

## INNOVATIVE STRATEGIES FOR IMPROVING PROBIOTIC VIABILITY USING AVOCADO-BASED PREBIOTICS AND NANOTECHNOLOGY

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### ABSTRACT

Probiotics are increasingly incorporated into functional foods; however, their efficacy is hindered by poor stability under processing and Gastrointestinal (GI) conditions. Avocado fruit and its waste contain fermentable fibers (pectin, resistant starch) and bioactive compounds that act as prebiotics. Current probiotic stabilization strategies often overlook the importance of nutrient synergy and sustainability. This review reveals a gap: conventional carriers (alginate, inulin, etc.) provide protection but little additional growth substrate, whereas avocado fibers offer both nourishment and encapsulation potential. Nanotechnology has emerged as a revolutionary field with immense potential in various industries, including the delivery of nutraceuticals. Combining avocado pectin/resistant starch with cellulose nanofibers (from plants or microbial cellulose) is innovative because it co-delivers prebiotics and mechanical defense. Future work should address remaining challenges: regulatory evaluation of edible nanocellulose and novel food status, comprehensive safety testing for ingestible nano-carriers, and deeper mechanistic studies on how avocado polysaccharides and phenolics modulate probiotic metabolism. Overall, integrating avocado-derived prebiotics into nanocellulose matrices appears to be a promising strategy for robust, targeted probiotic delivery and warrants further experimental and clinical validation.

**Keywords:** Avocados, Nutraceuticals, Nanotechnology, Regulatory, Probiotics, Prebiotics target delivery

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### INTRODUCTION

Nutraceuticals are plant-based products that derive a word from "nutrition" and "Pharmaceuticals." It's also have significant attention worldwide due to their biological activity, which eventually promotes health and well-being benefits to reduce the risk of chronic diseases of the human body. These bioactive molecules, frequently derived from natural sources, encompass a broad range of products like dietary supplements, functional foods, and herbal remedies [1]. The global interest in nutraceuticals has been increasingly acknowledged by the growing healthier option as a pivotal part of the profound connection between dietary choices and health outcomes in societies, as a great diet plays an important role, including habitual conditions, and improving the overall quality of life of human beings [2].

In recent times, the demand for nutraceuticals has immensely increased with consumers seeking natural remedies over conventional medicinal over pharmaceutical products to improve their healthy lifestyle [3]. This type of consumer interest can be attributed to several factors. Initially, the rising healthcare costs and the frequency of habitual conditions have led individuals to adopt visionary approaches to manage their health with nutraceutical products [4]. Secondly, the added frequency of nutrition-related diseases, like kidney infections and diabetes, has urged individuals to explore nutraceuticals as alternative curatives for the betterment of their health [5].

The rise of social media and digital platforms has increased the dispersion of information, empowering consumers to make informed opinions about their health and well-being regarding such products [6]. To explore the global interest in nutraceuticals by examining the key drivers of this phenomenon, the current market trends hold the scientific substantiation supporting their efficacy and safety for the betterment of human life [7]. Also, it will delve into the regulatory governing or rules of nutraceuticals across different regions. Due to the challenges and opportunities of any new product from nutraceuticals by manufacturers and including consumers far behind in knowledge [8, 9].

The multifaceted aspects of global interest in nutraceuticals also seek to give precious insight for experimenters or innovators, healthcare professionals, policymakers, and individuals seeking substantiation information on the part of nutraceuticals in promoting optimal health and complaint evaluations about those products on a day-to-day basis [10]. In recent times, nanotechnology has shown as a promising field of research in the researcher's mind, also it has not evolved only health sector but also expanded to different sectors within diverse disciplines, including drug formulation, electronics, and energy evaluation. One area that has considerable attention is the integration of nanotechnology within the nutraceutical industry. Nutraceuticals, which encompass functional foods, dietary supplements, and herbal products, also offer immense health benefits beyond normal nutrition in the human body. As per market viewpoints, the concept of nutraceuticals has gained substantial traction in a short period in the consumer mind [11].

The objectification of nanostructured materials in the nutraceutical industry presents instigative openings to enhance their efficacy, bioavailability, and targeted delivery [12]. To explore the different investigation approaches underlying the nutraceuticals incorporated into nanostructured materials, will highlight the advantages and challenges linked to their application or utilization of the product development process in the future. Nanostructured materials, defined as materials with dimensions in the nanometre range, retain unique physical, chemical, and natural properties compared to other bulk materials that have been used in manufacturing this type of product [13, 14]. The Influence of substandard materials at the nanoscale formulation production enables precise control over their physicochemical characteristics, similar to particle size, surface area, and surface charge, which can greatly impact their relations with natural systems [15].

These properties render nanostructured materials ideal candidates for enhancing the performance of nutraceuticals during formulation or product manufacturing. One of the crucial advantages of incorporating nanostructured materials in nutraceutical formulation is their potential to enhance bioavailability [16, 17]. Numerous bioactive composites are set up in nutraceuticals but have poor

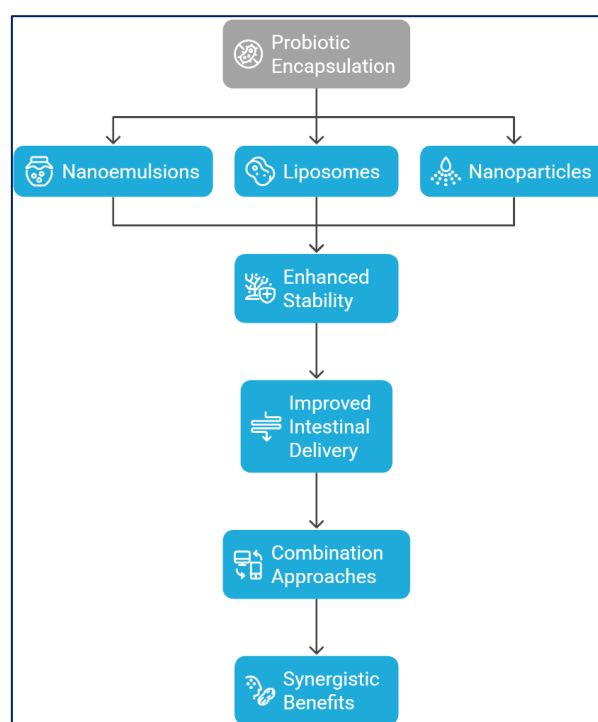
solubility and stability, limiting their bioavailability and bioaccessibility in the human body. By employing nanostructured carriers, similar to nanoparticles, liposomes, and nanoemulsions, it's possible to synthesize and cover these composites, enabling their effective delivery to target tissues like Pancreatic Islets, Skeletal Muscle, Endothelial Cells, Beta Cells kidneys, also adding their bioavailability to improve human health [18].

Likewise, nanostructured materials offer openings for the controlled and targeted release of bioactive composites. Through face variations and functionalization, nanocarriers can achieve something to respond to specific stimuli, similar to pH, temperature, or enzymes, enabling the release of nutraceuticals at the desired point of action in the human body. This targeted delivery approach aims to minimize off-target effects, enhance therapeutic efficacy, and reduce potential side effects associated with high-dose systemic administration [19]. In addition to bettered bioavailability and targeted delivery, nanostructured materials can also enhance the stability and shelf life of nutraceutical products. Numerous bioactive composites are prone to degradation, oxidation, or reactions with other constituents, leading to reduced efficacy and compromised quality. Still, the integration of nanostructured materials in nutraceuticals also presents certain challenges that need to be addressed. A thorough evaluation is necessary for safety

considerations, especially regarding the potential toxicity and long-term effects of nanomaterials [20].

Regulatory frameworks governing the use of nanomaterials in food and supplements need to be established to ensure consumer safety [21]. Likewise, large-scale products and cost-effectiveness of nanostructured materials for nutraceutical product development or operations pose specialized and profitable challenges that warrant further exploration and growth of the product. The incorporation of nanostructured materials in nutraceuticals offers significant potential for optimizing their effectiveness, bioavailability, and targeted delivery for the betterment of product success with the help of taking different measures as per policy or regulation [22]. Nanoencapsulation methods can enhance the stability, solubility, and controlled release of bioactive composites, opening new avenues for the development of innovative nutraceutical products for consumers. Still, careful consideration of safety, regulatory policy, and product aspects is essential to ensure the successful integration of nanostructured materials in nutraceutical products [23].

