



# **Gut Probiotics and Health of Dogs and Cats: Benefits, Applications, and Underlying Mechanisms**

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Abstract: Pets (mostly domestic dogs and cats) play an important role in the daily lives of humans and their health has attracted growing attention from pet owners. The intestinal microbiota, a complex microbial community with barrier-protective, nutritional, metabolic, and immunological functions, is integral to host health. Dysbiosis has been related to a variety of diseases in humans and animals. Probiotics have been used in functional foods and dietary supplements to modulate intestinal microbiota and promote host health, which has been introduced in pet dogs and cats in recent years. Various canine- and feline-derived probiotic strains have been isolated and characterized. The administration of probiotics has shown positive effects on the gut health and can alleviate some intestinal diseases and disorders in dogs and cats, although the underlying mechanisms are largely unresolved. In this review, we summarize the current knowledge on the benefits of probiotics and discuss their possible mechanisms in dogs and cats in order to provide new insights for the further development and application of probiotics in pets.

Keywords: probiotics; microbiome; intestinal health; companion animal; dog; cat



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## 1. Introduction

With the advancement of society and improvement in people's quality of life, an increasing number of households are raising pets or companion animalsthat are dominated by domestic dogs and cats. According to a report by the American Pet Products Association, 70% of American households owned pets by 2021 [1]. Evidence has indicated the benefits of pet ownership on mental and physical health [2–4]. It is a global trend that pet owners treat dogs and cats as family members. As dogs and cats have become an important part of human life, their health, especially intestinal health, is attracting increasing attention from pet owners [5].

The microbiota in the gastrointestinal tract (GIT) comprises a complex community of microorganisms, including bacteria, fungi, archaea, protists, and viruses [6]. The GIT microbiota contributes to maintaining intestinal and host homeostasis via multifaceted mechanisms, such as defense against intestinal pathogens, provision of nutrients, facilitation of nutrient digestion and absorption, improvement in barrier function, stimulation of intestinal development, and modulation of the immune system [7–9]. Bacterial metabolites, such as short-chain fatty acids (SCFAs), secondary bile acids, and tryptophan metabolites, are key mediators of host-microbiota interactions and play important roles in influencing host health [10,11]. SCFAs are products of bacterial fermentation of dietary fibers, mainly including acetate, propionate, and butyrate, which contribute to intestinal and host health by serving as energy substrates for colonic epithelial cells, maintenance of epithelial barrier integrity, regulation of energy metabolism, and anti-inflammatory effects [12,13]. Secondary bile acids derived from primary bile acids through deconjugation and dihydroxylation by intestinal bacteria possess metabolic and immunomodulatory properties [14,15]. However, it has been demonstrated that dysbiosis or dysfunction of the intestinal microbiota is associated with a variety of diseases, including intestinal disorders and diseases in humans

and farm animals [7,16,17]. Perturbation of intestinal microbiota, indicated by decreased microbiota diversity, imbalance of microbiota structure, or shifted metabolite profile, has also been observed in cats and dogs with gut-related disorders, such as chronic enteropathy, inflammatory bowel diseases (IBD), acute uncomplicated diarrhea, and acute hemorrhagic diarrhea syndrome [18,19].

The use of probiotics is a promising approach for manipulating the gut microbiota to promote host health [20–23]. Studies have shown that probiotics exert prophylactic or therapeutic activities in a variety of diseases such as IBD, antibiotic-associated diarrhea, colon cancer, allergy, type 2 diabetes, and atopic dermatitis in humans [20–22,24]. With positive effects such as improving growth performance, promoting intestinal health, and enhancing resistance against infections, probiotics have been explored as promising antibiotic alternatives in animal production [25,26]. An increase in nutrient digestion and absorption, improvement in intestinal morphology, optimization of the gut microbiota and suppressed inflammation have been linked with the health benefits of probiotics in farm animals [23]. Species and strains belonging to *Lactobacillus*, *Bifidobacterium*, *Enterococcus*, and *Saccharomyces* are commonly used probiotics in functional foods and dietary supplements [20,24,27].

