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From Kitchen to Clinic: Translational Evidence for Functional Foods in Chronic Disease Management

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ABSTRACT: Functional foods have gained attention as effective dietary strategies for preventing and managing chronic illnesses due to their abundance of bioactive compounds such as polyphenols, dietary fibers, probiotics, omega-3 fatty acids, carotenoids, and bioactive peptides. Increasing scientific research indicates that these elements provide antioxidant, anti-inflammatory, immunomodulatory, and metabolic regulatory benefits, which help in lowering the risk and progression of significant chronic conditions like cardiovascular disease, type 2 diabetes mellitus, obesity, cancer, neurodegenerative disorders, and gastrointestinal diseases. This review delves into the evolution of functional foods from traditional dietary practices in kitchens to their application in evidence-based clinical settings. It emphasizes the molecular mechanisms that contribute to their health benefits and critically assesses findings from both preclinical and clinical studies that support their therapeutic potential.

Additionally, the review addresses major obstacles to clinical translation, such as variations in bioactive composition, limited bioavailability, lack of standardization, regulatory inconsistencies, and the necessity for large-scale randomized clinical trials. The review also explores emerging advancements in precision nutrition, gut microbiome research, multi-omics technologies, artificial intelligence, and digital health as transformative methods for enhancing functional food interventions and facilitating personalized dietary strategies. By combining mechanistic insights with clinical evidence and future technological advancements, this review offers a comprehensive view of the role of functional foods in

managing chronic diseases and identifies research priorities essential for their integration into routine healthcare and public health nutrition.

1. INTRODUCTION

1.1 Functional Foods and Their Importance

Functional foods, a relatively new addition to modern nutrition, are foods that are natural or processed and contain biologically active compounds that can improve physiological functions and reduce the risk of chronic diseases when consumed in moderation as part of a balanced diet [1,2]. Examples of functional foods are fruits, vegetables, whole grains, legumes, fermented dairy products, nuts, seeds, marine products, and foods fortified with vitamins, minerals, probiotics, or other bioactive ingredients.

The health benefits of functional foods are largely due to bioactive compounds, including polyphenols, flavonoids, carotenoids, dietary fibers, omega-3 fatty acids, phytosterols, probiotics, prebiotics, and bioactive peptides, which have antioxidant, anti-inflammatory, antimicrobial, immunomodulatory, and metabolic regulatory properties that maintain physiological homeostasis and decrease the incidence of chronic diseases [3–5]. A large number of epidemiological and clinical studies have shown that habitual consumption of functional foods has beneficial effects on cardiovascular health, glucose metabolism, immune function, gut microbiota composition, and quality of life [2,6], and therefore are viewed as a key element in preventive healthcare and personalized nutrition.

1.2 Global Burden of Chronic Diseases

Chronic non-communicable diseases (NCDs), such as cardiovascular diseases, type 2 diabetes mellitus, obesity, cancer, chronic respiratory diseases, and neurodegenerative disorders, are one of the largest public health challenges in the world, causing nearly three-quarters of all global deaths and remaining the leading cause of mortality, according to the World Health Organization (WHO) [7]. The increasing burden of these diseases is linked to unhealthy eating patterns, physical inactivity, aging populations, environmental exposures, and rapid urbanization [8].

Diet is considered to be one of the most important modifiable risk factors associated with chronic disease etiology and progression; diets high in ultra-processed foods with high saturated fats, refined sugars, and sodium lead to obesity, insulin resistance, oxidative stress, and chronic inflammation, while dietary patterns focused on fruits, vegetables, whole grains, legumes, healthy fats, and fermented foods have been linked to lower disease incidence and better clinical outcomes [6,9]. Hence, nutritional interventions using functional foods are being integrated into clinical guidelines and public health policies and strategies for disease prevention and long-term management.

1.3 Translational Nutrition: From Kitchen to Clinic

Translational nutrition takes findings from nutritional science and applies them to clinical medicine to translate traditional dietary knowledge into clinically proven healthcare interventions [10]. The process from discovery of health-promoting foods and their bioactive components to elucidating molecular mechanisms, preclinical studies, randomized clinical trials, and evidence-based dietary recommendations in healthcare practice is often referred to as From Kitchen to Clinic [10].

