of probiotics in the gastrointestinal tract, thus amplifying their positive effects on mood regulation. Both psychobiotics and prebiotics can influence mental health through various mechanisms. They can modify the gut microbiome's composition and diversity, encouraging the growth of beneficial bacteria that produce essential neurotransmitters. They also contribute to the production of SCFAs, which play significant roles in brain health. Moreover, by strengthening the gut barrier and reducing systemic inflammation, they may prevent or alleviate neuroinflammation, often linked to mood disorders. As research continues to advance in this field, the roles of psychobiotics and prebiotics

are increasingly recognized as critical components of mental health strategies. The gut-brain connection highlights the importance of nutrition in regulating mood and cognitive function, emphasizing the potential of dietary interventions in mental health management⁵⁵.

5.3. Novel probiotics and prebiotics

The landscape of gut health is rapidly evolving with the introduction of novel probiotics and prebiotics, which are redefining our understanding of how dietary components can influence the gut microbiome. As research delves deeper into the intricacies of the human microbiome, these innovative substances are emerging as key players in promoting health and preventing disease. Novel probiotics are distinguished by their unique strains of live microorganisms that confer health benefits beyond those of traditional probiotics. While established probiotics such as *Lactobacillus* and *Bifidobacterium* have been extensively studied, recent discoveries are bringing attention to lesser-known strains that may offer enhanced therapeutic effects. For example, strains like *Lactobacillus rhamnosus* GR1 and *Lactobacillus casei* Shirota have demonstrated specific health benefits, including the modulation of immune responses and the reduction of gastrointestinal disorders⁵⁶.

The sourcing of these novel probiotics is diverse, with strains isolated from fermented foods, human feces, and environmental samples such as soil or plants. The identification of these strains often involves advanced microbiological techniques, including high-throughput sequencing, which allows researchers to characterize and catalog microbial communities in unprecedented detail. A significant aspect of novel probiotics is their ability to withstand gastrointestinal challenges, such as acidity and bile salts, which can hinder the survival of traditional strains. Strains like *Bifidobacterium infantis* have shown enhanced resilience, enabling them to colonize the gut effectively and exert beneficial effects. The functional characteristics of novel probiotics can also be enhanced through genetic engineering, allowing for the development of designer probiotics that can produce specific metabolites or bioactive compounds. For instance, engineered strains that produce butyrate, an SCFA known for its anti-inflammatory properties, can directly impact gut health and metabolic processes.

The ability of these probiotics to produce antimicrobial peptides and other compounds that inhibit pathogenic bacteria further underscores their therapeutic potential⁵⁷. Novel prebiotics have expanded beyond traditional soluble fibers like inulin and fructooligosaccharides to include a variety of compounds that promote gut health. Emerging research has identified new sources of prebiotics, including resistant starches, oligosaccharides from seaweed, and polyphenolic compounds from plant sources. These substances can selectively stimulate the growth of beneficial bacteria while inhibiting harmful microorganisms⁵⁸.

One area of interest is resistant starch, which is found in foods like green bananas, cooked and cooled potatoes, and legumes. This type of starch resists digestion in the small intestine and reaches the colon, where it undergoes fermentation by gut bacteria. The fermentation process produces SCFAs, including butyrate, which is essential for maintaining colonic health, regulating inflammation, and supporting the gut barrier function. Butyrate is particularly important as it serves as the primary energy source for colonocytes, promoting the integrity of the intestinal lining and preventing conditions like leaky gut. Polyphenols, which are abundant in a variety of fruits, vegetables, tea, and dark chocolate, are another exciting category of novel prebiotics. These compounds exhibit antioxidant properties and have been shown to influence gut microbiota composition positively. For instance, polyphenols from blueberries have been linked to increased levels of *Bifidobacterium* and *Lactobacillus*, thereby enhancing gut health and metabolic functions⁵⁹.

5.4. Synbiotics

The concept of synbiotics combinations of prebiotics and probiotics offers an innovative approach to maximizing gut health. By pairing beneficial live bacteria with nondigestible food components that nourish these bacteria, synbiotics can enhance the overall effectiveness of both components. The prebiotic component provides a food source for probiotics, enabling them to thrive and exert their health benefits. For example, a synbiotic formulation containing *Lactobacillus plantarum* alongside inulin can enhance the survival of the probiotic during digestion while simultaneously promoting the growth of beneficial gut bacteria. This synergistic effect can lead to improved gut microbiome diversity, enhanced immune responses, and better overall gut health. Ongoing research is focusing on identifying the optimal ratios and combinations of specific prebiotics and probiotics to maximize these benefits⁶⁰.

