



Use of Microorganisms as Nutritional and Functional Feedstuffs for Nursery Pigs and Broilers

Yi-Chi Cheng and Sung Woo Kim *D

Department of Animal Science, North Carolina State University, Raleigh, NC 27695, USA

* Correspondence: sungwoo_kim@ncsu.edu

Simple Summary: The use of microorganisms has become a trend as nutritional and functional feedstuffs become widely used in swine and poultry diets. Microorganisms, as coproducts obtained from the food industry and biorefineries, can reduce not only the burdens of the natural ecosystem but also the high costs of feedstuffs. It is possible to mitigate food and land competition with humans in the current global issues. These microorganisms could be promising and sustainable alternatives in animal diets because they contain highly valuable proteins, amino acids, fatty acid composition, and biogenic metabolites, which are beneficial for animal production. Microorganisms could be good alternatives to replace plant and animal-based protein supplements with high protein and a balanced amino acid composition. Lipid-rich microalgae and yeasts could be alternative energy feeds with valuable fatty acids used to enhance intestinal health and meat quality. In addition, microorganisms could be functional feed additives due to their cell contents and their cell wall bioactive components. However, there still are some limitations to using microorganisms, including the sources and dose of those microorganisms, which may cause negative effects on growth and health. Thus, this research focused on investigating the use of nutritional and functional microorganisms as feedstuffs and feed additives to replace conventional feedstuffs for enhancing the growth and intestinal health of nursery pigs and broilers.

Abstract: The objectives of this review paper are to introduce the structures and composition of various microorganisms, to show some applications of single cells as alternative protein supplements or energy feeds in swine and poultry diets, and to discuss the functional effects of microorganisms as feed additives on the growth performance and intestinal health of nursery pigs and broilers. Microorganisms, including bacteria, yeasts, and microalgae, have been commonly supplemented in animal diets because they are cost-effective, stable, and have quantitative production that provides nutritional and functional benefits to pigs and broilers. Microorganisms could be alternative antibiotics to enhance intestinal health due to bioactive components from cell wall components, which interact with receptors on epithelial and immune cells. In addition, bioactive components could be digested by intestinal microbiota to produce short-chain fatty acids and enhance energy utilization. Otherwise, microorganisms such as single-cell protein (SCP) and single-cell oils (SCOs) are sustainable and economic choices to replace conventional protein supplements and energy feeds. Supplementing microorganisms as feedstuffs and feed additives improved the average daily gain by 1.83%, the daily feed intake by 0.24%, and the feed efficiency by 1.46% in pigs and broilers. Based on the properties of each microorganism, traditional protein supplements, energy feeds, and functional feed additives could be replaced by microorganisms, which have shown benefits to animal's growth and health. Therefore, specific microorganisms could be promising alternatives as nutritional and functional feedstuffs in animal diets.

Keywords: bacteria; feedstuffs; feed additives; nursery pigs; microalgae; yeasts



Citation: Cheng, Y.-C.; Kim, S.W. Use of Microorganisms as Nutritional and Functional Feedstuffs for Nursery Pigs and Broilers. *Animals* **2022**, *12*, 3141. https://doi.org/10.3390/ ani12223141

Academic Editor: Yong Su

Received: 4 October 2022 Accepted: 8 November 2022 Published: 14 November 2022

Publisher's Note: MDPI stays neutral with regard to jurisdictional claims in published maps and institutional affiliations.



Copyright: © 2022 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (https:// creativecommons.org/licenses/by/ 4.0/).

1. Introduction

Animal diets make up 70% of the total costs of animal production [1]. Soybean meal (SBM) and corn are the main protein supplement and energy feed in animal diets, respectively. However, plant protein supplements contain anti-nutritional factors, including trypsin inhibitors, flatulence-producing compounds, and allergenic proteins, which restrict growth performance and intestinal development [2,3]. Some anti-nutritional factors could be eliminated via fermentation by using yeasts or bacteria to enhance nutrient bioavailability [4]. After fermentation, these microorganisms could be supplemented as coproducts in animal diets due to their valuable amino acids, vitamins, minerals, nucleotides, enzymes, and other metabolites [5–7]. In addition, the use of plant-based feedstuffs is dependent on seasonal availability and is limited to land use [8], whereas the use of microorganisms has fewer availability concerns and could be produced on a large scale in less time.