Metabolomics, nutrigenomics, proteomics, gut microbiome research, and systems biology have advanced the understanding of the molecular interactions between dietary bioactive compounds and human physiology considerably. Functional foods affect many biological pathways, including oxidative stress, inflammatory responses, lipid metabolism, glucose homeostasis, immune modulation, and the cellular signaling pathways NF- κ B, Nrf2, AMPK, PI3K/Akt, and MAPK [10,11]. In addition, precision nutrition is being supported by artificial intelligence, machine learning, digital health technologies, wearable biosensors, and multi-omics approaches that provide personalized dietary recommendations based on genetic, metabolic, microbiome, and lifestyle characteristics [11,12].

However, several barriers still hinder successful clinical translation of functional foods, including variability in the composition of bioactive compounds, poor bioavailability, lack of standardized manufacturing processes, limited high-quality clinical trials, and differences in international regulatory frameworks [1,2].

1.4 Aim and Scope of the Review

This review provides an overview of the current translational evidence in support of the use of functional foods for the prevention and treatment of chronic diseases, including classification of functional foods and major bioactive compounds, molecular mechanisms of action, evidence from pre-clinical and clinical studies on cardiovascular diseases, diabetes mellitus, obesity, cancer, neurodegenerative disorders, and gastrointestinal diseases, as well as challenges related to standardization, bioavailability, safety, and regulatory approval, and emerging advances in precision nutrition, gut microbiome research, artificial intelligence, and multi-omics technologies that will bridge the gap between nutritional science and clinical practice to translate functional foods from common dietary practices into evidence-based healthcare intervention. [10].

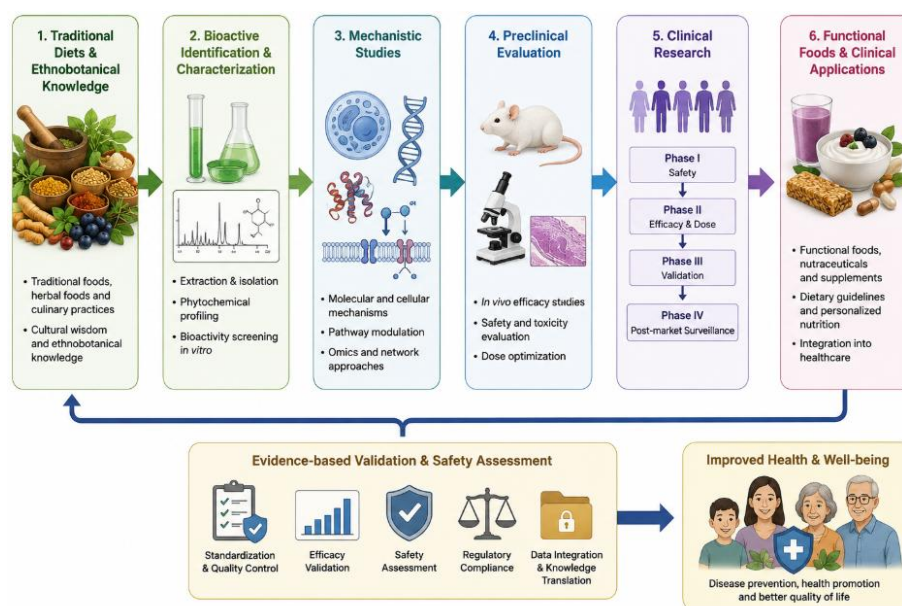


Figure 1. Translational pathway of functional foods from traditional diets to clinical applications.

2. FUNCTIONAL FOODS AND THEIR BIOACTIVE COMPONENTS

2.1 Definition and Classification

A rapidly emerging area of preventive nutrition is functional foods, which are defined as natural or processed foods that contain biologically active compounds that enhance physiological functions and reduce the risk of chronic disease when consumed regularly in the normal diet [11]. Functional foods are more in keeping with a holistic approach to disease prevention than pharmaceuticals, in that functional foods impact physiological functions through synergistic interactions between and among multiple nutrients, phytochemicals, and microbial components. The move from traditional dietary practices to evidence-based nutritional interventions for functional foods has been enhanced by advances in biotechnology, nutrigenomics, metabolomics, and microbiome research [12]. Functional foods can be categorized in a number of ways depending on their origin and formulation: natural functional foods (fruits, vegetables, whole grains, dairy), fortified functional foods (processed foods), and functional foods with added bioactive components (processed foods).