Continued exploration and collaboration between nanotechnology and the nutraceutical diligence scientific community will pave the way for the consummation of this innovative formulation of nutraceutical products and the advancement of a substantiated and effective nutraceutical future world on a global scale [24, 15].



**Fig. 1: Innovative and advanced delivery systems for enhancing probiotic viability and functional activity**

Probiotics play a crucial role in maintaining gut health, immune function, and overall well-being. However, one of the key challenges in probiotic-based formulations is their stability during storage, processing, and digestion. Prebiotics, which serve as non-digestible food components that promote the growth of beneficial microorganisms, have been increasingly studied for their potential to enhance probiotic viability [24]. Avocado-derived prebiotics, rich in dietary fibers, polyphenols, and oligosaccharides, offer a novel approach to improving the stability and functionality of probiotics. The unique polysaccharide composition and antioxidant properties of avocado-derived prebiotics help protect probiotics from environmental stressors while promoting their growth in the gut [25, 26].

Avocado seed contains up to 58.4% dietary fiber, including complex carbohydrates that resist digestion but ferment in the colon, producing beneficial Short-Chain Fatty Acids (SCFAs) like acetate

and butyrate. Avocado peel and pulp are rich in polyphenolic compounds that possess antimicrobial and antioxidant properties, potentially enhancing probiotic viability under stress. As a sustainable by-product of the food industry, avocado waste (seed, peel) offers a low-cost, eco-friendly source of functional prebiotics, reducing agricultural waste [27].

Probiotics, despite their documented health benefits, often fail to deliver therapeutic effects due to poor survival during food processing, storage, and Gastrointestinal (GI) transit. Exposure to acidic pH, bile salts, high temperatures, and oxidative stress drastically reduces the viability of probiotics before they reach the colon. Studies report up to 99% loss in viability of *Lactobacillus* strains during industrial processes and GI passage. This instability compromises product efficacy, shelf life, and consumer trust, highlighting a critical need for innovative stabilization approaches

that ensure both survivability and functionality of probiotics throughout the delivery process [28, 29].

*Lactobacillus rhamnosus* retained  $9.03 \times 10^7$  CFU/g in avocado-cacao fruit paste after 40 d at 4 °C. Electrospun fibers of gum arabic/pullulan showed ~97.8% survival of probiotics at 28 d. Consumption of 140g of avocado daily for 12 w increased fecal acetate by 18% and elevated populations of *Prevotella*, *Ruminococcus*, and other fiber-fermenting species [30].

Nanotechnology, particularly nanoencapsulation strategies, has already been widely explored for enhancing the bioavailability and targeted delivery of nutraceuticals [26, 30]. Integrating avocado-

derived prebiotics with cellulose-based matrices provides a sustainable, biocompatible, and effective means of ensuring probiotic protection. The synergy between nanotechnology and prebiotic carriers represents a promising avenue for delivering stable, bioavailable, and highly effective probiotic formulations in functional foods and nutraceuticals [31].

Nanocarriers are nanoscale delivery systems, typically ranging between 1–1000 nm, designed to transport bioactive compounds such as probiotics or nutraceuticals. These include liposomes, polymeric nanoparticles, nanoemulsions, dendrimers, and solid lipid nanoparticles, which enhance bioavailability, protect probiotics from degradation, and allow for targeted or sustained release [32].

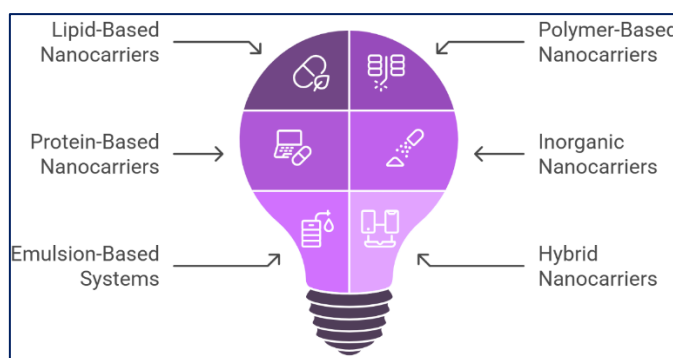


Fig. 2: Detailed insight into the types of nanocarriers employed in modern nutraceutical and probiotic formulations

A systematic literature search was conducted using PubMed, Scopus, Web of Science, and Google Scholar, covering publications from January 2018 to March 2025. The search strategy included combinations of terms such as “avocado prebiotic,” “avocado seed fibre,” “cellulose matrix,” “nanocellulose probiotic encapsulation,” “nano-encapsulation+probiotic,” and “stability of probiotics.” Inclusion criteria were limited to peer-reviewed articles published in English that focused on probiotic stability, encapsulation technologies, or the prebiotic effects of avocado or its components, including both experimental and review articles offering quantitative data or mechanistic insights. Exclusion criteria involved non-peer-reviewed reports, editorials, studies lacking relevant outcome measures related to probiotic viability, or those focusing solely on the pharmacological applications of avocado without a microbiome-related context.

### Nanostructures in nutraceuticals

Nanostructured materials, including nanoparticles, liposomes, nanoemulsions, and nanofibers, offer unique advantages in nutraceutical delivery [33]. These materials can ameliorate bioavailability, enable controlled release, and grease targeted delivery of bioactive composites. Still, challenges similar to safety, regulatory considerations, scale-up and manufacturing issues, and stability need to be addressed for the successful integration of nanostructured materials in nutraceutical product formulations. Addressing these limitations will pave the way for the development of innovative and effective nutraceutical products, furnishing enhanced health benefits to consumers [34].

Nanotechnology has revolutionized nutraceutical formulations by offering innovative delivery systems that enhance the solubility, stability, and bioavailability of bioactive compounds [35]. Cellulose matrices, specifically nano-fibrillated cellulose and nanocrystalline cellulose, have gained attention as ideal carriers for encapsulating probiotics due to their high surface area, mechanical stability, and controlled release properties. These matrices act as protective barriers, shielding probiotics from harsh gastric conditions while ensuring their gradual release into the intestinal environment [36].

Avocado-derived prebiotics, when integrated within cellulose matrices, further improve probiotic stability by providing a

nourishing substrate that enhances probiotic survival. Avocado-derived prebiotics have been shown to increase *Lactobacillus acidophilus* viability by up to 37% under simulated gastrointestinal conditions when incorporated within nanocellulose-based encapsulation systems. Nanocarrier encapsulation has demonstrated improvements in survival rates of probiotic strains such as *Bifidobacterium longum*, increasing viability from 25% to 68% after 4 h simulated gastric exposure, when embedded in chitosan-coated lipid nanoparticles [37].

The synergistic effect of prebiotics and cellulose nanostructures can modulate the hydration properties, swelling behaviour, and mechanical resistance of the encapsulation system. Additionally, the bioactive compounds in avocado extracts, such as flavonoids and phenolic acids, exhibit antimicrobial and antioxidant properties, further safeguarding probiotics from oxidative stress and microbial contamination [38]. This innovative approach paves the way for the development of next-generation probiotic delivery systems with enhanced efficiency and functionality [39].

Cellulose matrices refer to structured scaffolds or encapsulating materials made from natural or modified cellulose. These matrices act as carriers that physically protect probiotics during processing and gastrointestinal transit by providing mechanical stability and controlling release kinetics. Common forms include microcrystalline cellulose, bacterial cellulose, and nanocellulose gels or films, each offering biocompatibility, non-toxicity, and enhanced structural integrity for encapsulation purposes [40].