Animal-based feedstuffs are commonly supplemented in nursery diets to enhance growth performance, nutrient digestibility, and intestinal health [3,9,10]. Although animalbased feedstuffs have positive effects on growth and health development in pigs and broilers, these feedstuffs are expensive and in short supply [11,12]. It is important for nutritionists to seek alternative feedstuffs so that animal producers can reduce the cost burden while maintaining the growth performance of pigs and broilers. For alternative feedstuffs in pig and poultry diets, some key points need to be considered, including nutritional values, availability, palatability, and consistency [13,14]. Among alternative feedstuffs, coproducts from the food industry, insects, and some microorganisms can replace expensive feedstuffs in animal diets.

Coproducts from the food industry are convenient and easily available for delivery to feed mills. In the research from Kwak and Kang [15] using finishing pigs, a food waste mixture (70% food waste, 10% poultry litter, and 13% bakery coproducts) with an aerobic microbial culture could be supplemented in diets, replacing corn and SBM, without adverse effects on their growth and meat quality. However, supplementing bakery meal as an alternative energy feed reduces growth performance and the digestibility of AA in diets fed to nursery and growing pigs [16–18]. Candy coproducts could partially replace whey permeate without negative effects on the growth performance of nursery pigs [19]. The concern of using food coproducts is the variable nutrient composition by different processes and sources; therefore, it is important to analyze nutrient composition before formulation. Insect meal contains high protein and lipid content and is used to replace SBM and animal-based protein supplements [20,21]. Even though some studies demonstrated that corn-insect diets had better growth performance than corn-SBM diets in poultry diets, the price of insect meal remains high due to the low production [21,22]. Some microorganisms have been commonly supplemented in animal diets because they are cost-effective, stable, and their quantitative production provides nutritional and functional benefits to pigs and broilers. In addition, specific microorganisms could be divided into groups with different characteristics and functions. Further details are reviewed in the following sections.

1.1. Bacteria

Bacteria are unicellular and relatively small with a size range of 0.5 to 5.0 μm [23]. Bacteria are rich in lipids, proteins, and amino acids (Table 1). In addition, bacteria are categorized into two groups, Gram-positive and Gram-negative, based on different cell wall structures (Figure 1A). Peptidoglycan (PGN) is the major component (40 to 60%) of the cell wall and is made of N-acetylglucosamine (NAG), N-acetylmuramic acid (NAM), and short peptide chains, including L-alanine, D-glutamic acid, either L-lysine or diaminopimelic acid (DAP), and D-alanine [24,25] (Figure 1B). In the cell wall, cross-linking of the PGN envelope enhances the strength of the structure (Figure 1C). Gram-positive bacteria have a thicker cell wall due to more PGN envelopes in the cell wall than Gram-negative bacteria [26], whereas Gram-negative bacteria have an outer membrane, called the lipopolysaccharide (LPS), which may be toxic and affect animal health [23,27]. Based on properties of bacterial

cells, they were commonly used to replace fish meal in aquacultural diets without negative effects on intestinal health and growth [28]. Some studies demonstrated that the thick cell wall is non-digestible in mono-gastric animals; however, some bacterial cell walls can be utilized by the intestinal microbiota to enhance intestinal health [27,29].

Table 1. Characteristics and nutrient content of bacteria, yeasts, and microalgae.

	Bacte	eria	Yeast	Microalgae
DM ¹ , %	90 to	95	93	94
CP, % DM basis	50 to	80	12 to 53	10 to 70
Lipid, % DM basis	7 to	15	1 to 40	3 to 71
Total Fiber, % DM basis	3 to	6	2 to 40	10 to 66
	Gram + ²	Gram – ²		
Cell wall contents	20 to 80 nm; PN ³ (40 to 60%); 8 to 10 nm; Teichoic acid PN (10 to 20%); (up to 40%); LPS ⁴ ; Arabinogalactan Lipoprotein (10 to 20%) Lipoprotein		 Mannoprotein (35 to 40%); 1,3 β-glucan (50 to 55%); 1,6 β-glucan (5 to 10%); Chitin (up to 3%) 	Polysaccharide (1 to 12%) Soluble protein (up to 4.5%)
References	[30,3	31]	[23,32,33]	[34–36]