Probiotics have been introduced to companion animals in recent years, and current evidence suggests the benefits of probiotics in health promotion and disease prevention in dogs and cats, although related research is not thorough relative to that in humans and livestock species [18,28,29]. Herein, we review the recent advances in the use of probiotics in pet dogs and cats to provide new insights into the further development and application of probiotics in pets.

### 2. The Microbiota in the GIT of Dogs and Cats

#### 2.1. The Normal GIT Microbiota of Dogs and Cats

Advancements in high-throughput microbiome techniques such as 16S rRNA gene sequencing, metagenomics, metatranscriptomics, metabolomics, culturomic, and multiomics make it possible to decipher the composition and function of the microbiota, discover microbiota-associated biomarkers, and reveal host-microbiota interactions [5,6,30–32]. An integrated use of culturomic and metagenomic analyses has identified 305 strains of commensal lactic acid bacteria in domestic dogs [33]. Characterization of the normal intestinal microbiota and the identification of microbiota-derived biomarkers associated with GIT diseases are important for understanding host-microbiota interactions and the development of probiotics in cats and dogs.

The canine and feline GIT microbiota remain largely unexplored. Available data suggest that the composition of the GIT microbiota is generally similar between dogs and cats [34,35] (Table 1). Microbial diversity and load increase along the GIT. The numbers of bacteria in the stomach, small intestine, colon, and feces of dogs and cats are generally  $10^4 - 10^5$  CFU/g,  $10^5 - 10^9$  CFU/g,  $10^9 - 10^{11}$  CFU/g, and  $10^8 - 10^{11}$  CFU/g, respectively [36]. In healthy dogs and cats, the stomach microbiota is dominated by *Helicobacter* and *Lacto*bacillus [28,36]. Firmicutes, Proteobacteria, Bacteroidetes, Fusobacteria, and Actinobacteria are the major phyla in the intestines of dogs and cats, with variations in abundance in each segment [19,28,36,37]. The fecal microbiota of healthy cats and dogs is dominated by Firmicutes, Proteobacteria, Bacteroidetes, Fusobacteria, and Actinobacteria [18,36]. At the genus level, Lactobacillus, Bifidobacterium, Enterococcus, Streptococcus, and Pediococcus are dominant in canine feces [38,39]. Moreover, a core microbiota composed of Firmicutes, Bacteroidetes, and Fusobacteria is present in the feces of healthy dogs, among which many bacteria belonging to Clostridia and Bacilli classes are SCFA producers (e.g., Faecalibac*terium* and *Blautia*) or lactic acid bacteria (e.g., *Lactobacillus*) with probiotic properties [18]. Prevotella and Bacteroides in the phylum Bacteroidetes were abundant in dog feces. Fusobac*terium* is associated with health and is, thus, a potential therapeutic target [18]. Similarly, another study showed that Fusobacterium, Prevotella 9 (a sub cluster of the Prevotellaceae family), and *Bacteroides* are the core intestinal microbiota in both dogs and cats [40]. Many

*Bifidobacterium* spp. and lactic acid-producing bacteria have been identified in canine feces [41]. *Lactobacillus* is widespread in the canine and feline intestines and can be the origin of probiotics [27,42].

	Stomach	Small Intestine	Colon	Feces	References
The number of total bacteria	$10^4$ – $10^5$ CFU/g (dogs and cats)	10 <sup>5</sup> –10 <sup>9</sup> CFU/g (dogs and cats)	$10^9$ – $10^{11}$ CFU/g (dogs and cats)	10 <sup>8</sup> –10 <sup>11</sup> CFU/g (dogs and cats)	[36]
Dominant bacteria	<i>Helicobacter</i> and <i>Lactobacillus</i> (dogs and cats)				[28,36]
		Protec Bacter Fusol Actine	nicutes obacteria roidetes oacteria obacteria and cats)		[19,28,36,37]
				Firmicutes, Proteobacteria, Bacteroidetes, Fusobacteria, and Actinobacteria (dogs and cats); a core microbiota composed of Firmicutes, Bacteroidetes, and Fusobacteria (dogs); Fusobacterium, Prevotella 9, and Bacteroides (dogs and cats)	[18,36,40]

Table 1. Distribution of bacterial microbiota along the GIT of dogs and cats.

