

Aerobes in the Gut Microbiota – Roles, Interactions, and Implications for Host Health

Sania Ikram¹, Zoha Waheed Abbasi¹, Aroosa Imtiaz Khan², Javeria Kulsoom¹, Muhammad Amir Khan³, Shahid Ullah Khan^{4*}

¹Department of Biological Sciences, National University of Medical Sciences (NUMS), Rawalpindi, Punjab, Pakistan

²Department of Pharmacy, Riphah International University, Islamabad, Punjab, Pakistan

³Department of Foreign Medical Education, Fergana Medical Institute of Public Health, 2A Yangi Turon Street, Fergana 150100, Uzbekistan

⁴Department of Biomedical Sciences, Dubai Medical College for Girls, Dubai Medical University, Dubai 19099, United Arab Emirates

Email: shahid.zkd35@yahoo.com

Abstract – The human gut contains a dynamic and rich microbial community comprising aerobic and anaerobic bacteria that live in symbiosis with the host and contribute to host physiology and gut homeostasis. Anaerobes, which predominate in the gut microbiota and have been well characterized, are contrasted with relatively lesser-known aerobic bacteria. Recent data indicate that aerobes, including facultative anaerobes, perform essential functions during early colonization, nutrient metabolism, immune modulation, and resistance against pathogens. However, the interactions between aerobes, anaerobes, and host immune responses remain poorly understood. Factors, such as diet, antimicrobial exposure, oxygen gradients, and host physiology, determine the composition and function of these microbes. A significant research gap exists in the form of limited methodologies for examining aerobic bacteria in the predominantly anaerobic gut environment. The improvements in sequencing technologies, metagenomics, transcriptomics, and in vitro gut models have helped understand microbial dynamics in gut environment. This review evaluates current methodologies for studying gut aerobes, their limitations, and their role in microbial ecology, health, and disease.

Keywords – Gut-microbiota, Aerobic bacteria, Anaerobes, Microbial Interaction, Oxygen utilization, Immune modulation, Gut homeostasis.

1. Introduction

This The human gut hosts a diverse microbial community, including bacteria, archaea, viruses, and fungi, which play a crucial role in shaping host physiology [1,2]. Among them, bacteria are the most studied due to their abundance and metabolic versatility [3]. While anaerobes dominate, a small but significant fraction of aerobic bacteria thrives in localized oxygen-rich pockets, particularly in the cecum and terminal ileum [4-6]. These aerobes contribute to immune regulation, gut barrier maintenance, and metabolic interactions with other microbes. Though outnumbered by strict anaerobes, they play a vital role in maintaining microbial equilibrium [7-9].

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The gut microbiota regulates immunity, metabolism drug processing and mucosal integrity with factors like diet, antibiotics and genetics shaping its composition [20-22]. Imbalance in gut lead to obesity, allergies and inflammatory disease [23-26]. Facultative aerobes (Escherichia, Enterococcus, Lactobacillus) support carbohydrate

degradation, SCFA production and redox balance, aiding anaerobic growth [27,28]. Gut microbes interact competitively or cooperatively with each other producing antimicrobial metabolites and engage in cross feeding [29,30]. Culture independent approaches like metagenomics, meta transcriptomics and single cell sequencing now deeper insight into their roles and interaction.

Early life colonization of facultative aerobes is critical for establishing a stable gut microbiota [31]. Factors like delivery mode, breastfeeding and antimicrobial exposure influence this transition, impacting long term health [32-35]. This review explores gut aerobes, their study methods, functions, interaction and therapeutic applications

2. Distribution of Aerobes in Various Gut Regions:

The human GI tract hosts a diverse microbial community influenced by oxygen levels, pH, nutrients, and host defenses [36]. Aerobic bacteria vary across gut regions, impacting microbial interactions and host physiology. In the upper GI tract, the mouth harbors aerobes like Neisseria, Streptococcus, and Rothia, forming biofilms [37]. The esophagus carries transient aerobes, while the stomach, despite its acidity, supports Helicobacter pylori and low number of Streptococcus and Lactobacillus [38,39]. The small intestine transition from aerobes tot aerobes with, Enterococcus, Lactococcus and Streptococcus, Escherichia coli and Bacillus dominating duodenum, jejunum and ileum followed by anaerobic dominant environment in colon where oxygen diffusion creates microaerophilic niches where Aerobes like Actinobacter and Pseudomonas reside in intestinal

mucous influencing host immunity and gut homeostasis [40].