### Approaches and limitations

Several approaches will be expressed in nanomaterials used in the preparation of pharmaceuticals and nutraceuticals, as the utilization of the nanomaterials will be encapsulated for protection and controlled release. It also enhances solubility. To achieve the specific delivery of drug formulations [42]. The increase in the absorption of the formulations is used as a bioavailability enhancer. Nanotechnology enables the controlled release, combination therapy, intracellular delivery, and protection of sensitive nutraceutical compounds from degradation. It enhances their stability and biocompatibility to ensure safe and effective delivery, including facilitating their transport across the blood-brain barrier [43].

**Table 1: A comparative overview of commonly employed probiotic stabilization methods is presented below**

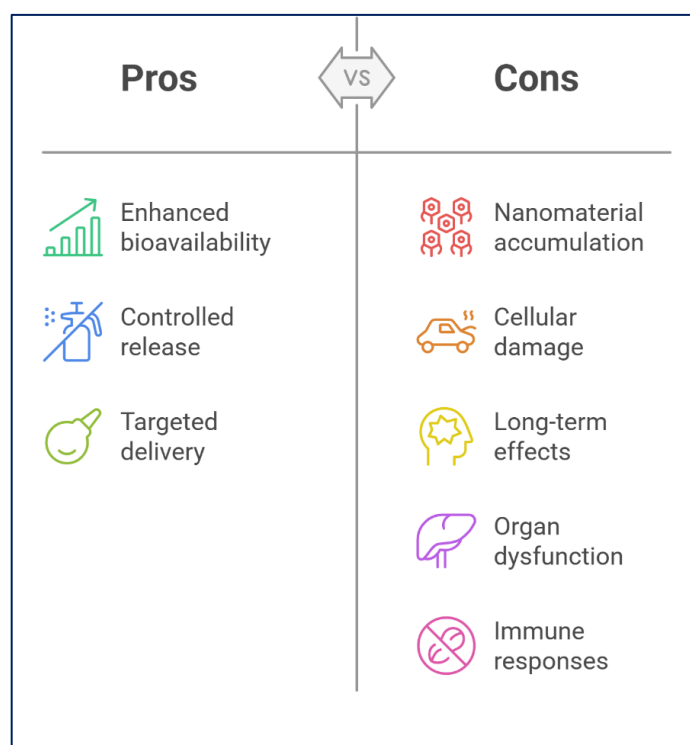
Method	Key materials	Viability post-GI transit	Thermal resistance	Limitations	Reference
Freeze-drying	Cryoprotectants (e. g., skim milk, sucrose)	~40–70%	Low	Requires refrigeration, low resistance to GI acids	[41]
Spray-drying	Maltodextrin, whey protein	~30–60%	Moderate	High thermal stress, oxygen exposure	
Alginate encapsulation	Sodium alginate, CaCl <sub>2</sub>	~60–80%	Moderate	Poor mechanical strength, slow release	
Nanocellulose-based delivery	Bacterial cellulose, plant nanofibers	~85–98% (up to 120 °C)	High	Still under regulatory evaluation	
Avocado	Avocado	~99.8% in SGF; stable over 40 d at 4 °C	Excellent	Requires optimization of extraction/processing	
Prebiotic+Nano-fiber Systems	pectin/starch+cellulose				

**Table 2: Functional roles of nanotechnology in nutraceutical delivery**

Section	Description	Examples/Data	References
Controlled Release and Protection	Nanocarriers protect nutraceuticals from heat, light, pH, and enzymatic degradation, enabling sustained and targeted release.	Curcumin-loaded PLGA nanoparticles showed >70% protection and 24h sustained release.	[44]
Combination Therapy	Co-delivery of multiple bioactives to enhance synergistic therapeutic outcomes.	Curcumin+piperine delivery improved bioavailability by 2000%.	
Intracellular Targeting	Functionalized nanocarriers target specific cells or receptors to enhance uptake and action inside cells.	Gold nanoparticles delivered polyphenols, boosting antioxidant activity 3.2×.	
BBB Penetration	Nanoparticles facilitate crossing of the blood-brain barrier for neuroprotective nutraceutical delivery.	Chitosan-coated NPs delivered resveratrol with 4× higher brain accumulation.	
Biocompatibility and Safety	Use of natural and biodegradable polymers ensures low toxicity and high safety for long-term use.	Chitosan, PLGA, and alginate-based systems showed no cytotoxicity in in vivo studies.	

The multiple challenges or limitations in these cases like the potential cytotoxicity of certain nanomaterials. In most cases, there is a huge gap in the regulatory approval process. There is always a question in those cases about the long-term effects of nanomaterials-based nutraceutical formulations, which also leads to affecting the overall nutraceutical formulation. One more consideration of the cost associated is higher, as potentially [45]. Enhancing the consistency, quality, and performance across different batches of

nanomaterials incorporating formulation is a tough or challenging job all the time. There are other considerations, are as ethical concerns, environmental impact, limited understanding, interdisciplinary collaboration, and biodegradability, that will have diluted this sector for a long time. Also, establishing a standardized method will always be a potential focus for scientists and researchers to improve the safety and efficacy of nanomaterial-based nutraceutical formulations [46].

**Fig. 3: A critical analysis of the functional benefits and limitations in the advancement of nanostructured nutraceutical delivery systems**

**Table 3: Comparative efficacy of avocado-derived prebiotics in nanocarrier systems for probiotic delivery**

S. No.	Prebiotic source	Nanocarrier type	Probiotic strain	Survival/Efficacy (%)	References
1	Avocado seed oil	Solid lipid nanoparticles	L. acidophilus	60–64%	[47-52]
2	Avocado fibers	Alginate-cellulose matrix	L. rhamnosus	68% survival @ pH 2	
3	Avocado oil+lecithin	Liposomes	B. breve	72% GI survival	
4	Avocado+MCC	Polymeric nanocarriers	L. casei	70% GI survival	
5	Avocado peel fibers	Nanocellulose hydrogel	L. plantarum	75% viability after 5 h	
6	Avocado polysaccharides	PLEVs (plant extracellular vesicles)	L. reuteri	78% cellular uptake	
7	Avocado oil residue	Lipid-based nanosuspension	B. lactis	65% GI survival	
8	Avocado skin extract	Chitosan nanoparticles	L. fermentum	62% acid tolerance	
9	Avocado+pectin	Alginate nanobeads	S. thermophilus	59% after 4h, pH 1.5	
10	Cold-pressed avocado fibers	Gelatin nanofibers	L. gasserii	73% encapsulation efficiency	
11	Avocado mash residues	Starch nanocarriers	L. delbrueckii	66% viability	
12	Avocado peel pectin	Polysaccharide nanogels	L. bulgaricus	68% survival after lyophilization	
13	Avocado pulp extract	β-glucan-based nanocapsules	L. johnsonii	64% stability @ RT	
14	Avocado seed residue	Biopolymer nanocoatings	B. coagulans	61% under storage	
15	Avocado pit hydrolysates	Zein nanoparticles	L. casei Shirota	69% survival at pH 3	
16	Avocado oil	Nanoemulsions	L. salivarius	67% bile resistance	

### Clinical trials and toxicities reported

The essence of nanostructured materials in nutraceuticals has shown promising potential for perfecting their delivery and efficacy [53]. Still, it's essential to consider the implicit toxin associated with these nanomaterials to ensure public health and safety. While nanostructured materials offer multitudinous advantages, there

have been reports of certain toxins. Nanomaterial accumulation. Some studies have indicated that certain nanomaterials used in nutraceuticals can accumulate in organs and tissues, raising concerns about the potential long-term effects. For example, nanoparticles may accumulate in the liver, lungs, or feathers, potentially leading to adverse health effects if not duly cleared from the body [54].