Despite similarities between the canine and feline intestinal microbiota, differences have been observed between the two species. The diversity of fecal microbiota has been reported to be higher in cats [41,43]. A study showed that the abundances of *Enterococcus, Fusobacterium*, and *Megamonas* were higher in the feces of dogs, while multiple genera, including *Bifidobacterium, Faecalibacterium, Carnobacterium, Adlercreutzia, Alistipes, Collinsella, Coprococcus, Desulfovibrio, Oscillospira, Parabacteroides, Peptococcus, Peptostreptococcus, Ruminococcus, Slackia, and Sutterella, were more abundant in cats [43]. Another study also identified more abundant and prevalent <i>Bifidobacterium* in the guts of cats [40]. Dogs and cats have evolved into carnivores and have some anatomical and metabolic characteristics in common, feeding on high-protein diets and having relatively short GITs compared with humans and livestock species [44]. Domestic cats are obligate carnivores that rely on high-protein diets, whereas dogs are metabolically omnivorous and can digest, absorb, and metabolize higher amounts of carbohydrates [44]. The differences in the diet composition could be one of the factors that contribute to shaping divergent gut microbiota between the two species.

#### 2.2. Perturbation of Intestinal Microbiota of Dogs and Cats in Intestinal Diseases

An intact intestinal microbiota is essential for maintaining host homeostasis, whereas dysbiosis in the gut microbiota can be linked to a variety of diseases and disorders, such as IBD, diarrhea, obesity, autoimmune diseases, and cancers in humans [7,17]. Current evidence indicates that dysbiosis of the intestinal microbiota is related to various intestinal disorders in dogs and cats, such as chronic intestinal diseases, acute uncomplicated diarrhea, acute hemorrhagic diarrhea syndrome, and bowel cancer [5,35,45,46].

Altered gut microbiota structures and/or metabolite profiles were observed in dogs and cats with intestinal inflammation [35,37,45–47]. For example, decreased microbiota diversity, reduction in potentially beneficial bacteria (e.g., *Fusobacterium* and *Faecalibacterium* 

*praunitzii*), and overgrowth of *Clostridium perfringens* have been observed in dogs with chronic enteropathy [10,18]. Reductions in *Ruminococcaceae, Blautia, Faecalibacterium*, and *Turicibacter* have been observed in acute and chronic enteropathy in dogs [48]. Decreased *Bifidobacterium* but increased *Escherichia coli* have been reported in cats with IBD [19]. The disturbances of intestinal microbiota in canine IBD are similar to those in human IBD, characterized by reduced diversity, reduced Firmicutes, increased Proteobacteria, and reduced production of SCFAs and secondary bile acids [37,49]. The identification of microbial signatures associated with health and diseases can promote the development of probiotics. For instance, *Enterococcus hirae*, a dominant species of *Enterococcus* in the small intestinal mucosa of healthy shelter kittens, was found to be negatively associated with enteritis in kittens and was confirmed to have probiotic properties [50,51].

#### 3. Mechanisms of Action of Probiotics

Probiotics are live microorganisms with beneficial effects on host health when administered in adequate amounts, as defined by the International Scientific Association for Probiotics and Prebiotics [52]. Probiotics directly target the GIT, but their beneficial effects on the host can be local or systemic [20]. The most commonly studied probiotics include members of lactobacilli (e.g., *Lactobacillus acidophilus, Lacticaseibacillus casei, Lactobacillus delbrueckii* subsp. *bulgaricus, Lacticaseibacillus rhamnosus, Lactiplantibacillus plantarum,* and *Limosilactobacillus reuteri*), *Bifidobacterium (Bifidobacterium adolescentis, Bifidobacterium longum* subsp. *infantis, Bifidobacterium breve, Bifidobacterium longum*, and *Bifidobacterium bifidum*), and *Saccharomyces (Saccharomyces boulardii* and *Saccharomyces cerevisiae*), which have been used as probiotics for a long time with validated safety and efficacy [20,24,27]. Additionally, *Akkermansia muciniphila, Faecalibacterium prausnitzii, Prevotella copri, Bacteroides* spp., *Anaerobutyricum hallii, Parabacteroides goldsteinii, Roseburia,* and *Propionibacterium* are potentially next-generation probiotics [20,53,54].