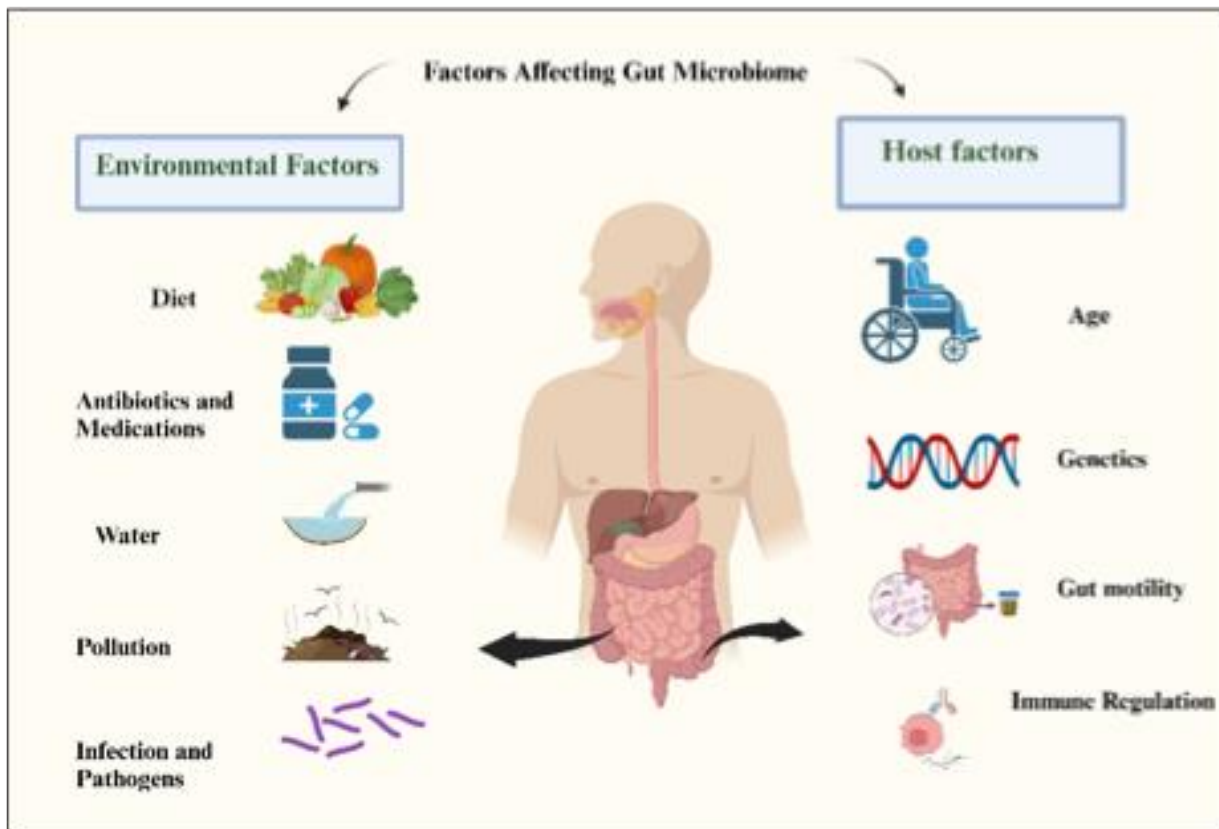


Figure 1. Environment and host factors affecting the Human Gut Microbiome

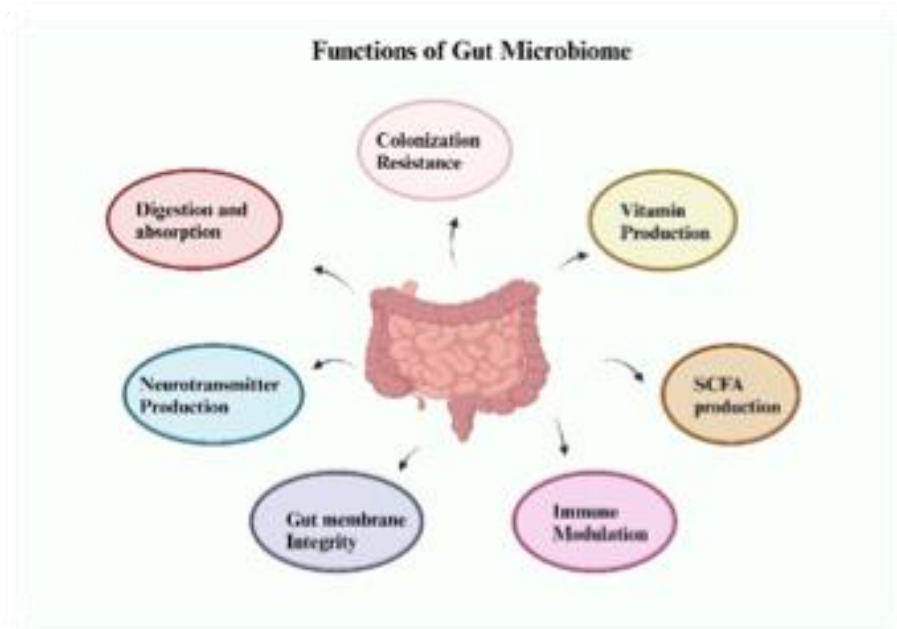


Figure 2. Functions of Gut Aerobes showing some essential features.

3. Methods for Studying Aerobic Bacteria:

Scientists study aerobic gut bacteria using both culture-dependent and culture-independent methods. Culture-based approaches isolate bacteria for detailed phenotypic and genotypic analysis [41,42]. Selective and differential media, such as MacConkey agar, enable targeted bacterial growth, while high-throughput Culturomics expands the range of culturable species [43]. Oxygen gradient cultivation further aids in studying bacteria with varying oxygen tolerances. Despite providing valuable physiological insights, culture-dependent methods are limited to bacteria that can be grown in the lab. Recent advancement in automated culturing techniques have further improved bacterial isolation efficiency, making it easier to study fastidious organisms.

Culture-independent techniques, such as 16S rRNA sequencing and metagenomics, allow for comprehensive microbial analysis without the need for cultivation [44,45]. 16S rRNA sequencing identifies bacterial taxa, while metagenomics sequences entire microbial genomes, revealing functional potential. Meta transcriptomics analyzes gene expression, providing insights into bacterial activity under specific conditions, while meta proteomics examines protein functions within the gut microbiota [46]. Advances in single-cell genomics have identified novel bacterial species, furthering knowledge of microbial contributions to gut homeostasis. Integrated machine learning with sequencing data has enhanced the ability to predict bacterial function and interactions offering new perspectives on gut microbiome dynamics. Combining Culturomics with sequencing-based approaches bridges the gap between microbial presence and function, enhancing our understanding of gut bacteria and their role in health and disease.