5.5. Implications for Health and Disease Management

The implications of novel probiotics and prebiotics extend to various health conditions, including gastrointestinal disorders, metabolic diseases, and mental health issues. Specific strains of novel probiotics may play a role in managing conditions such as Irritable bowel syndrome (IBS), Inflammatory bowel disease (IBD), and even obesity. Research indicates that particular strains can modulate inflammation, restore microbial balance, and enhance gut barrier function, all of which are crucial for alleviating symptoms of these disorders. Moreover, there is growing evidence supporting the connection between gut health and mental well-being, often referred to as the gut-brain axis. Novel probiotics have shown promise in influencing mood and cognitive function, potentially providing therapeutic benefits for anxiety and depression⁶¹. By restoring the balance of gut microbiota and promoting the production of neurotransmitters such as serotonin, these probiotics may serve as adjunct therapies in mental health treatment. The exploration of personalized nutrition is also an important consideration in the application of novel probiotics and prebiotics. Individual microbiomes vary widely based on factors such as genetics, diet, and lifestyle, suggesting that tailored probiotic and prebiotic interventions could yield more effective health outcomes. Understanding how different microbial communities respond to specific strains and compounds will pave the way for personalized dietary recommendations and interventions aimed at optimizing gut health. The introduction of novel probiotics and prebiotics represents a significant advancement in the field of nutrition and health. By leveraging the latest findings in microbiome research, these innovative substances have the potential to transform dietary strategies and therapeutic approaches, contributing to improved health outcomes and enhanced quality of life for individuals across diverse populations⁶².

6. FUTURE DIRECTIONS FOR RESEARCH

The future of gut microbiome research holds tremendous potential, with a multitude of avenues for exploration that could significantly advance our understanding of the complex interplay between the microbiome and human health. As researchers continue to uncover the intricate relationships within microbial communities, several key areas are poised for further investigation. One promising direction is the development of personalized microbiome interventions tailored to individual needs. As microbiome profiles can vary greatly between individuals due to genetic, environmental, and lifestyle factors, personalized approaches could enhance the efficacy of probiotics and prebiotics. Future studies could focus on identifying specific microbial signatures associated with various health outcomes, allowing for targeted dietary recommendations and interventions. Personalized nutrition could lead to optimized health benefits by selecting strains of probiotics or types of prebiotics that align with an individual's unique microbiome composition. A deeper understanding of the mechanisms through which the gut microbiome influences health is crucial. Research could focus on elucidating the biochemical pathways involved in microbiome-host interactions, including how microbial metabolites like short chain fatty acids (SCFAs) affect immune function, metabolism, and mental health. By identifying specific compounds produced by gut bacteria that mediate health effects, researchers could develop targeted therapies that leverage these mechanisms. Understanding these pathways may also provide insights into how dysbiosis contributes to disease, opening avenues for preventative measures⁶³.

Longitudinal studies examining the gut microbiome over time are essential to understanding how shifts in microbial composition affect health throughout an individual's life. Research could investigate how factors such as diet, age, medication, and environmental exposures influence the microbiome's evolution. Such studies could provide critical insights into the timing and nature of interventions needed to maintain a healthy microbiome, particularly during key life stages like infancy, childhood, and aging. Further exploration of the gut microbiome's role in chronic diseases is another significant area for future research. While current studies have established links between dysbiosis and conditions such as obesity, diabetes, and inflammatory bowel disease, there remains much to uncover about the microbiome's involvement in other diseases, including autoimmune disorders and neurodegenerative conditions. Understanding how specific microbial communities or metabolites contribute to disease pathogenesis could lead to novel therapeutic strategies and preventive measures⁶⁴.

The development of innovative therapeutic approaches utilizing the microbiome is an exciting frontier. Beyond faecal microbiota transplantation (FMT), future research could explore new methods for administering probiotics, such as encapsulation technologies that enhance the survival of beneficial strains during gastrointestinal transit. Furthermore, the creation of synbiotics combinations of prebiotics

and probiotics tailored for specific health conditions could maximize their health benefits. Researchers could also investigate the therapeutic potential of microbial consortia or engineered probiotics designed to produce beneficial compounds directly in the gut⁶⁵.