**Table 4: Exploring controlled release techniques in avocado-derived nutraceuticals**

Technique	Material used	Encapsulation method	Particle size (nm)	Release mechanism	Targeted site	Release duration	Stability improvement	Bioavailability impact	Potential applications	Ref.
Liposomal Encapsulation	Phospholipids and Avocado Oil	Thin-film hydration	100-200	pH-dependent release	Small Intestine	8-12 h	Enhanced oxidative stability	Improved absorption	Functional beverages	[55, 56]
Nanoemulsions Technology	Avocado Seed Extract	Ultrasonication	50-150	Diffusion-based	Gut	4-8 h	Improved thermal stability	Increased bioavailability	Yogurt, smoothies	
Alginate Bead Encapsulation	Alginate-Cellulose	Ionotropic gelation	200-500	Enzyme-triggered release	Microbiota Colon	10-14 h	Enhanced moisture retention	Controlled probiotic delivery	Probiotic capsules	
Solid Lipid Nanoparticles	Avocado Oil and Lipids	Hot homogenization	100-300	Temperature-sensitive	Stomach	6-10 h	Improved antioxidant retention	Prolonged release in digestion	Functional desserts	
Chitosan Nanoparticles	Chitosan and Avocado Fiber	Ionic gelation	150-400	Electrostatic attraction	Small Intestine	10-15 h	Enhanced acid resistance	Improved nutrient stability	Nutrient-rich powders	
Polymeric Nanoparticles	Avocado-Cellulose Matrix	Solvent evaporation	100-250	Gradual diffusion	Colon	12-16 h	Strong pH resistance	Enhanced gut colonization	Fiber-enriched supplements	
Nano encapsulation (Protein)	Whey Protein and Avocado	Spray drying	200-400	Enzyme-activated release	Small Intestine	8-12 h	Enhanced protein stability	Optimal nutrient absorption	High-protein snacks	
Nanostructured Lipid Carriers	Avocado Extracts	Cold homogenization	50-200	Lipid-melting mechanism	Gut Lining	8-12 h	Reduced oxidative degradation	Increased probiotic viability	Gut-supportive drinks	
Nano-Coated Capsules	Avocado Polysaccharides	Electrospinning	150-350	pH-sensitive coating	Colon	10-14 h	Improved probiotic protection	Gradual probiotic release	Medicinal probiotic tablets	
Microencapsulation	Avocado Fiber and Starch	Spray cooling	300-600	Controlled diffusion	Stomach and Colon	12-18 h	Enhanced water retention	Delayed nutrient breakdown	Nutraceutical powders	

**Table 5: The different characterization techniques for nanotechnology used for nutraceuticals**

Characterization techniques	Explained overview	References
Electron microscopy-scanning electron microscopy (SEM)	Provides high-resolution imaging of the face morphology and flyspeck size of nanostructured materials, and the Transmission Electron Microscopy (TEM) enables detailed imaging of the internal structure, size, and shape of nanoparticles and nanoscale structures.	[57-60]
X-ray Diffraction(XRD)	Determines the crystal-clear structure and crystalline phases of nanostructured materials, furnishing insight into their composition and stability.	
Dynamic light scattering (DLS)	Measures the flyspeck size distribution and size-dependent parcels of nanoparticles in the result, aiding in understanding their stability and behaviour.	
Fourier transform infrared spectroscopy(FTIR)	Analyses the chemical bonds and functional groups present in nanostructured accoutrements, easing identification and characterization of their molecular composition.	
Ultraviolet-visible spectroscopy(UV-VIS)	Assesses the optic parcels and immersion characteristics of nanostructured materials, aiding in their quantification and stability evaluation.	
Zeta Potential Analysis	Determines the face charge and stability of nanoparticles and colloidal systems, furnishing insight into their dissipation and implicit relations.	
Thermogravimetric analysis(TGA)	Evaluates the thermal stability, corrosion, and weight loss of nanostructured materials under controlled temperature conditions.	
Differential scanning calorimetry(DSC)	Measures the changes in heat inflow associated with phase transitions, melting, and crystallization behaviour of nanostructured materials, a benefit in their thermal analysis.	
Atomic force microscopy(AFM)	Provides high-resolution imaging and topographical mapping of nanostructured materials, allowing for the characterization of face roughness and morphology.	
Raman spectroscopy	Analyses the vibrational modes of nanostructured materials, enabling identification of molecular structures and chemical composition.	



The underlying mechanisms of nanostructured materials in nutraceuticals involve various factors that contribute to their enhanced efficacy and delivery. These mechanisms are driven by the unique properties and behaviour of nanomaterials at the nanoscale. There are some crucial underlying mechanisms. Increased surface area, nanostructured materials retain a high surface area-to-volume ratio compared to their bulk counterparts. This increased surface area allows for less commerce with target spots in the body, easing bettered immersion and bioavailability of bioactive composites [61]. Enhanced solubility. Numerous bioactive composites used in nutraceuticals have poor solubility in water or natural fluids. Nanostructured carriers, similar to nanoparticles or liposomes, can enhance the solubility of these composites by encapsulating them within their structures, adding their dispersibility and ensuring their emptiness for total involvement [62]. Controlled release, nanostructured materials can offer controlled release of bioactive molecules. By recapitulating the molecule within nanocarriers, release can be regulated based on factors such as pH, temperature, or enzymatic activity at the target point. This controlled release allows for sustained and slow remedial effects [63]. Targeted delivery, nanostructured carriers can be designed for targeted delivery of bioactive molecules to specific tissues, cells, or organs. This can be achieved through surface variations, where ligands or targeting moieties are attached to the nanocarrier, easing specific interactions with target cells or receptors. Targeted delivery enhances the localization and concentration of bioactive compounds at specific sites, maximizing their therapeutic effects and minimizing off-target effects [64-66]. Protection and stability, Nanostructured materials give protection to preserved bioactive molecules, shielding them from degradation, enzymatic exertion, or harsh environmental conditions. This protection preserves the stability and integrity of the composites, ensuring their optimal functionality and bioactivity. Enhanced permeation and cellular uptake, nanostructured materials can overcome barriers to cellular uptake and saturation, particularly in natural membranes. Their small size and unique surface properties smooth relations with cell membranes, promoting effective uptake and intracellular delivery of bioactive composites [67]. Synergistic effects, in some cases, the combination of nanostructured materials with bioactive composites may lead to synergistic effects. The nanomaterials can enhance the exertion or effectiveness of the bioactive composites, potentially through bettered solubility, stability, or cellular uptake, leading to enhanced remedial issues. These foundational mechanisms accentuate the eventuality of nanostructured materials in nutraceuticals to ameliorate the delivery, stability, and bioavailability of bioactive composites. Understanding these mechanisms is pivotal for the rational design and development of nano-grounded nutraceutical formulations or products, optimizing their remedial efficacy and ensuring safe and effective use for the human body [68, 69].

### Different synergistic combinations

The synergistic combination of avocado-derived prebiotics, cellulose matrices, and advanced nanotechnology offers a groundbreaking approach to enhancing probiotic stability, bioavailability, and targeted delivery [70, 71]. Cellulose matrices act as biodegradable carriers, protecting probiotics from gastric acid, bile salts, and oxidative stress, while avocado-derived prebiotic fibers (oligosaccharides and polysaccharides) improve water retention, creating an optimal environment for probiotic survival. Additionally, avocado polyphenols provide antioxidant protection, preventing oxidative damage and enhancing gut microbiota diversity. When combined with nanoencapsulation techniques such as liposomal carriers, alginate-cellulose hybrids, and protein-based nanocarriers, probiotics gain enhanced resistance to environmental factors and controlled release mechanisms for effective gut colonization. Furthermore, avocado seed extracts rich in tannins and phenolic compounds contribute to gut barrier protection, while nanoemulsions ensure uniform probiotic dispersal. This multi-layered probiotic delivery system not only extends shelf-life but also optimizes metabolic activity and therapeutic efficacy, paving the way for next-generation probiotic formulations with enhanced functional and health benefits [72].