 1 DM—dry matter. 2 Gram +/ — Gram-positive (+)/negative (–) bacteria. 3 PN—peptidoglycan. 4 LPS—lipopolysaccharide.

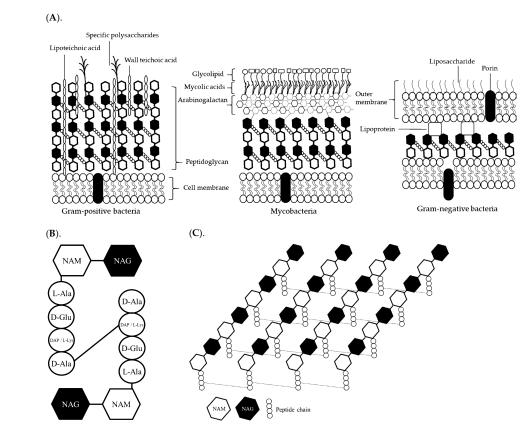


Figure 1. The structure of Gram-positive and Gram-negative bacterial cell walls (**A**), peptidoglycan (PG) structure (**B**), and the cross-linking of PG chains in bacteria (**C**). Concepts were based on Koch (2006) [37]; Kang et al. (2016) [38]; Pazos and Peters (2019) [25]. LPS, lipopolysaccharides; NAM, N-acetylmuramic acid; NAG, N-acetylglucosamine; L-Ala, L-alanine; D-Glu, D-glutamic acid; DAP, Diaminopimelic acid; L-Lys, L-lysine; D-Ala, D-Alanine.

1.2. Yeasts

Yeast sizes are variable from 2 to 50 μ m in length and 1 to 10 μ m in width [32]. Yeasts could grow under aerobic respiration, so they are generally used for brewing to produce alcohol [39]. In both inner and outer cell walls (Figure 2), β -glucans and mannoprotein are the major components [23] (Table 1). Chitin is a minor component, contributing approximately 1 to 3% in the yeast cell wall, and β -1,6 glucan links to the inner and outer walls, strengthening the cell structure [40]. Interestingly, yeast cell walls could stimulate animals to secrete protease and glucanase to release cell contents and cause fragmentation of cell walls [41]. Some studies demonstrated that β -glucans and mannoprotein from yeasts can enhance growth performance and intestinal health in pigs [42–44].

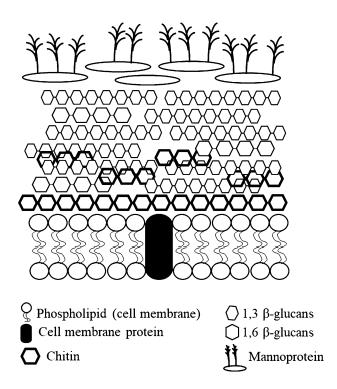
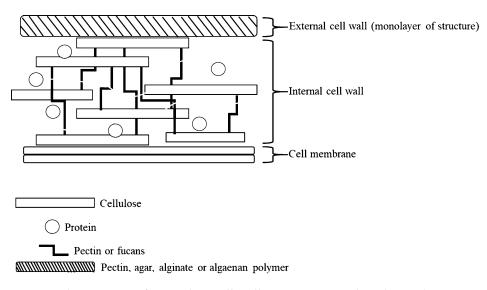


Figure 2. The structure of yeast cell wall. Concepts were based on Lipke and Ovalle (1998) [45]; Kogan et al. (2008) [46]; Morphology (2012) [32].

1.3. Microalgae

Within the microorganisms, microalgae are rich in essential fatty acids (FA), vitamins, and minerals [34] (Table 1). Cell wall components of microalgae are primarily made of cellulose, with pectin, fucan, xylan, and mannan as minor components [47,48] (Figure 3). Pyrrophyta contains two flagella and chlorophyll with carotenoid and xanthophyll as bioactive components, which accumulate starch via photosynthesis [49]. Compared to other types of microalgae, Chrysophyta includes cellulose, silica, and calcium carbonate in the cell wall and are able to accumulate lipids, including omega-3 [50], therefore, they are generally used as energy feeds [51–53].

There are several advantages of using nutritional and functional microorganisms: (1) Microbial production is an applicable and economical technology to obtain stable products and maintain the cell culture [56,57]. (2) It is eco-friendly because microorganisms that are considered as coproducts from the food industry or biofuel production could be recyclable and supplemented in animal diets [56,58]. (3) Microorganisms are high in nutrients, such as protein, AA, fats, and vitamins, and can be useful in animal diets. However, there are some issues with using microorganisms in animal diets due to low digestibility, heavy metals, and toxicity [12]. Although the cell wall is non-digestible in



pigs, bioactive components could be extracted to enhance intestinal health in pigs and broilers [59–61].

Figure 3. The structure of microalgae cell wall. Concepts were based on Velazquez-Lucio et al. (2018) [54]; Pôjo et al. (2021) [55].

The hypothesis of this review is that supplementing microorganisms as nutritional and functional feed additives in nursery diets is feasible. To achieve this, the objectives are as follows: (i) To introduce the structures and composition of various microorganisms, (ii) to show some applications of microorganisms as alternative protein supplements or energy feeds in animal diets, and (iii) to discuss the functional effects of microorganisms as feed additives on the growth performance and intestinal health of pigs and broilers.