2.2 Major Bioactive Compounds

Functional foods exert their health-promoting effects through bioactive compounds that are involved in the regulation of oxidative stress, inflammation, metabolism, and immune responses. Polyphenols (flavonoids, phenolic acids, stilbenes, and lignans) are the most abundant phytochemicals in fruits, vegetables, tea, cocoa, and spices, and they have antioxidant and anti-inflammatory activities through modulation of signaling pathways like nuclear factor kappa B (NF- κ B) and nuclear factor erythroid 2-related factor 2 (Nrf2) that contribute to cardiovascular and metabolic health [14].

Dietary fibers from whole grains, fruits, vegetables, and legumes enhance gastrointestinal function, glucose regulation, lipid metabolism, and gut microbiota composition; fermentation by intestinal microorganisms produces SCFAs that regulate immune responses and metabolic homeostasis [15]. Probiotics and prebiotics also contribute to gut health by altering microbial composition.

Probiotics and prebiotics are also important for maintaining gut health by altering microbial composition and improving intestinal barrier function; fermented foods like yogurt, kefir, and fermented vegetables are good sources of beneficial microorganisms that promote immune and metabolic regulation [13]. The main dietary source of EPA and DHA is from marine sources, and these fatty acids have cardioprotective, anti-inflammatory, and neuroprotective properties through regulation of lipid metabolism and inflammatory mediators [16]. Additional bioactive compounds include bioactive peptides, carotenoids, and phytosterols that provide antioxidant defense, cholesterol regulation, and immune modulation [17].

2.3 Mechanisms of Action

There are a number of interconnected mechanisms by which functional foods exert their biological effects, including antioxidant activity (the neutralization of ROS and the enhancement of endogenous antioxidant defense systems, reducing oxidative damage linked to chronic diseases), and the regulation of inflammatory responses (the suppression of pro-inflammatory pathways and the reduction of cytokine production) [14].

Functional food components also modulate inflammatory responses by downregulating pro-inflammatory pathways and inhibiting cytokine production; polyphenols and omega-3 fatty acids reduce inflammatory mediators that are associated with chronic disease progression, such as IL-6, TNF- α , and NF- κ B signaling pathways [16].

Additionally, it also impacts metabolic health through enhanced insulin sensitivity, lipid metabolism, and energy balance; dietary fibers and polyphenols regulate glucose homeostasis, while probiotics modulate gut microbiota-derived metabolites that influence immunity, metabolism, and brain function [15]. Additionally, there is emerging evidence that dietary bioactive compounds can influence gene expression and epigenetic mechanisms, providing an additional layer of support for the development of precision nutrition approaches [18].

In summary, functional foods serve as the scientific link between traditional dietary practices and contemporary healthcare by incorporating nutritional, molecular, and clinical approaches to improve chronic health.

Table 1. Classification of functional foods, bioactive compounds, food sources, and health benefits

Functional Food Category	Major Bioactive Compounds	Representative Food Sources	Major Health Benefits
Fruits and Vegetables	Polyphenols, flavonoids, carotenoids, vitamins	Berries, citrus fruits, tomatoes, leafy vegetables	Antioxidant protection, cardiovascular health, reduced inflammation
Whole Grains	Dietary fiber, phenolic compounds, beta-glucans	Oats, wheat, barley, brown rice	Improved glycemic control, cholesterol reduction, gut health
Fermented Foods	Probiotics, peptides, organic acids	Yogurt, kefir, kimchi, fermented vegetables	Improved gut microbiota, immune regulation, digestive health
Marine Foods	Omega-3 fatty acids, peptides	Fish oil, salmon, seaweed	Cardioprotection, anti-inflammatory and neuroprotective effects
Nuts and Seeds	Phytosterols, polyphenols, healthy fats	Almonds, walnuts, flaxseed, chia seeds	Improved lipid metabolism, antioxidant effects

Legumes	Fiber, resistant starch, bioactive peptides	Beans, lentils, chickpeas, soy products	Blood glucose regulation, metabolic health
Tea and Cocoa Products	Catechins, flavonoids, polyphenols	Green tea, black tea, cocoa	Vascular protection, antioxidant activity
Fortified Functional Foods	Added vitamins, minerals, probiotics, omega-3	Fortified milk, cereals, beverages	Nutrient improvement and disease prevention