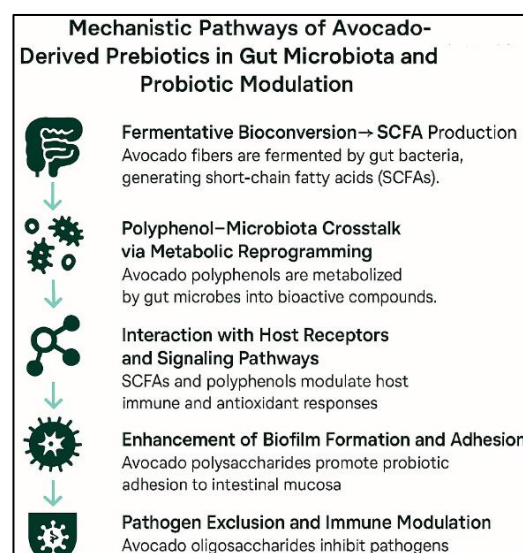


Fig. 4: Mechanistic pathways by which avocado-derived prebiotics modulate gut microbiota and enhance probiotic viability

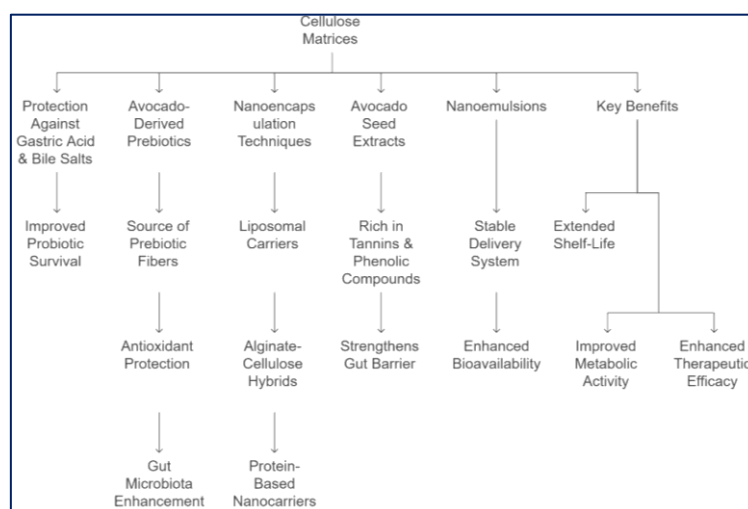
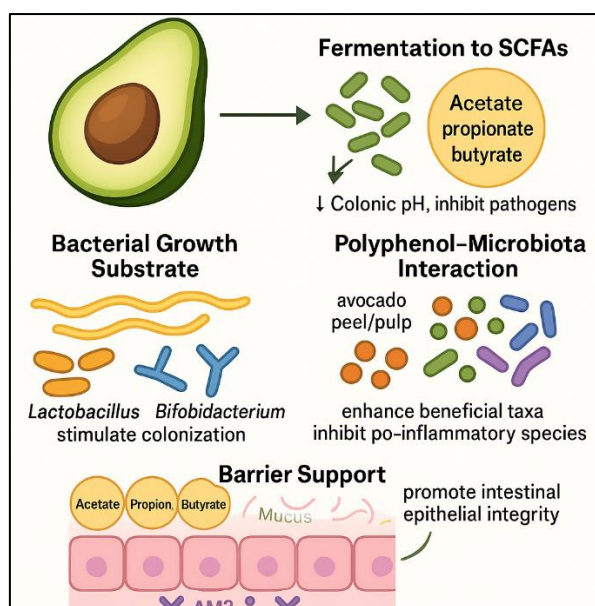


Fig. 5: Comprehensive framework for improving probiotic viability using cellulose matrices and supportive biotechnological approaches



**Fig. 6: Mechanistic explanation of how avocado-derived prebiotics support probiotic activity**

### Microbiota SCFA barrier link

Avocados have emerged as a functional food with significant potential to modulate gut health through both prebiotic and probiotic pathways. Rich in dietary fiber, avocados serve as an effective substrate for beneficial gut bacteria such as *Lactobacillus* and *Bifidobacterium*, promoting their colonization and activity within the gastrointestinal tract. The fermentation of these fibers by the resident microbiota leads to the production of SCFAs, including acetate, propionate, and butyrate. These SCFAs are critical for lowering colonic pH, thereby inhibiting the growth of harmful pathogens, while simultaneously acting as signalling molecules and energy sources for colon epithelial cells [73, 74].

Beyond fiber, the polyphenolic compounds present in avocado peel and pulp further contribute to gut microbiota modulation. These polyphenols enhance the abundance of beneficial microbial taxa and suppress pro-inflammatory species, promoting a favourable microbial balance. Moreover, the SCFAs, particularly butyrate, play a key role in supporting the integrity of the intestinal barrier. They stimulate the production of mucins and antimicrobial peptides, reinforce tight junctions, and help maintain epithelial health, thus preventing microbial translocation and systemic inflammation [75, 76].

Avocados provide a synergistic combination of fibers and bioactive compounds that support the growth of beneficial microbes, facilitate the production of health-promoting SCFAs, modulate inflammatory responses, and reinforce intestinal barrier function. These findings underscore the promising role of avocados in dietary strategies aimed at enhancing gut health and overall well-being [77-79].

### Approaches of targeted delivery

Targeted delivery refers to the precise and precise delivery of remedial agents or bioactive molecules to specific cells, tissues, or organs in the body. In the environment of nutraceuticals, targeted delivery aims to optimize the distribution and attention of bioactive molecules to the required area, maximizing their remedial efficacy while minimizing implicit side effects. Targeted delivery enables nutraceuticals to exert their effects directly at the intended point of action of the human body organ [80]. By directing bioactive molecules to specific cells, tissues, or organs, the desired remedial issues can be achieved more effectively. For illustration, targeting the gastrointestinal tract can enhance the absorption of nutrients or promote gut health, while targeting specific cells can modulate cellular functions or signalling pathways. Targeted delivery systems can ameliorate the bioavailability of bioactive molecules by overcoming barriers to immersion, metabolism, or elimination [81].

By widely delivering the molecules to specific spots, their attention at the target point can be increased, using effective uptake and application. This enhances the remedial effects and reduces the amount of the bioactive composites. By directing nutraceuticals specifically to the target point, the exposure of on-target tissues or cells to the bioactive composites can be minimized [82, 83]. This targeted delivery reduces the threat of adverse responses and enhances the safety profile of the nutraceutical formulations. Targeted delivery systems can cover bioactive molecules from declination or unseasonable release. Targeted delivery of probiotics is essential for maximizing therapeutic efficacy and gut colonization. Conventional probiotics often suffer from low survival rates due to premature degradation in the acidic stomach environment. However, cellulose matrices act as intelligent carriers, enabling probiotics to bypass gastric degradation and reach the intestine intact. When combined with avocado-derived prebiotics, these matrices provide an optimal environment for probiotic adherence, colonization, and metabolic activity in the gut microbiome [84]. The structural properties of cellulose-based nanocarriers allow for pH-responsive release, ensuring probiotics are released only upon reaching the intestinal lumen, where they exert their beneficial effects. Additionally, avocado-derived prebiotics serve as a selective energy source, fostering the growth of probiotics and enhancing their survival in the gut. This targeted delivery system could revolutionize probiotic-based nutraceuticals, ensuring higher efficacy and prolonged retention in the gastrointestinal tract [85, 86].

### Stability and shelf-life enhancement

Stability and shelf-life enhancement are basic considerations when joining nanostructured materials in nutraceutical formulations [87]. There are how nanostructured materials can contribute to soundness improvement and amplified stability of nutraceutical formulations, nanostructured materials can grant protection to bioactive molecules against degradation [88]. Epitome of the molecules inside nanostructured carriers, comparable to nanoparticles or liposomes, shields them from outside components, such as oxygen, humidity, light, or enzymatic effort, which can corrupt or modify their chemical structure. This assurance enhances the intelligence and stability of the bioactive compounds, prolonging their effectiveness over an extended duration. Nanostructured materials can help diminish oxidation of bioactive molecules, particularly those delicate to oxidative degradation. One of the critical challenges in probiotic formulations is maintaining viability during storage, transportation, and digestion [89, 90]. Probiotics are sensitive to environmental conditions, including temperature, humidity, pH fluctuations, and oxidative stress, which can

significantly reduce their potency. The integration of avocado-derived prebiotics with cellulose matrices offers a two-fold advantage: (1) stabilizing probiotic cells during storage and (2) ensuring their protection during gastrointestinal transit. The prebiotic fibers from avocado function as natural cryoprotectants and moisture regulators, preventing dehydration and improving the resilience of probiotic strains against temperature variations. Furthermore, cellulose-based encapsulation provides a physical barrier against external aggressors while facilitating a gradual and controlled release of probiotics in the gut. This novel formulation can significantly extend the shelf-life of probiotic-based nutraceuticals, making them more commercially viable and effective in real-world applications [91].