2. Microorganisms as Functional Feed Additives

2.1. Introduction of Functional Feed Additives

The intestinal microbiota is an indicator of intestinal health, which assists digestion and absorption of nutrients, the development of the intestinal immune system, and the inhibition of the colonization of harmful microbiota [62–64]. Young animals are susceptible to pathogenic infections due to their immature gastrointestinal (GI) tract and microbiota community. Therefore, diet composition plays a critical role in developing the balance of intestinal microbiota [65,66]. Antibiotics have been supplemented in young animal diets to avoid disease and enhance the growth rate [67]. The role of antibiotics is to inhibit pathogen replication and destroy cell wall synthesis [68]. However, the use of antibiotics gives rise to pathogens developing antibiotic resistance and affecting the intestinal microbial population [69]. Many countries have banned the use of antibiotics due to chemical residues in animals and antibiotic resistance transferred to humans [68]. Consequently, alternative antibiotics, including prebiotics, probiotics, or postbiotics derived from bacteria, yeast, and microalgae, can be a safer alternative for use in swine production [29,42,43]. Probiotics are live microorganisms, which benefit animal growth and the intestinal microbial community [27,70]. Prebiotics are polysaccharides obtained from the cell walls of microorganisms [35]. On the other hand, postbiotics are metabolites and cell contents extracted from probiotics [71]. Probiotics, prebiotics, and postbiotics not only balance intestinal microbiota diversity [27,42] but also have positive effects on the immune system by preventing intestinal inflammation [29,43].

2.2. Mechanism and Application

2.2.1. Bacteria

The mechanism of immune response is complicated and varies based on different types of bacteria. Among the bacterial cell walls, PGN, teichoic acid (TA), and S-layers are the main cell wall components in most Gram-positive bacteria, whereas *Mycobacteria* contain PGN, arabinogalactans (AG), and mycolic acids, which make up the top layer [30]. Different from the Gram-positive bacteria, Gram-negative bacteria contain less peptidoglycan, porin, and lipopolysaccharides (LPS) in the cell wall. Peptidoglycans, TA, and AG interact with receptors on the epithelial and immune cells, including Toll-like receptor 2 (TLR2) and a cluster of differentiation 14 (CD14), which is a co-receptor of TLR2, and on the intestinal epithelial cells (IECs) [71–73]. Peptidoglycans are cleaved by PGN hydrolases into small fragments and recognized by nucleotide binding and oligomerization domain proteins (NOD). The NOD inhibits nuclear factor- κ B (NF- κ B) and proinflammatory cytokines, interleukin (IL) 6, IL-8, and tumor necrosis factor-alpha (TNF- α) [74,75]. Different from PGN, AG binds to C-type lectin receptor (CLR) on the surface of immune cells, dendritic cells, and macrophages that decrease the activity of NF- κ B and inhibit the release of proinflammatory cytokines [76].

When animals are infected by LPS from pathogenic bacteria, LPS stimulates TLR4 and NF-κB releasing interferons and pro-inflammatory cytokines. Probiotics, prebiotics, and postbiotics could act as immunostimulators to stimulate TLR to inhibit NF-κB and activate an anti-inflammatory response [77–79]. For example, selected *Bacillus* sp. not only help to decrease diarrhea caused by Escherichia coli (E. coli) infection but also improve growth performance [27,80] (Table 2). However, it is observed that not all *Bacillus* sp. exhibit positive effects on growth performance [81]. In addition, Lactic acid bacteria (LAB), including *Enterococcus* sp. and *Lactobacillus* sp., commonly occur in the GI tract due to a favorable environment and strong adhesion to the intestinal epithelial cells [82], which stimulate the inflammatory response to release cytokines and chemokines as pathogens enter the body [79]. This can increase lymphocyte proliferation and macrophage phagocytosis to reduce aggregated pathogens in the intestine [82]. Taras et al. [83] demonstrated that Enterococcus faecium fed to sows can reduce the mortality of newborn piglets and post-weaning diarrhea. Enterococcus faecium reduced serum IgG [67] and tended to reduce chlamydial infection in newborn piglets from infected sows [84]. Therefore, bacteria and their bioactive components could be functional feed additives for beneficial immune responses in nursery pigs.