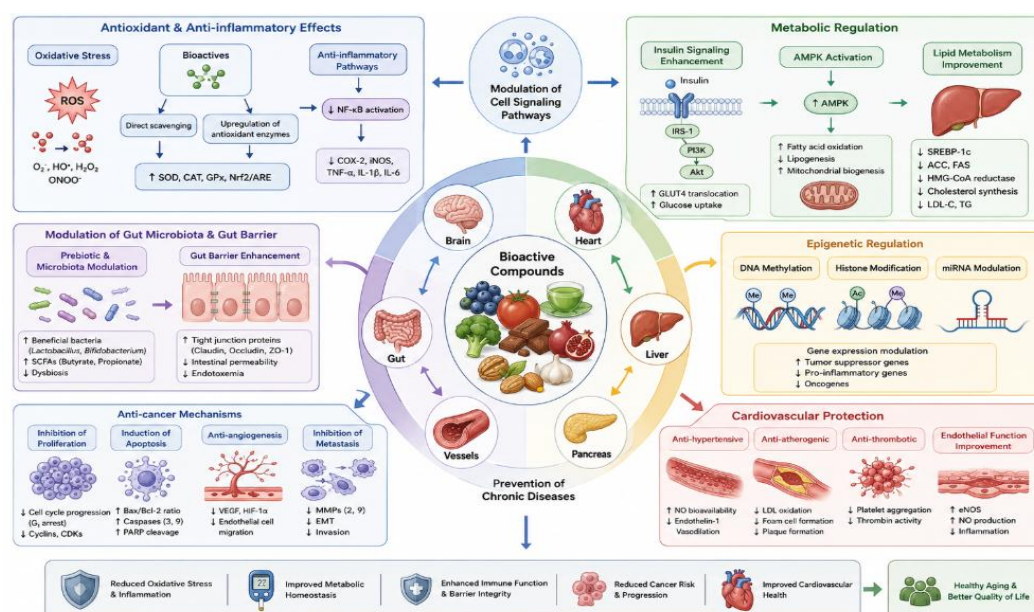


Figure 2. Molecular mechanisms of functional food bioactives in chronic disease prevention.

3. CLINICAL EVIDENCE FOR FUNCTIONAL FOODS IN CHRONIC DISEASE MANAGEMENT

In addition to basic nutrition, functional foods contain bioactive compounds (polyphenols, dietary fibers, probiotics, omega-3 fatty acids, and phytochemicals) that offer health benefits beyond basic nutrition. Growing clinical evidence indicates that consumption of functional foods can contribute to the prevention and management of chronic disease by enhancing metabolic regulation, reducing inflammation, and modulating oxidative stress.

3.1 cardiovascular diseases

Foods that have been found to be beneficial in CVD include foods rich in omega-3 fatty acids (fatty fish, flaxseed), soluble fibers (oats, barley), plant sterols, and polyphenols (berries, tea, cocoa), which have been linked to reduced LDL cholesterol, improved endothelial function, and lowered inflammatory markers. Clinical evidence has indicated that consumption of oat β -glucan can lower serum cholesterol, and omega-3 fatty acids can reduce triglycerides and cardiovascular risk factors.

3.2 Type 2 Diabetes Mellitus

Functional foods are effective in improving glucose metabolism and insulin sensitivity, dietary fibers, resistant starch, probiotics and polyphenol-rich foods help maintain postprandial glucose levels and decrease insulin resistance; clinical trials employing whole grains, legumes, cinnamon, and fermented foods have shown improved glycemic control and blunted inflammatory responses in subjects with Type 2 diabetes.

3.3 Obesity and Metabolic Syndrome

Functional foods can aid weight management by increasing satiety, improving lipid metabolism, and modulating gut microbiota, and high-fiber foods, probiotics, green tea catechins, and protein-rich functional foods have demonstrated positive effects on body weight reduction, abdominal fat accumulation, and metabolic parameters, respectively. Modulation of gut microbiota through probiotics and prebiotics is an emerging strategy for the treatment of obesity-related disorders.

3.4 Cancer Prevention

Bioactive compounds found in functional foods, such as antioxidant, anti-inflammatory, and anticancer bioactive compounds in cruciferous vegetables (sulforaphane), berries (anthocyanins), turmeric (curcumin), and green tea (catechins), have shown promise in reducing oxidative DNA damage and modulating pathways associated with cancer development, although clinical evidence for protective effects exists.

3.5 Neurodegenerative Disorders

While these functional foods may support brain health by reducing neuroinflammation and oxidative stress, omega-3 fatty acids, flavonoids, polyphenols, and fermented foods have been researched for their role in cognitive function and neuronal protection, and clinical studies have shown the components of the Mediterranean diet, along with berries, nuts, and fish consumption are correlated with better cognitive function and decreased risk for neurodegenerative disorders, such as Alzheimer disease.