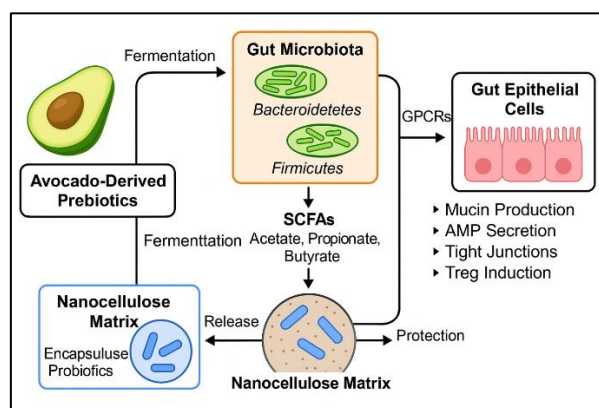
Avocado is rich in polysaccharides such as pectin, cellulose, and oligosaccharides, as well as polyphenolic compounds. These constituents act as prebiotic substrates that: a) Protect probiotic viability under oxidative and acidic conditions. b) Enhance colonic fermentation, increasing probiotic colonization potential. c) Reduce moisture sensitivity when incorporated into matrices, improving powder hygroscopicity and extending shelf life during storage. Nanocarriers serve as physical and chemical protectants for probiotics by a) Shielding them from gastric acid and bile salts during GI transit. b) Creating hydrated protective matrices that prevent desiccation and oxygen exposure. c) Acting as barriers to water activity ( $a_w$ ), which is critical in preserving probiotic cultures during storage [92].

**Table 6: Various nanocarrier systems applied in avocado-derived cellulose nanotechnology**

Types of nanocarriers used:		
Nanocarrier	Functionality	References
Alginate-cellulose beads	pH-responsive swelling, strong gelation, and moisture resistance	[93]
Lipid nanoparticles	Protection against thermal and oxidative stress	
Chitosan-coated liposomes	Mucoadhesive delivery and bile salt resistance	
Zein nanoparticles	Hydrophobic matrix, suitable for dry probiotic formulations	

Stability in the Simulated Gastric and Intestinal Fluid (SGF/SIF) models has been significantly improved using avocado-derived matrices. Improved survival from 30% → 65–78% in acidic pH (pH 1.5–3.0). Bile salt resistance enhanced by formation of hydrophobic outer shells from avocado lipids. Freeze-drying or spray-drying

probiotic cultures with avocado fibre matrices has shown improved rehydration capacity and retention of CFU over time. Nanocarrier coatings also serve as antioxidant and anti-humidity shields, preventing degradation caused by light, temperature, and air exposure [94].

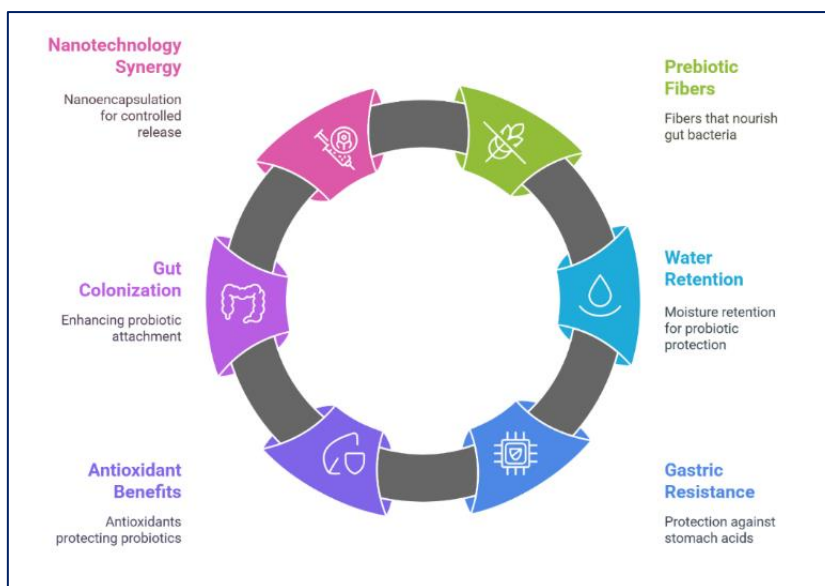


**Fig. 7: Biochemical and technological pathways enhancing probiotic stability using avocado-derived prebiotics and nanocellulose system**

**Table 7: Overview of biochemical mechanisms and technological strategies involving avocado-derived prebiotics and nanocellulose matrices for enhancing probiotic stability**

Aspect	Details	References
Avocado-derived prebiotics	Avocado peels and seeds contain 3–26% dietary fibre (mainly insoluble cellulose/hemicellulose) and polyphenols (3–5 mg GAE/g). These components resist digestion in the upper gut and are fermented by colonic microbiota.	[95-97]
Gut Microbiota	Bacteroidetes (e. g., <i>Bacteroides</i> spp.) hydrolyse complex polysaccharides into sugars and lactate, producing acetate and propionate. Firmicutes (e. g., <i>Faecal bacterium prausnitzii</i> ) further metabolize these products to produce butyrate.	
Fermentation	The fermentation process yields Short-Chain Fatty Acids (SCFAs) with a typical colonic profile of 60:20:20 for acetate: propionate: butyrate. Avocado fiber consumption has been shown to increase fecal butyrate levels significantly.	
SCFA Production	SCFAs bind to G-protein-coupled receptors (GPR41/FFAR3, GPR43/FFAR2, GPR109A) on intestinal epithelial and immune cells, activating pathways that modulate metabolism and inflammation.	
SCFA Signalling Pathways	Activation of GPR41/43 by SCFAs leads to the production of chemokines and cytokines, enhancing mucin production, tight junction integrity, and regulatory T-cell responses, thereby maintaining intestinal homeostasis.	
Immune Modulation	Prebiotics lower local pH, inhibiting pathogens and favoring acid-tolerant probiotics. They also physically protect probiotics, buffer gastric acid, and bind bile acids, reducing their detrimental effects on probiotic membranes.	
Probiotic Survival Enhancement	Nanocellulose-based matrices (e. g., cellulose nanofibrils, methylcellulose) encapsulate probiotics, enhancing their stability and survival through the gastrointestinal tract. These matrices are pH-responsive, releasing probiotics in the colon.	
Nanocellulose Matrices	Studies have shown that nanocellulose-alginate capsules significantly improve probiotic survival under gastric conditions compared to alginate alone, with higher viability and controlled release in the intestinal environment.	
Comparative Efficacy		