Micro-	Species	F 1 4 44	D 1 (Lervel 9/	I	mprovemen	it, %	Animal	D (
organism	Species	Feedstuff	Product	Level, %	ADG	ADFI	FE	Model	Reference
Bacteria	Bacillus sp.	probiotic	$1.0\times 10^{10}~\text{CFU/g}$	0.01	11.1	2.94	8.28	Nursery pigs	[80]
Bacteria	<i>Bacillus</i> sp.	probiotic	$1.0\times 10^9~\text{CFU/kg}$	0.05 0.05	18.83 2.44	$20.5 \\ -0.80$	-0.91 1.21	Nursery pigs	[81]
Bacteria	Bacillus sp.	probiotic	$3.2 imes 10^9 \ \mathrm{CFU/kg}$	0.04	2.65	0.81	1.94	Growing- finishing pigs	[85]
Bacteria	<i>Bacillus</i> sp.	probiotic	$6.0\times 10^8~\text{CFU/g}$	0.05 0.03	9.80 4.93	$-0.44 \\ -0.15$	10.4 5.11	Growing pigs	[86]
Bacteria	Bacillus sp.	probiotic	$2.4 imes 10^8 \ \mathrm{CFU/g}$	0.45 0.30 0.15	9.12 6.92 2.20	4.87 4.06 1.01	4.17 2.78 1.24	Nursery pigs	[87]
Bacteria	Bacillus sp.	probiotic	$3.2 imes 10^9 \ \mathrm{CFU/kg}$	0.04	2.41	-10.6	6.63	Nursery pigs	[88]
Bacteria	Clostridium butyricum	probiotic	$1.0 imes 10^9 \ \mathrm{CFU/kg}$	0.05	2.54	1.76	0.00	Broilers	[89]

Table 2. Examples of bacteria as functional feed additives.

Micro-	Species	E J. (Due du at	Level, %	I	mprovemen	ıt, %	Animal	D . (
organism	Species	Feedstuff	Product	Level, 70	ADG	ADFI	FE	Model	Reference
Bacteria	Enterococcus faecium	probiotic	$2.0 imes 10^9 \ \mathrm{CFU/kg}$	0.05	4.93	5.14	0.00	Broilers	[89]
Bacteria	Enterococcus faecium	probiotic	$2.0 imes 10^9 \ \mathrm{CFU/kg}$	0.01	5.50	-0.77	6.01	Nursery pigs	[90]
Bacteria	Lactobacillus sp.	postbiotic	$6.0\times 10^{10}CFU/g$ and medium	0.20	26.1	20.0	9.52	Nursery pigs	[91]
Bacteria	Lactobacillus sp.	probiotic	$1.0\times 10^{10}\text{CFU/g}$	0.50	4.56	36.2	1.10	Nursery pigs	[88]
Bacteria	Lactobacillus sp.	probiotic	$5.0 imes 10^9 ext{ CFU/kg}$	0.10 0.15 0.20	6.06 3.56 -1.89	-7.20 -7.12 -10.2	12.5 10.3 8.44	Nursery pigs	[92]
Bacteria	Lactobacillus sp.	probiotic	$2.4\times 10^5\text{CFU/g}$	0.10 0.50 0.75 1.00	16.0 1.91 3.35 4.16	10.5 0.84 1.75 1.16	5.16 1.03 1.63 2.90	Nursery pigs	[93]

Table 2. Cont.

2.2.2. Yeasts

Beta-glucans, mannoprotein, and chitin are the main cell wall components of yeasts. Many studies have investigated how the yeast cell wall and its metabolites could modulate immune responses through pattern recognition receptors (PRR), including TLR and CLR [94,95]. Beta-glucans derived from yeast cell walls bind to the TLR2 and CLR family and dectin-1 receptor on enterocytes and immune cells [96,97]. Activated receptors give rise to Ig secretion and increase the number of goblet cells for the maintenance of intestinal structural integrity [98]. Li et al. (2006) [99] demonstrated that pigs fed β -glucans can inhibit secretion of TNF- α and IL-6 due to increased IL-10; therefore, nutrients would be utilized for increase the number of LAB, which improve intestinal health and alleviate pathogens infected [100].

Mannoprotein, located on the external cell wall, contains oligomannoside chains to produce mannose oligosaccharides in yeasts [46]. Mannose has the ability to bind to the mannose-specific lectin-type receptors on pathogenic bacteria or viruses to prevent colonization of the intestinal villi [60]. Additionally, mannose also binds to TLR4 and dectin-2 receptors to activate the immune responses releasing anti-inflammatory cytokines to avoid inflammation [96,97]. Yeasts provided to sows during the gestation and lactation period could improve the growth performance of offspring [101–103] because of positive effects on the establishment of beneficial microbiota in the GI tract [104]. Therefore, nursery pigs continuously fed yeasts in their diets showed enhanced digestibility of nutrients [43,104] and had positive effects on intestinal health and morphology by β -glucans, which can enhance growth performance [43] (Table 3). Although the high dose of β -glucans may reduce growth performance in pigs [99], yeasts and their metabolites may have functions similar to antibiotics to enhance growth and reduce inflammation [105].