4. Intestinal Colonization and their Functional Roles:

Intestinal colonization by aerobic bacteria plays a crucial role in gut health by modulating oxygen levels, interacting with immune cells, and producing beneficial compounds. These interactions help maintain a balanced microbiota, preventing pathogenic colonization and supporting overall intestinal function. For instance, *Lactococcus lactis* can perform aerobic respiration in the presence of exogenous heme, reducing intestinal oxygen levels. This process promotes autophagy in intestinal cells, lowers *Salmonella* load, and improves gut microbiota composition, thereby enhancing immune balance and reducing inflammation [47-49]. Similarly, aerobic spore-formers like *Bacillus subtilis* and *Alkalihalobacillus clausii* interact with intestinal and immune cells, producing antimicrobial compounds and contributing to immune system development [50].

Gut aerobic interaction shape microbial stability and host health through spatial organization, metabolic exchanges and environmental factors. *Aeromonas ZOR0001* form spatial aggregates, but *Aeromonas MB4* disrupts them, reducing *Enterobacter* abundance [51-53]. Metabolic cross-feeding such as between *Acetobacter fabarum* and *Lactobacillus brevis* enhances microbial

growth and efficiency. Furthermore, *Brevibacillus massiliensis* detoxifies aldehydes via a tungsten-containing oxidoreductase, enhancing gut resilience [54,55]. Additionally aerobic bacteria modulate host immune responses, influencing inflammation and gut barrier integrity. These interactions collectively shape microbiome composition, offering insights for micro-biome targeted therapies to improve gut health.