Probiotics have been shown to promote health in many diseases, including IBD, diarrhea, colorectal cancer, allergy, type 2 diabetes, and atopic dermatitis, with strain-specific activity [21,24,55]. Probiotics have been proposed to have multiple mechanisms of action. Probiotics confer benefits to host health by interacting with the resident microbiota or by communicating with host cells. Probiotics can improve the balance of the intestinal microbiota, enhance the integrity of the epithelial barrier, and maintain immune homeostasis directly or indirectly [21,37,55,56]. Probiotics inhibit the colonization of pathogens via colonization resistance and production of antimicrobial molecules such as bacteriocins and organic acids (e.g., lactic acid) [20]. Interestingly, strains belonging to the Lactobacillus and Bifidobacterium genera do not produce butyrate per se, but can cross-feed other commensals to produce butyrate and increase levels of SCFAs, which have multifaceted and important physiological activities [12,57]. Moreover, the beneficial effects of probiotics can be imparted, at least in part, by the induction of host defense peptides (HDPs). Some probiotics, such as L. casei, L. paracasei, L. plantarum, B. breve, A. muciniphila, Bacteroides thetaiotaomicron, and E. coli Nissle 1917, stimulate the production of host-derived HDPs in humans and animals [58–61]. HDPs, also known as antimicrobial peptides, are a group of small cationic amphipathic peptides with a ubiquitous expression in epithelial cells and phagocytes [62]. With antimicrobial and immunomodulatory activities, HDPs constitute an important component of host immune defense [62,63].

The integrity of the intestinal epithelial layer is integral to intestinal homeostasis as the first physical barrier against the external environment and is largely maintained by the mucus layer and tight junctions linking adjacent epithelial cells [8,64,65]. In addition to the inhibition of pathogen colonization and stimulation of HDPs and SCFAs, probiotics contribute to the improvement in epithelial barrier function by promoting the production of mucins and tight junction proteins directly or indirectly [8,20,56,66]. Studies have demonstrated that *Lactobacillus*, *Bifidobacterium*, *A. muciniphila*, and SCFAs stimulate the expression, production, and secretion of mucins [8].

Moreover, probiotics modulate immune functions by stimulating T-cell differentiation, regulating pro- and anti-inflammatory cytokine profiles, and inducing sIgA production [21,55,67]. Furthermore, some probiotic strains, such as *Streptococcus thermophiles* and *Lactobacillus* spp., can produce  $\beta$ -galactosidase and bile salt hydrolase, which are involved in lactose digestion and bile acid metabolism, respectively [20]. Although the mechanisms of action of probiotics are diverse, it is noteworthy that they are heterogeneous and strainspecific [67]. A better understanding of probiotics—host interactions and precise probiotic mechanisms could accelerate the selection of effective probiotics [68].

### 4. Application of Probiotics in Dogs and Cats

Probiotics derived from dogs, cats, and other species have been evaluated in healthy animals and those with intestinal diseases in the form of monostrain or multistrain. The most commonly studied probiotics in dogs and cats are *Lactobacillus*, *Bifidobacterium*, and *Enterococcus* [27,47,69]. Indigenous microbiota is an important source of probiotics owing to the host specificity. Several strains isolated from dogs and cats have exhibited probiotic properties and hold promise for future use [27,28]. The utilization of probiotics in dogs and cats has been shown to improve the gut microbiota balance, modulate inflammation, enhance immune function, and protect against infections caused by enteropathogens [28,70–72].