Table 3. Examples of yeasts and microalgae as functional feed additives.

Missographism	Smaailaa	F 1.4.44	D 1 <i>i</i>	L 1 . 9/	Im	provemen	ıt, %	Animal	
Microorganism	Species	Feedstuff	Product	Level, %	ADG	ADFI	FE	Model	Reference
				0.05	10.6	5.16	5.38		
N I	Saccharomyces cerevisiae	Postbiotic	Glucans extracted	0.10	18.7	13.3	4.93	Broilers	[106]
Yeast				0.15	10.5	10.0	1.35		
				0.20	10.6	4.55	10.8		
Yeast	Saccharomyces cerevisiae	Prebiotic	Cell wall extract	0.30	4.71	-0.26	6.25	Broilers	[107]

Missographism	Smaaina	F 1.4.44	D 1 <i>i</i>	T 1 0/	Imj	provemen	t, %	Animal	D (
Microorganism	Species	Feedstuff	Product	Level, %	ADG	ADFI	FE	Model	Reference
Yeast	Saccharomyces cerevisiae	Prebiotic	Cell wall extract	0.03	-15.7	0.85	5.88	Nursery pigs	[108]
Yeast	Saccharomyces cerevisiae	Probiotic	$1.0 imes 10^9 \ \mathrm{CFU/g}$	0.10	4.48	20.1	-15.4	Nursery pigs	[109]
Yeast	Saccharomyces cerevisiae	Postbiotic	yeast culture	0.50 1.00 2.00	$6.31 \\ -6.80 \\ -8.01$	$9.48 \\ -1.07 \\ -5.47$	0.00 -2.79 0.93	Nursery pigs	[43]
Microalgae	Arthrospira platensis	Postbiotic	Spray dried algae, Setalg (Pleubian, France)	1.00	-0.25	4.21	-3.28	Nursery pigs	[65]
Microalgae	Arthrospira platensis	Postbiotic	Spirulina powder, NeoEnBiz Co. (Bucheon, Republic of Korea)	0.25 0.50 0.75 1.00	1.79 1.91 3.35 4.16	1.12 0.84 1.75 1.16	0.67 1.03 1.63 2.90	Broilers	[110]
Microalgae	Aurantiochytrium limacinum	Postbiotic	ALL-G Rich, Alltech Inc. (Lexingtong, KE, USA)	1.00	1.12	-0.38	1.22	Finishing pigs	[111]
Microalgae	Chlorella sp.	Postbiotic	Spray dried algae, Setalg	1.00	0.51	-1.40	1.64	Nursery pigs	[65]
Microalgae	Haematococcus pluvialis	Postbiotic	Novasta, AstaCarotene AB (Stockholm, Sweden)	0.04 0.18 0.90	$-2.00 \\ -1.77 \\ -0.74$	$-2.20 \\ -3.64 \\ -2.43$	0.00 1.90 1.27	Broilers	[112]
Microalgae	<i>Schizochytrium</i> sp.	Postbiotic	JB5, JINIS Co., Ltd. (Jeonju, Republic of Korea)	0.50 1.00	-1.33 1.56	2.23 0.48	-3.36 1.12	Nursery pigs	[113]

Table 3. Cont.

2.2.3. Microalgae

The bioactive components of microalgae are variable across different species. Many microalgae contain high omega-3 FA, which is converted to polyunsaturated fatty acids (PUFA), eicosapentaenoic acid (EPA), and docosahexaenoic acid (DHA) [114]. Both PU-FAs are beneficial to the integrity of cell membranes and reduce inflammation and oxidation [115]. In addition, the use of DHA-rich microalgae could improve meat quality by changing FA composition and reducing the backfat thickness [116,117]. Kibria and Kim (2019) [113] demonstrated that nursery pigs fed Schizochytrium sp. developed an enhanced immune system and displayed improved nutrient digestibility and feed efficiency (Table 3). Betacarotene, one type of carotenoid produced by *Schizochytrium* sp. and the family *Chrysophyceae*, could be antioxidants and immunomodulators [118,119]. In addition, the flavonoid is another common bioactive component, which has anti-inflammatory effects to inhibit NOD-, LRR-, and pyrin domain-containing protein 3 releasing pro-inflammatory cytokines [120]. Furbeyre et al. (2017) [65] demonstrated that pigs fed Chlorella and Arthrospira platensis had reduced incidence of diarrhea and provided antibiotic function to maintain the intestinal morphology in newly nursery pigs, which may increase beneficial microbiota in the intestine and nutrient digestibility, respectively.