**Fig. 8: Avocado-derived nutrients and their impact on probiotic growth and function**

### Overview of preclinical and clinical studies

Preclinical and clinical studies play a vital part in the assessment of nanostructured materials in nutraceuticals, giving valuable insights into their safety, efficacy, and potential benefits. These considerations encompass comprehensive testing and assessment in both laboratory and human settings, gathering scientific evidence to support decision-making regarding the incorporation of nanostructured materials in nutraceutical formulations. Here, the centrality of preclinical and clinical considerations within the assessment of nanostructured materials in nutraceuticals [98]. Preclinical studies include laboratory-based tests conducted on cells, tissues, or in silico models, sometimes recently testing in people. These studies evaluate the security, bioavailability, pharmacokinetics, and potential toxicities of nanostructured materials. To evaluate the possible adverse effects and toxicities of nanostructured materials, comprehensive toxicity studies, repeated-dose toxicity assessments, genotoxicity evaluations, and investigations of potential organ-specific or systemic toxicities are conducted. Additionally, research on the Absorption, Distribution, Metabolism, and Excretion (ADME) of nanostructured materials helps determine their bioavailability and systemic exposure. These studies are essential for gaining insights into the fate and behaviours of these materials within the body [99]. To evaluate the helpful adequacy of nutraceutical details containing nanostructured materials. This includes assessing their ability to convey bioactive compounds, target cells or tissues, and illustrate the required organic or physiological impacts. Preclinical considers help within the optimization of definition parameters, such as molecule estimate, surface alterations, and discharge energy, to improve the execution and adequacy of nanostructured materials in nutraceuticals. Clinical considerations include the assessment of nanostructured materials in human subjects. They are conducted in a few stages, including Stage I, II, and III trials, to survey security, viability, and tolerability [100]. Clinical trials evaluate the safety profile and tolerability of nanostructured materials in human subjects. This involves monitoring adverse events, potential side effects, and allergic reactions, ensuring the overall safety of the nutraceutical formulations. Clinical considers examine the assimilation, conveyance, digestion system, and disposal of nanostructured materials in people, giving insights into their pharmacokinetics and bioavailability profiles [101]. Clinical trials assess the restorative adequacy of nutraceutical details containing nanostructured materials. They evaluate the craved wellbeing results, adequacy endpoints, and clinical endpoints, illustrating the adequacy of the nanostructured materials in improving wellbeing conditions. Clinical studies about offering assistance decide the suitable dose

regimen for nutraceutical details, considering factors such as viability, safety, and persistent population variability. These studies guide the foundation of dosing suggestions for clinical utilize [102].

A few clinical studies about centre on long-term security checking, surveying the potential antagonistic impacts related to drawn exposure to nanostructured materials in nutraceuticals. Through conducting preclinical and clinical studies, researchers and regulatory experts can gather scientific evidence regarding the safety, effectiveness, and potential benefits of nanostructured materials in nutraceuticals. These studies provide a foundation for evidence-based decision-making, regulatory approvals, and recommendations for the use of nanostructured materials in nutraceutical formulations, ensuring their safe and effective application in human health [103].

### Applications for emerging technology in nutraceuticals

Relevant examination and exercises offer important bits of knowledge into the application of nanostructured materials in useful foods, highlighting their benefits and complexities. These studies demonstrate the promise of nanoemulsions in enhancing the bioavailability of bioactive compounds, like curcumin, surpassing traditional formulations. Nanoparticles act as targeted delivery systems, allowing precise delivery of compounds to specific cells or organs. Controlled-release nanofibers have been shown to guarantee in amplification of the discharge of bioactive over time for applications like wound healing and sedative delivery. Nanoencapsulation improves stability and shelf life by securing delicate compounds from corruption and intelligent [108, 109].

The integration of nanotechnology, prebiotics, and probiotics is at the forefront of nutraceutical innovations. Recent advancements in probiotic-prebiotic nanocomposites highlight their potential in personalized nutrition, gut health optimization, and disease prevention [110]. Case studies have demonstrated that encapsulating probiotics in prebiotic-functionalized cellulose nanofibers significantly improves viability, enhances metabolic activity, and extends probiotic shelf-life in functional food applications. For instance, research on plant-based prebiotics combined with cellulose nanofibers has shown increased resistance to gastric fluids and improved probiotic adherence to intestinal mucosa. By incorporating avocado-derived prebiotics, the field can advance eco-friendly, plant-based probiotic stabilizers that align with consumer demand for sustainable and natural nutraceutical solutions. Moreover, the use of biodegradable cellulose matrices ensures minimal environmental impact while enhancing bioavailability, retention time, and probiotic functionality [111].

**Table 8: Comprehensive overview of next-generation probiotic formulations using avocado-derived prebiotics and nanotechnology**

Component	Source	Bioactive compounds	Function	Technique /Technology	Encapsulation material	Release mechanism	Stability improvements	Health benefits	Potential applications	References
Avocado-Derived Prebiotics	Avocado pulp and peel	Oligosaccharides, polysaccharides	Supports gut microbiota growth and improves water retention	Fiber extraction and purification	Alginate-cellulose hybrids	Controlled release in the intestinal environment	Enhanced resistance to gastric acid and bile salts	Improved digestion and gut microbiota balance	Functional foods, probiotic beverages, dietary supplements	[104-107]
Avocado Polyphenols	Avocado pulp and seed	Phenolic compounds, flavonoids	Antioxidant protection against oxidative stress	Solvent extraction and encapsulation	Polymeric nanoparticles	Gradual release in the gut	Prevents oxidative damage, extends probiotic viability	Reduced inflammation, improved immune response	Anti-inflammatory supplements, functional teas	
Cellulose Matrices	Plant-derived cellulose	Microcrystalline cellulose (MCC)	Provides a biodegradable protective barrier for probiotics	Microencapsulation via spray-drying and freeze-drying	Biopolymer films	Sustained release in the intestinal tract	Improved probiotic survival in harsh conditions	Strengthened gut barrier, improved nutrient absorption	Yogurt, fermented foods, dietary capsules	
Nanoencapsulation Techniques	Various nanocarriers	Lipids, proteins, and polysaccharides	Controlled release and improved resistance to stress	Liposomal carriers, nanoemulsions, protein-based carriers	Chitosan-alginate nanoparticles	Timed release triggered by pH levels	Enhanced protection from gastric acid and bile salts	Extended probiotic viability, optimized therapeutic effects	Probiotic capsules, functional beverages	
Avocado Seed Extracts	Avocado seed	Tannins, phenolic acids	Gut barrier protection and anti-inflammatory properties	Tannin extraction and nanoparticle formulation	Nano-coating polymers	Gradual release for prolonged activity	Strengthens the intestinal lining, reduces inflammation	Enhanced gut health, reduced risk of intestinal disorders	Digestive enzyme boosters, anti-inflammatory formulations	
Nanoemulsions	Avocado oil	Fatty acids, bioactive lipids	Uniformly disperses probiotics in the digestive tract	Ultrasonic emulsification, microfluidization techniques	Oil-in-water emulsions	Rapid dispersion in gut fluids	Enhanced bioavailability, stable dispersion in the GI tract	Enhanced nutrient absorption, improved probiotic colonization	Probiotic drinks, nutraceutical formulations	
Probiotic Strains	Bifidobacteria, Lactobacillus	Lactic acid bacteria (LAB)	Strengthens immunity and balances gut flora	Encapsulation with cellulose and avocado fibers	Alginate-chitosan beads	Targeted release at specific gut regions	Prolonged viability under environmental stress	Improved digestion, immune support, and prevention of infections	Functional foods, capsules, yogurt, and kefir	
Carrier Systems	Natural biopolymers	Alginate, chitosan, gelatin	Form protective matrices for probiotics	Spray-drying, coacervation, and emulsion techniques	Hybrid biopolymer structures	Controlled release with pH or enzyme triggers	Improved resistance to temperature and mechanical stress	Prolonged shelf-life, improved probiotic efficacy	Functional powders, dietary supplements, fortified foods	
Probiotic Metabolites	Fermented avocado extract	Organic acids, short-chain fatty acids	Supports gut microbiota growth and enhances gut lining	Biofermentation techniques	Polysaccharide-encapsulated matrices	Slow-release mechanisms based on microbial activity	Enhanced probiotic metabolic activity and effectiveness	Improved gut flora balance, reduced risk of gastrointestinal issues	Fermented foods, kombucha, probiotic shots	
Functional Additives	Natural antioxidants	Vitamin E, C, and beta-carotene	Prevents oxidative degradation of probiotics	Blending with polymer-based matrices	Protein-polysaccharide blends	Antioxidant-protected gradual release	Enhances stability in heat, light, and oxygen exposure	Improved probiotic survival, better therapeutic efficiency	Enriched capsules, dietary blends, probiotic-enriched foods	

### Recent advances and developments in nanotechnology

Recent advances in nanotechnology have significantly transformed the nutraceutical industry, particularly in enhancing the stability, bioavailability, and targeted delivery of probiotics and prebiotics. The integration of nanostructured cellulose matrices with avocado-derived prebiotics has emerged as a promising approach to improving probiotic survival and gut microbiome modulation. Innovations such as nanoencapsulation, nanoemulsions, and lipid-based nanocarriers

ensure that probiotics are protected from harsh gastric conditions and released in a controlled manner at the desired site in the intestine [112]. Additionally, polyphenol-functionalized nanocarriers derived from avocado extracts enhance antioxidant stability, further safeguarding probiotics from oxidative stress. Breakthroughs in nanofiber technology now allow for personalized probiotic formulations, designed to meet specific microbiome profiles. Moreover, bioengineered nanoparticles with pH-sensitive coatings are being developed to enable targeted probiotic delivery, ensuring

maximal efficacy. As research continues, the combination of nanotechnology, artificial intelligence, and functional food innovations is expected to revolutionize precision nutrition, paving the way for next-generation nutraceuticals with enhanced health benefits [113].