#### 4.1. Dogs

Multiple canine-derived bacterial strains with probiotic properties were isolated and identified. A recent study isolated dozens of Lactobacillus strains from healthy dogs and revealed their antagonistic activities against Campylobacter in vitro [73]. L. plantarum strain RW1 isolated from canine feces displayed tolerance to low pH and bile salts, good adhesion to intestinal epithelial cells, and anti-inflammatory activity induced by Salmonella infection [74]. Ligilactobacillus animalis, identified as the dominant lactic acid bacterium in canine feces, showed various inhibitory effects against enteropathogens [75]. Other studies have isolated bacterial strains from dogs (mainly belonging to the genera Lactobacillus and Enterococcus) and characterized their probiotic properties in vitro [69,76–82]. The characteristics of probiotics are associated with host specificity, and it has been proposed that isolation from indigenous microbiota can yield the most optimal strains of probiotics [27,28]. For instance, canine-derived Enterococcus faecalis and Enterococcus faecium exhibited better adhesion to the intestinal mucosa of dogs than chicken-derived strains of the same species [83]. Similarly, a canine-derived *L. acidophilus* strain showed a higher adhesive ability to canine colonic mucus than strains derived from pigs and humans [80]. Moreover, some studies have confirmed the survivability of potential probiotic strains in gastric acid and the intestines of dogs [84,85].

Studies have indicated that the use of probiotics improves intestinal microbiota composition and changes microbial functions in healthy dogs by adopting 16S rRNA gene sequencing, metagenomics and metabolomics [86,87] (Table 2). Dietary supplementation with L. reuteri ZJF036 altered the fecal microbiota composition with increased relative abundances of Lactobacillus, but decreased relative abundances of Turicibacter and Blautia in healthy beagle dogs [88]. Feeding S. cerevisiae yeast increased the abundance of Turicibacter, decreased the abundance of *E. coli*, increased fecal butyrate, and shifted the microbial functional gene profile in healthy adult beagle dogs [89]. The administration of Lactiplantibacillus paraplantarum L-ZS9 changed the intestinal microbial diversity, composition, and metabolism in dogs [90]. Healthy dogs fed the probiotic mixture Slab51® had decreased fecal C. perfringens and increased fecal Bifidobacterium and Lactobacillus [70]. Metabolomics has been used to identify intestinal metabolic changes related to probiotic cheese feeding in healthy beagle dogs [87]. Probiotics also exert immunomodulatory activities in dogs [70,71]. Feeding the probiotic mixture Slab51 elevated fecal IgA and plasma IgG [70]. Multistrain probiotics containing three strains improved the levels of serum IgG and fecal sIgA [71]. However, in some cases, the administration of probiotics did not influence intestinal microbiota or immunological parameters in dogs [91,92].