Some microorganisms have been used as functional feed additives with clear mechanisms in immune responses. Their functional cell wall components, including peptidoglycan, teichoic acid, β -glucans, mannoprotein oligosaccharides, and flavonoids, have positive effects on animal's intestinal health to increase growth and nutrient digestibility. In addition, yeasts and microalgae have prebiotic effects on polysaccharides from the cell wall. Polysaccharides could be digested by intestinal microbiota to produce short-chain fatty acids (SCFA) and enhance energy utilization [35,121]. Short-chain fatty acids bind and activate G protein-coupled receptors related to lipid and glucose metabolism [31] that increase fatty acid oxidation in muscle and reduce fat deposition in adipose tissue [122]. Shen et al. (2009) [43] reported that increased SCFA production may be correlated with improved marbling scores. A recent study demonstrated that SCFA infusion in the ileum increased dressing weight and improved carcass traits by reducing N excretion and regulating lipid metabolism in growing to finishing pigs [123]. However, there are some concerns surrounding the use of microorganisms that may contain biogenic toxins, such as purines and heavy metals [124]. New technology could reduce these toxins and break the cell wall of microalgae to release functional metabolites as valuable feed additives in the future.

3. Single Cell Protein (SCP)

3.1. Introduction of SCP

Conventional protein supplements, soybean meal (SBM), and animal-based protein supplements, including meat and bone meal, blood plasma, and fish meal, are mainly utilized in animal diets. Soybean meal has anti-nutritional factors, including the trypsin inhibitor, glycinin, and flatulence-producing oligosaccharides, which reduce nutrient digestibility in diets fed to pigs [2,3,125]. Although animal-based protein supplements are highly digestible and can improve health and growth in pigs and broilers [126], they are relatively expensive and in short supply [11,127]. Therefore, SCP is the sustainable and economic choice to replace conventional protein supplements.

Single-cell proteins not only contain high protein content in the cell, namely, 30 to 50% in yeast, 50 to 80% in bacteria, and 60 to 70% in microalgae, but can also be efficiently produced [128]. The SCP has highly valuable protein and AAs, similar to SBM and animal protein supplements [12,129] (Table 4). Furthermore, the use of microorganisms is more eco-friendly and can reduce land usage and carbon production [130]. Many studies have demonstrated that SCP could be a beneficial alternative protein supplement to animals, reducing the portion of conventional protein supplements in diets to enhance growth performance [131,132], animal health [133,134], and meat quality [135,136].

			Conventi	onal Pr	otein Sup	plemen	ıt			Bact	eria		
		Fish	Meal	•	bean leal		ood sma	v	bacterium micum		lophilus otrophus	v	lococcus ulatus
CP, %		63.0		48.0		78.0		76.8		79.9		68.0	
	Arg	3.80	83%	3.50	117%	4.40	64%	4.09	61%	3.61	80%	4.56	123%
	His	1.40	30%	1.30	43%	2.50	36%	1.55	23%	1.54	34%	1.54	42%
	Ile	2.60	57%	2.10	70%	2.70	39%	3.35	50%	3.32	73%	3.01	81%
	Leu	4.50	98%	3.60	120%	0.40	6%	5.38	80%	5.45	120%	5.06	137%
Essential	Lys	4.60	-	3.00	-	6.90	-	6.74	-	4.54	-	3.70	-
AA, %	Met	1.70	37%	0.70	23%	0.80	12%	1.26	19%	1.83	40%	1.72	46%
	Phe	2.50	54%	2.40	80%	4.30	62%	2.78	41%	4.22	93%	2.70	73%
	Thr	2.60	57%	1.90	63%	4.50	65%	3.32	49%	3.97	87%	2.87	77%
	Trp	0.60	13%	0.70	23%	1.40	20%	0.56	8%	0.77	17%	2.21	60%
	Val	3.10	67%	2.20	73%	5.10	74%	4.61	68%	4.91	108%	3.94	106%
	Ala	3.90	85%	2.10	70%	4.00	58%	6.26	93%	6.04	133%	4.64	125%
	Asp	5.40	117%	5.40	180%	7.40	107%	6.68	99%	7.52	166%	5.66	153%
NT	Cys	0.60	13%	0.70	23%	2.60	38%	0.35	5%	0.57	13%	0.45	12%
Non-	Glu	7.90	172%	8.50	283%	10.90	158%	8.86	131%	10.50	231%	7.35	198%
essential	Gly	4.70	102%	2.00	67%	2.80	41%	3.33	49%	5.69	125%	3.34	90%
AA, %	Pro	2.90	63%	2.50	83%	4.30	62%	2.32	34%	NA	NA	2.57	69%
	Ser	2.40	52%	2.40	80%	4.20	61%	2.34	35%	2.60	57%	2.37	64%
	Tyr	1.90	41%	1.60	53%	3.90	57%	1.81	27%	3.48	77%	2.43	66%
Reference		[1	37]	[1	37]	[1	37]	[138	,139]	[140	,141]	[135,1	42,143]

Table 4. Composition of crude protein (CP) and amino acids (AA-to-lysine ratio) of conventional protein supplements and bacterial protein supplements.