3.6 Gastrointestinal Disorders

The clinical benefits of functional foods with respect to gut microbiota and intestinal barrier function are evident in conditions like IBS, IBD, and constipation, where probiotics, prebiotics, fermented foods, and dietary fibers have all been shown to have positive effects, particularly with specific probiotic strains.

Table 2. Major Clinical Studies Evaluating Functional Foods in Chronic Diseases

Functional Food / Bioactive Component	Disease Condition	Major Findings	Clinical Evidence
Oat β -glucan	Cardiovascular disease	Reduced LDL cholesterol and improved lipid profile	Clinical trials showed cholesterol-lowering effects with regular intake
Omega-3 fatty acids (EPA/DHA)	Cardiovascular disease	Reduced triglycerides and improved heart health markers	Multiple human studies demonstrated cardioprotective effects
Whole grains and dietary fiber	Type 2 diabetes	Improved glycemic control and insulin sensitivity	Associated with reduced HbA1c and better glucose regulation
Probiotics (Lactobacillus, Bifidobacterium)	Diabetes and metabolic syndrome	Improved gut microbiota balance and inflammation control	Clinical studies reported metabolic improvements
Green tea catechins	Obesity and metabolic syndrome	Enhanced fat oxidation and reduced body weight parameters	Randomized trials showed modest weight-management benefits
Turmeric/Curcumin	Cancer prevention and inflammation	Reduced inflammatory markers and oxidative stress	Human studies support anti-inflammatory activity
Berries (anthocyanins)	Neurodegenerative disorders	Improved cognitive function and antioxidant capacity	Clinical trials reported cognitive benefits in older adults
Fermented foods	Gastrointestinal disorders	Improved intestinal microbiota and digestive health	Clinical evidence supports probiotic effects

4. TRANSLATIONAL CHALLENGES AND EMERGING TECHNOLOGIES

Although functional foods hold promise for their health benefits, their implementation in clinical settings faces a number of challenges related to bioavailability, safety, personalization, and technological integration, which presents new opportunities for advancing functional food development and use with innovative technologies such as AI, digital health, and microbiome analysis.

4.1 Bioavailability and Standardization

A key issue in functional food research is the reduced bioavailability of numerous bioactive compounds. Aspects such as inadequate absorption, swift metabolism, and interactions with other dietary elements can diminish their efficacy. Standardizing active compounds, optimizing dosages, and creating sophisticated delivery systems like nanoencapsulation and microencapsulation are crucial approaches to improve their stability and biological function.

4.2 Safety and Regulatory Considerations

For functional foods to be accepted and used widely, their safety and quality must be guaranteed. Regulatory obstacles include variations in composition, possible contamination issues, and the lack of uniform norms. To ensure consumer safety and validate health claims, sufficient toxicological evaluations, strict quality control, and evidence-based legislation are required.

4.3 Precision Nutrition and Gut Microbiome

The composition of the gut microbiome, lifestyle, and genetics all affect how each person reacts to functional meals. Precision nutrition methods predict individual responses using metabolic data, microbiome analysis, and customized food plans. Using probiotics, prebiotics, and dietary changes to alter gut microbiota is a promising strategy for managing and preventing specific diseases.

4.4 Artificial Intelligence and Digital Health in Functional Food Research

Functional food research is being revolutionized by artificial intelligence and digital health technologies, which allow data-driven discovery and personalized recommendations based on dietary patterns, clinical data, microbiome information, and molecular interactions, along with the real-time monitoring of dietary behaviors and health outcomes by wearable devices and digital health platforms, accelerating the development of personalized functional food interventions.

5. FUTURE PERSPECTIVES

Personalized Functional Foods

Personalized nutrition based on individual genetics, metabolism, lifestyle, and gut microbiome profiles is the future of functional foods, with customized dietary formulations that may help prevent disease and improve therapeutic outcomes.

Multi-omics Approaches

The combination of genomics, metabolomics, proteomics, and microbiomics will aid in the identification of bioactive compounds and their mechanisms of action, facilitating the development of evidence-based functional foods.

Sustainable Functional Food Development

Future functional foods will be developed from sustainable approaches using plant-based ingredients and food by-products, environmentally friendly processing methods, and improved nutritional and ecological benefits.

Future Research Directions

Large-scale clinical trials, standardized evaluation methods, improved bioavailability, and the integration of artificial intelligence to discover and design the next generation of functional foods should be emphasized in future studies.

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