Probiotics	Main Outcomes	Animals	References
L. reuteri ZJF036	fecal Lactobacillus $\uparrow$ fecal Turicibacter and Blautia $\downarrow$	healthy juvenile beagles; 75 $\pm$ 5-day-old; male	[88]
S. cerevisiae CNCM I-5660	fecal total biogenic amines and ammonia $\downarrow$ fecal pH $\downarrow$ fecal butyrate $\uparrow$ dysbiosis index $\downarrow$ fecal abundance of <i>Turicibacter</i> $\uparrow$ fecal abundance of <i>E. coli</i> $\downarrow$	healthy adult beagle dogs; 5-year-old; mixed sex healthy dogs; mixed breeds; 2.5 to 4 years old; mixed sex	[89]
probiotic mixture Slab51 <sup>®</sup> containing <i>S.</i> thermophilus DSM 32245, <i>B. lactis</i> DSM 32246, <i>B. lactis</i> DSM 32247, <i>L. acidophilus</i> DSM 32241, <i>L. helveticus</i> DSM 32242, <i>L.</i> paracasei DSM 32243, <i>L. plantarum</i> DSM 32244, and <i>L. brevis</i> DSM 27961	fecal <i>C. perfringens</i> ↓ fecal <i>Bifidobacterium</i> and <i>Lactobacillus</i> ↑ fecal IgA and plasma IgG↑		
cheese added with <i>L. reuteri</i> KACC 92293 and <i>B. longum</i> KACC 91563	fecal SCFAs $\uparrow$	healthy beagles; mixed sex	[87]
A mixture of <i>B. subtilis</i> and <i>B. licheniformis</i>	fecal score↓ fecal odor↓ fecal biogenic amines↓	healthy beagle dogs; 4-year-old; mixed sex	[93]
W. cibaria JW15	fecal ammonia emission $\downarrow$	healthy beagles; 1 to 2 years old; mixed sex	[94]
multistrain probiotics containing <i>L. casei</i> Zhang, <i>L. plantarum</i> P-8, and <i>B. animalis</i> subsp. <i>lactis</i> V9	fecal microbial diversity ↑ fecal beneficial bacteria ↑ fecal opportunistic pathogenic bacteria ↓ changes in microbial functional genes	dogs with diarrhea; German Shepherd and Belgium Shepherd dogs; age of 4 months to 13 years; mixed sex	[86]
probiotic mixture Slab51®	mast cells in colonic mucosa $\downarrow$	dogs with colonic dysmotility; mixed breeds; 2 to 10 years old; mixed sex	[95]
probiotic mixture containing <i>L. fermentum,</i> <i>L. rhamnosus,</i> and <i>L. plantarum</i>	frequency of diarrhea $\downarrow$	dogs treated with non-steroidal anti-inflammatory drugs (NSAIDs); mixed breeds; age of 3 months to 14 years; mixed sex	[96]
L. rhamnosus MP01, L. plantarum MP02	gastrointestinal infections $\downarrow$	puppy; German shepherd and Yorkshire; 1-month-old; mixed sex	[77]
probiotic mixture Visbiome <sup>®</sup> (L. plantarum DSM 24730, S. thermophilus DSM 24731, B. breve DSM 24732, L. paracasei DSM 24733, L. delbrueckii subsp. bulgaricus DSM 24734, L. acidophilus DSM 24735, B. longum DSM 24736, and B. infantis DSM 24737)	clinical recovery $\uparrow$ normalization of gut microbiome $\uparrow$	dogs with acute hemorrhagic diarrhea; mixed breeds; averagely aged 5.5 or 6 years	[97]
probiotics compound Visbiome®	duodenal and colonic expression of E-cadherin, occludin, and zonulin ↑	dogs with idiopathic IBD; mixed pure breeds; mean age of 6.2 and 4.6 years; mixed sex	[98]

Table 2. Effects of probiotics on the intestinal health of dogs.

The up arrow sign  $\uparrow$  indicates upregulation while the down arrow sign  $\downarrow$  indicates downregulation.

Moreover, studies have suggested improved fecal quality and reduced nitrogen fermentation through probiotic treatments in dogs [89,93,94]. Feeding *S. cerevisiae* yeast decreased the fecal pH and concentrations of total biogenic amines and ammonia [89]. Dietary supplementation with *Bacillus subtilis* and *Bacillus licheniformis* improved the fecal score, reduced nitrogen fermentation products in feces, and reduced fecal odor in dogs [93]. Dietary supplementation with *Weissella cibaria* JW15 decreased fecal ammonia emission in dogs [94]. In addition, probiotics do not seem to affect the nutrient digestibility or growth performance of dogs [70,88,89,94], with one study indicating that a multistrain probiotic compound improved feed intake and weight gain in dogs [71].

Probiotics confer health benefits in dogs with various intestinal disorders (Table 2). Studies have shown that multistrain probiotics improved clinical remission, suppressed inflammation, and increased epithelial barrier function in dogs diagnosed with IBD [47]. The probiotic mixture Slab51<sup>®</sup> alleviated the signs of canine colonic dysmotility [95]. A lactic acid bacteria product containing *Limosilactobacillus fermentum*, *L. rhamnosus*, and *L. plantarum* isolated from dogs decreased the frequency of diarrhea in dogs administered with non-steroidal anti-inflammatory drugs [96]. Feeding probiotics containing *Lactobacillus* and *Bifidobacterium* strains increased fecal microbiota diversity, improved microbiota structure, and regulated microbial functional pathways in dogs with diarrhea [86]. Dietary supplementation with *L. rhamnosus* MP01 or *L. plantarum* MP02 isolated from canine milk prevented gastrointestinal infections in puppies [77]. Oral administration of a probiotic mixture accelerated clinical recovery and intestinal microbiota normalization in dogs with acute hemorrhagic diarrhea [97]. A probiotic compound therapy increased the intestinal expression of tight junction proteins in dogs with idiopathic IBD [98].