### Growing interest in precision nutrition

The growing interest in precision nutrition is revolutionizing the nutraceutical industry by enabling personalized dietary interventions based on an individual's genetics, microbiome, metabolism, and lifestyle. The integration of avocado-derived prebiotics, probiotics, and nanotechnology aligns perfectly with this shift, offering customized gut health solutions that cater to specific nutritional deficiencies and microbiota imbalances [114]. Precision nutrition leverages advanced omics technologies (genomics, metabolomics, and microbiomics) to tailor functional foods and supplements that enhance probiotic efficacy and gut microbiome composition. By incorporating prebiotic-functionalized cellulose matrices, probiotic formulations can be designed to target specific populations, such as individuals with digestive disorders, immune deficiencies, or metabolic syndromes [115, 116]. Moreover, nanotechnology-driven smart delivery systems ensure controlled and site-specific probiotic release, maximizing absorption and therapeutic benefits. As consumer demand for personalized wellness solutions grows, the development of precision-engineered probiotic-prebiotic combinations using avocado bioactive and cellulose-based carriers is set to transform the future of gut health and functional nutrition, making tailor-made nutraceuticals more accessible and effective [117, 118].

### Future directions and emerging technologies challenges

Avocado-derived prebiotics, including pectin, resistant starch, and polyphenols, exhibit promising potential to enhance probiotic viability and gut health, especially when integrated with nanotechnology. The current review underscores their role in improving probiotic delivery, protection during gastrointestinal transit, and synergistic interactions that modulate gut microbiota composition. However, to fully harness these advantages, several areas require further exploration. More in-depth investigations into the exact biochemical and molecular pathways through which avocado-derived compounds enhance probiotic survival and gut colonization are needed. While *in vitro* data are promising, controlled animal studies and human clinical trials are essential to validate efficacy, dosage, and safety. Research must clarify optimal concentrations and combinations of avocado prebiotics and probiotics for different populations. Future studies should develop novel nanomaterials (e. g., cellulose nanofibers, chitosan composites) for co-delivery systems that enhance both prebiotic and probiotic effects. Integration into personalized nutraceutical products tailored for specific gut microbiota profiles or disease conditions could become a future application area. There are different potential industrial applications, like as the incorporation of avocado-derived prebiotics into yogurts, snack bars, or fermented drinks with encapsulated probiotics, and nanotechnology-assisted capsules with avocado-derived prebiotics and probiotics can improve shelf life and therapeutic effectiveness. Smart food packaging materials containing avocado-derived compounds may help preserve probiotic-rich foods during storage and application in animal feeds to improve gut health in livestock and pets.

In the USA, Europe, Japan, Health Canada, China, and India, several objections and unborn perspectives are shaping the geography of nutraceuticals, nanotechnology, precision nutrition, and substantiated health. Common objections carry regulatory complications, ensuring consumer security, and maintaining quality control norms amidst rapid-fire technological creations. Moreover, there is a need for increased public awareness and education regarding the benefits and risks associated with these emerging fields. Unborn perspectives rotate around the integration of artificial intelligence, engaging literacy, and data analytics to enable individualized health recommendations and interventions. Collaboration between academia, industry, and regulatory bodies will play a pivotal part in furthering invention and easing market access. Featuring sustainability and environmental considerations will also be vital, promoting eco-friendly practices and reducing the ecological

footprint of this diligence. Also, addressing ethical considerations, guarding consumer birthrights, and ensuring equal access to substantiated health results will be an ongoing process. Despite the criticisms, the convergence of technology, research, and regulatory developments presents promising opportunities for improving healthcare, enhancing disease prevention, and transforming the way individuals approach their health and well-being in these countries.

In terms of looking for the new product development might be looking forward this regulatory challenges, the companies should be focused on the direction of the avocado extracts, and nanocarriers must be thoroughly evaluated for GRAS (Generally Recognized as Safe) certification, especially for oral use. Regulatory frameworks regarding nano-encapsulation in foods vary widely across countries, with inconsistent safety thresholds. Clear guidelines are needed for labelling nano-enabled nutraceuticals to maintain consumer trust and transparency.

### LIMITATIONS

Despite the growing interest in this field, several limitations were observed across the studies reviewed, as most studies were *in vitro* or animal-based; few human clinical trials exist. Different probiotic strains, matrices, and avocado derivatives used across studies reduce comparability. Many studies lacked statistical power due to small cohorts. No standard methodology for evaluating prebiotic-probiotic interactions with avocado compounds. Particularly for nano-encapsulated systems, long-term consumption safety remains underexplored. While nanotechnology shows promise, the long-term biocompatibility and potential toxicity of nanocarriers like nanocellulose or liposomes remain under-investigated. Most articles do not provide dose-response relationships for avocado-derived components or their synergistic effects with probiotics. There is insufficient mechanistic insight into how specific avocado compounds modulate probiotic gene expression or gut epithelial interactions. The technological feasibility of using avocado-based nanomaterials at an industrial scale (cost, production time, reproducibility) is not yet established. Regulatory guidelines for combining plant-derived prebiotics and nanotechnology in food or supplement applications are still evolving and country-specific. Publication bias may exist, where studies showing no improvement in probiotic viability with avocado-based systems are underreported.

### CONCLUSION

The period of nanotechnology has opened new possibilities for perfecting nutraceutical delivery systems for the future. The use of nanostructured materials as carriers offers several advantages, including enhanced bioavailability, targeted delivery, stability improvement, and controlled release. The implicit operations of nanotechnology in nutraceuticals have been explored through useful approaches, such as nanoemulsions, nanoparticles, nanofibers, nanoencapsulation, nanosensors, and nanocomposites. These inventions hold significant promise for enhancing the efficacy, security, and consumer acceptance of nutraceutical products. Still, along with the tremendous openings, some objections need to be managed, involving regulatory frameworks, actions on public health, ethical considerations, and intellectual property rights. Regulatory bodies play an overcritical part in ensuring the security, quality, and efficacy of these nutraceuticals' formulations, while public perception and acceptance are essential for their success in the market. Likewise, environmental sustainability considerations are getting increasingly important in the evolution and commercialization of nanotechnology-grounded nutraceuticals. Looking ahead, the future of nutraceutical delivery linkages on continued research and development, global adaptation, personalized nutrition approaches, and the incorporation of emerging technologies. With ongoing innovations, collaboration, and responsible practices, the era of nanotechnology in advancing nutraceutical delivery shows tremendous potential to revolutionize the field of nutrition and well-being, leading to improved health outcomes and enhanced quality of life for people.

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## AUTHORS CONTRIBUTIONS

Jatin M: Conceptualization, data acquisition, analysis, interpretation, writing-original draft, Bishop Adhikari: Data alignments and design of the fig. and table of the paper, writing-original draft. Piyush Kumar: Helped to analyse and collect data for the paper. For R. Rajesh Kumar: Review and editing, Supervision, Critical evaluation, and Validation.

## CONFLICT OF INTERESTS

The author is reporting no conflict of interest.

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