Aerobes in the human gut, though less abundant than anaerobes, play essential roles in detoxification, nutrient metabolism, and maintaining microbial balance. They aid in bile acid conversion, crucial for fat absorption, and contribute to vitamin K synthesis, essential for blood clotting and bone health [56,57]. Additionally, aerobes help regulate microbial balance by interacting with anaerobes, preventing pathogenic overgrowth, and competing for resources to defend against infections. However, their survival and activity are influenced by several factors, including oxygen availability, diet, pH, and host immune responses. Understanding these factors and their functional contributions can provide insights into therapeutic strategies for gut-related disorders.

5. Conclusion and Future Perspectives

Aerobic bacteria play a vital role in gut health by modulating oxygen levels, producing beneficial metabolites, and interacting with immune cells. They contribute to microbial balance, preventing pathogenic colonization and supporting intestinal homeostasis. Culture-dependent and culture-independent methods have provided significant insights into their diversity, metabolic activities, and ecological roles. Advances in sequencing technologies, metagenomics, and proteomics have expanded our understanding of their functions, while AI-driven imaging and high-throughput culturing techniques have improved bacterial identification and characterization [61]. Despite these advancements, challenges remain in linking microbial presence to function and understanding their real-time dynamics in the gut environment [62-65].

Moreover, personalized microbiome research will play a crucial role in understanding individual variations in gut aerobic bacterial composition and function. Future studies should focus on longitudinal analyses to track microbial changes over time and their correlation with health outcomes. The integration of machine learning and AI in microbiome research will enhance predictive modeling, enabling the development of precision medicine approaches [66,67]. Additionally, exploring novel bioengineering techniques to modify gut bacteria for therapeutic benefits could revolutionize microbiome-based treatments. Expanding research on host-microbe interactions at the cellular level will provide deeper mechanistic insights, paving the way for innovative strategies to manipulate gut microbiota for disease prevention and treatment. Furthermore, advancing multi-omics approaches, including metagenomics, metatranscriptomics, and metabolomics, will provide a comprehensive understanding of the functional roles of aerobic gut bacteria. These techniques will help unravel microbial metabolic pathways, resistance mechanisms, and their influence on host physiology. Integrating these

datasets with clinical findings will bridge the gap between microbiome research and translational medicine, facilitating targeted interventions for gut-related disorders.

Table 1. Methods for studying Intestinal Microbes Pros and Cons.

| Method | Description | Advantages | Limitations | References |
|--|---|--|---|------------|
| SELECTIVE AND DIFFERENTIAL MEDIA | Uses specialized media to distinguish between colonies | Enables isolation of viable bacteria for subsequent phenotypic examination | Most intestinal bacteria are not culturable | [58] |
| High throughput Phenotypic identification (MALDI-TOF MS) | Uses mass spectrometry to quickly identify bacterial species | Quick and Precise identification of cultured isolates | Need pure cultures, will not identify unculturable organisms | [58] |
| 16SrRNA Gene Sequencing | Sequencing all the genetic element with in sample to enable full analysis of bacterial speies | Global evaluation of microbial diversity and functional genes. Reveals new organisms | Very expensive and computationally intensive. Difficulty in assembling and interpreting complex data. | [59] |
| Meta proteomics | Examines the entire protein complement of a microbial community. | Direct evidence of microbial function and activity. | Highly technical with difficult sample preparation. Needs advanced instrumentation and bioinformatics software. | [60,61] |
| AI-Based Imaging Techniques | Leverages artificial intelligence to examine imaging information to identify bacteria | Facilitates quick, non-culture-based identification | Quality and amount of training data dependent. Can consume large amounts of computational power. | [61] |

Additionally, future studies should investigate the impact of environmental factors, such as diet, antibiotics, and lifestyle, on gut aerobic bacteria. Understanding how these factors shape microbial communities will aid in designing dietary and probiotic interventions to restore microbial balance. Collaborative research between microbiologists, clinicians, and bioinformaticians will be essential in developing novel microbiome-based therapeutics

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Conflict of Interest

Authors have no conflict of interest to declare.

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