#### 4.2. Cats

*Bifidobacterium* and other lactic acid-producing bacteria, which are largely detected in the feces of healthy cats, are potential sources for the selection of probiotics [41]. A recent study found that strains of *L. plantarum*, *L. rhamnosus*, *L. acidophilus* and *B. adolescentis* isolated from various biotopes of healthy cats showed in vitro probiotic properties such as adhesive activity and antimicrobial activity against causative agents of surgical infection in cats [27]. Another study identified several strains of *L. reuteri*, *L. fermentum*), *E. faecium* and *Pediococcus pentosaceus* from the feces of dogs and cats, although in vivo confirmation of safety and effectiveness is needed for further use [42]. *Bacteroides* sp. CACC 737 isolated from feline feces has potential probiotic properties, as revealed by functional genome analysis [99]. Freeze-dried *E. hirae* isolated from a healthy kitten decreased the diarrhea rate but did not influence the growth or composition of major bacterial phyla [50].

The administration of probiotics promoted the intestinal health of healthy dogs (Table 3). Supplementation with *L. acidophilus* CECT 4529 in the diet improved fecal quality and decreased fecal coliform counts in healthy adult cats [100]. Multistrain probiotics consisting of *S. boulardii* and *Pediococcus acidilactici* increased the production of butyric acid and total SCFAs, reduced concentrations of inflammatory markers myeloperoxidase and calprotectin, and improved the activity of antioxidant enzymes superoxide dismutase and glutathione in the feces of healthy short-haired cats [101]. Interestingly, this study showed that the use of the probiotic mixture did not change fecal microbiota structure in cats [101]. Feeding *E. faecium* SF68 decreased the diarrhea rate in shelter cats but did not affect the diarrhea rate of shelter dogs [102]. Dietary supplementation with *L. acidophilus* DSM13241 increased the numbers of *Lactobacillus* and *L. acidophilus* and decreased the numbers of *Clostridium* spp. and *E. faecalis* in the feces of healthy adult cats [103].

Probiotics	Main Outcomes	Animals	References [100] [101]
L. acidophilus CECT 4529	fecal quality ↑ fecal Lactobacilli count ↑ fecal Coliform count ↓	healthy adult Maine Coon cats; averagely aged 43.2 or 44.6 months; mixed sex	
a mixture of <i>S. boulardii</i> and <i>P. acidilactici</i>	fecal butyric acid and total SCFAs ↑ fecal inflammatory markers ↓ fecal antioxidant capacity ↑	healthy short-haired domestic cats; 2 to 4 years old; mixed sex	
E. hirae 1002-2	diarrhea rate $\downarrow$	healthy kitten; aged < 12 weeks; mixed sex	[50]
L. acidophilus DSM13241	fecal <i>Lactobacillus</i> ↑ fecal <i>Clostridium</i> and <i>E. faecalis</i> ↓	healthy domestic shorthair cats; 4 to 5.5 years old	[103]
B. licheniformis-fermented product	clinical signs↓ fecal <i>Clostridium</i> cluster XIVa ↑ fecal <i>C. perfringens</i> ↓	cats with chronic diarrhea; mixed breeds; age of 3 to 15 years; mixed sex	[104]
multi-strain probiotic (SLAB51 <sup>TM</sup> )	clinical symptoms↓ <i>Lactobacillus</i> and <i>Streptococcus</i> ↑	cats with chronic constipation and idiopathic megacolon; mixed breeds; 6 to 14 years old; mixed sex	[105]
E. faecium strain SF68	fecal quality $\uparrow$	cats administered amoxicillin-clavulanate; healthy young adult cats; purpose-bred cat; mixed sex	[106]
<i>E. faecium</i> strain SF68	diarrhea rate↓	shelter cats	[102]

Table 3. Effects of probiotics on the intestinal health of cats.

The up arrow sign  $\uparrow$  indicates upregulation while the down arrow sign  $\downarrow$  indicates downregulation.