Future research on the gut microbiome holds promise for advancing public health by investigating how various diets and lifestyle factors such as dietary patterns, physical activity, sleep, and stress affect gut microbial diversity and function. These insights could guide dietary guidelines and lifestyle recommendations to support a healthy microbiome and prevent disease. Exploring the interaction between gut bacteria and the immune system could yield new strategies for managing autoimmune diseases and allergies, with particular attention to the influence of gut bacteria on immune cells like T-cells and regulatory cells. Technological advancements in metagenomics, metabolomics, and bioinformatics are transforming microbiome research by enabling detailed analyses of microbial communities and their functions. As sequencing technologies become more accessible, large-scale studies will facilitate the identification of microbial patterns linked to health outcomes globally. Enhancing public awareness of the microbiome's role in health is essential for encouraging lifestyle choices that promote gut health. Moving forward, interdisciplinary approaches combining microbiology, nutrition, immunology, and psychology will be key in developing innovative microbiome-focused strategies to enhance health and improve quality of life worldwide⁶⁶.

7. CONCLUSION

The gut microbiome is an essential aspect of human health that contains trillions of microorganisms, including bacteria, viruses, fungi, and archaea. These microorganisms are functioning together collectively to influence physiologic processes. The diversity of the microbiome increases metabolic fermentation and mediates both immune functioning, mood and or aspect of mental health when dysbiosis is present. The literature identifies the gut microbial compounds involved in nutrient metabolism, vitamin synthesis, and fermentation of fibre to produce short-chain fatty acid (SCFA) including those such as butyrate, acetate, and propionate. SCFA are recognized in the literature as mediators in gut integrity and inflammation signalling and signalling in other organ systems mediating in aspects of appetite and energy regulation and or immune system mediation. Dysbiosis is associated with many health conditions. For example, gut microbiomes associated or found in obese humans tend to show a greater Firmicutes to Bacteroidetes ratio which allows for energy preservation, which increases energy harvest from food. Within the framework of inflammatory bowel disease (IBD), dysbiosis occurs as a result of decreased diversity of beneficial bacteria and an excessive abundance of harmful species, leading to increased flare-ups and worsening symptoms. Future research will expand into personalized feeding strategies based on microbial phenotype, including probiotics and prebiotics specifically tailored to individuals microbial signature highlighting the importance of microbiome profile as a predictor to obtain desired benefits from dietary and therapeutic processes in aiding health conditions. Understanding the mechanisms involved in intestinal-microbiome-host interactions will be critical, particularly following the gut-brain axis indicating how gut microbes impact neurophysiology and mental health; they modify/influence neurotransmitter levels through the gut to affect stress. This will generate preliminary pathways for microbiome fed dietary strategies related to mental health, in addition to targeted microbiome therapies.

Longitudinal studies are essential for tracking gut microbiome dynamics over time, especially concerning lifestyle changes and health transitions. This research is crucial during significant life stages, such as infancy, childhood, and aging, where the microbiome significantly impacts development and health maintenance. Additionally, further exploration of the gut microbiome's role in chronic diseases like autoimmune disorders, cardiovascular diseases, and metabolic syndrome can provide insights into disease progression and inform innovative therapeutic strategies. Investigating the impact of various diets and lifestyle factors on the gut microbiome will be essential for developing effective public health strategies. Research could focus on understanding how dietary patterns, physical activity, sleep, and stress influence gut microbial diversity and function. This knowledge could inform dietary guidelines and lifestyle recommendations aimed at promoting a healthy microbiome and preventing related diseases. The interplay between the gut microbiome and the immune system presents another compelling area for future research.

Understanding how gut bacteria influence immune responses may help us to develop new mechanisms to treat autoimmune diseases and allergies. Future research may address how the microbiome shapes

the development and functioning of different immune cell types, including T cells and regulatory cells, and if and how we might be able to modulate those interactions for therapeutic purposes. Advances in technology, such as metagenomics, metabolomics, and bioinformatics will continue to change the landscape of microbiome research to improve our understanding of how to analyze microbial communities and their functional capacities in increasingly sophisticated ways. As sequencing technologies become much more affordable and broadly accessible, it will increase the feasibility of population-scale studies to identify patterns of microbial composition associated with health and disease across the globe. Increasing public awareness about the microbiome's role in health is critical to being able to engage in positive dietary and lifestyle change. Going forward, research should focus on understanding how to communicate to the public about gut health and dietary interventions that help the microbiome. This knowledge can help shape choices that will lead general populations to being more mindful of promoting diversity of the microbiome for their overall health. The future of the gut microbiome will hold critical advances in understanding its complexities. Recruiting interdisciplinary approaches to research involving microbiology, nutrition, immunology, and psychology will be paramount to grasping the gut microbiome's implications for health. Providing targeted research focused on the microbiome could develop innovation that uses and possibly capitalizes on the microbiome to enhance health, prevent, and develop conditions.

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