Positive effects of probiotics have also been noted in cats with intestinal disorders (Table 3). Oral administration of *Bacillus licheniformis*-fermented products relieved diarrhea, enriched bacteria belonging to *Clostridium* cluster XIVa, and reduced *C. perfringens* in the feces of cats with chronic diarrhea [104]. A pilot study revealed that the oral administration of a multi-strain probiotic (SLAB51<sup>TM</sup>) composed of eight lactic acid bacteria strains ameliorated clinical signs and increased the abundance of fecal *Lactobacillus* and *Streptococcus* in cats with chronic constipation and idiopathic megacolon [105]. Feeding *E. faecium* strain SF68 improved the fecal quality in cats treated with amoxicillin–clavulanate, which can cause vomiting or diarrhea [106].

Although diverse aspects of probiotics have been elucidated, not all activities have been confirmed for every probiotic strain. There are inherent differences between probiotic strains, and the efficacy of probiotics can be influenced by many factors such as species, disease conditions, age, and sex [20,21,71,102,107]. A recent study showed that the bioactivity of probiotics might be sex-specific due to variations in the endocrine and central nervous systems between males and females [107].

#### 5. Future Directions

Probiotics exert health-promoting effects via direct interaction with the host cells and resident microbiota, or indirectly via microbial metabolites. The intestine hosts the largest and the most complex microbial community. More precise and accurate measurements of microbiota composition and further characterization of microbial functions in the intestine would expand the range and accelerate the selection of probiotics for use in cats and dogs [20]. Although multiple bacterial strains with probiotic characteristics have been

isolated from cats and dogs, their health-promotion effects are largely unconfirmed in vivo and there is a shortage of commercial canine/feline-derived probiotics products on the market, which necessitates further and accelerated studies.

It is worth noting that the microbiota can be influenced by many factors, such as species, age, sex, breed, diet, surgery, and antibiotic interventions [17,19,46,47,108–111]. These factors should be considered when evaluating the in vivo efficacy of probiotics, given that probiotics interact with the gut microbiota. It has been indicated that the administration dose or duration does not impact probiotic health benefits, as illustrated by the application of *L. fermentum* in dogs [72]. Whether this finding applies to other probiotics requires further investigation.

Although several mechanisms of action of probiotics have been proposed, not all have been verified in dogs and cats, warranting more comprehensive measurements and analysis in further research. Moreover, probiotic activities can be heterogeneous and strain-specific; therefore, a precise mechanistic investigation is needed.

Stability is one of the concerns in the application of probiotics. Probiotics such as Lactobacillus, Bifidobacterium, A. muciniphila, and F. prausnitzii are sensitive to stressful conditions such as heat, oxygen, pH, enzymes, and bile salts, which compromise their efficacy during processing, storage, distribution, and passage through the GIT [112]. Encapsulation has emerged as an effective strategy to enhance the viability, stability, and efficacy of probiotics [24,112]. Emulsions, gels, powder granules, nanofibers, electrospray capsules, and nano-coatings are commonly used probiotics encapsulation methods [24]. Each approach has specific advantages and disadvantages. However, nano-based encapsulation technology offers higher protection and delivery efficacy for probiotics in contrast with traditional encapsulation methods [113,114]. Furthermore, the single-cell nano-coating encapsulation strategy is considered a more advanced technology than bulk encapsulation methods with nanomaterials such as nanofibers and nanoparticles [113]. Research on encapsulation techniques utilized in probiotics for dogs and cats remains limited. A recent study has indicated that different microencapsulation techniques are differently efficient in maintaining the viability of probiotics during storage and passage through the GIT of cats [115]. Further studies are needed to verify encapsulation techniques for future applications in dogs and cats.

## 6. Conclusions

Maintaining the intestinal health of dogs and cats is becoming increasingly important as pet owners pay more attention to the health of their pets. The utilization of probiotics is an effective strategy to enhance the health of dogs and cats. The administration of probiotics can improve the balance of the intestinal microbiota, suppress inflammation, enhance immune function, and alleviate intestinal disorders in dogs and cats. Various bacterial strains derived from dogs and cats have shown probiotic properties in vitro, but their health benefits need to be confirmed in vivo. Moreover, the mechanisms of action of probiotics need to be further elucidated. Finer characterization of the intestinal microbiota of dogs and cats under different health conditions may facilitate the discovery of novel probiotics for use in pets.

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