3.2. Application of SCP 3.2.1. Bacteria

Based on the nutrient composition, bacterial protein supplements provide a similar amino acid composition to SBM and fish meal [138,144]. There are some bacterial protein supplements used in pig and poultry diets, including *Corynebacterium glutamicum* [138,139], *Methylobacterium extorquens* [145], and *Methylococcus capsulatus* [135,146] (Table 4). Supplementing *Methylococcus capsulate* up to 12 % to replace SBM and fish meal in nursery diets may improve growth performance and meat quality due to changes in FA composition in meat [131,135]. In broiler diets, supplementing *Methylophilus methylotrophus* negatively affected growth performance, whereas intestinal health, microbial community, and disease resistance were improved [147–149].

Corynebacterium glutamicum and *E. coli* are mainly used to produce AA [150,151]. These bacteria have been considered waste after AA production, but they contain high levels of protein and AA [139]. Escherichia coli is Gram-negative bacteria with double layers of membrane and contains LPS. Some strains of E. coli may cause low nutrient digestibility and inflammation by binding to TLR4 on enterocytes and activating inflammatory effects [96,152]. However, Corynebacterium glutamicum is a Gram-positive and endotoxin-free bacterium, which is generally recognized as safe. This bacterium could be considered a single-cell protein supplemented in nursery diets that can improve growth performance and stimulate immune responses by increasing immunoglobulins (Ig) due to the bioactive components from the CGCM cell wall [139]. Within the same genus, Corynebacterium ammoniagenes supplemented up to 1% replacing SBM in poultry diets enhanced the daily gain and feed efficiency; however, increasing the inclusion of Corynebacterium ammoniagenes caused negative effects on growth and meat quality as a result of the low digestibility of protein and AA [132,153]. The overall result of supplementing bacterial protein supplements improved ADG by 0.12%, while reducing ADFI by 0.7% and FE by 0.41% in pigs and broilers (Table 5).

Micro-	Species	Product	Level, %	Improver	nent, %		Animal Model	Reference
organism		Tiouuct	20101, 10	ADG	ADFI	FE		Reference
Bacteria	Corynebacterium ammoniagenes	Protide, CJ	1.00 3.00 5.00	3.69 -1.02 -3.07	$1.82 \\ -1.18 \\ -1.50$	2.08 0.00 -1.56	Broilers	[132]
Bacteria	Corynebacterium glutamicum	Bacteria and medium	2.50 5.00	$-4.32 \\ -8.38$	-1.89 1.89	$-2.08 \\ -10.4$	Nursery pigs	[138]
Bacteria	Corynebacterium glutamicum	Lysed bacteria	0.70 1.40 2.10	$-1.27 \\ -1.27 \\ 11.4$	0.00 5.31 8.78	$-3.64 \\ -9.09 \\ 0.00$	Nursery pigs	[139]
Bacteria	Methylococcus capsulatus	BP, Dansk Bioprotein	10.7 12.0 8.00 4.00	5.59 8.48 5.65 -0.22	1.31 7.82 2.42 4.55	3.44 -1.26 -2.52 3.77	Growing pigs Nursery pigs	[131]
Bacteria	Methylococcus capsulatus	BBP, Norferm AS	6.00 4.00 2.00	2.68 5.01 3.98	$-3.08 \\ -0.57 \\ 0.81$	5.88 5.29 3.53	Broilers	[142]
Bacteria	Methylophilus methylotrophus	Bacteria and medium	10.0 20.0	-1.44 9.41	0.20 -0.08	1.78 1.78	Nursery pigs	[154]
Bacteria	Methylophilus methylotrophus	Bacteria and medium	9.60 19.2	$\begin{array}{c} 4.01 \\ -14.2 \end{array}$	2.23 -6.95	9.60 -8.09	Broilers	[140]
Bacteria	Methylophilus methylotrophus	Bacteria and medium	3.65 6.35 9.00 13.6	$0.15 \\ -1.00 \\ -8.00 \\ -13.0$	$-4.00 \\ -4.00 \\ -8.00 \\ -13.0$	$-4.00 \\ -3.00 \\ -1.00 \\ 0.00$	Broilers	[149]

Table 5. Single-cell proteins from bacteria and their impacts on growth performance.

3.2.2. Yeasts

Torula yeast (*Candida utilis*) and brewer's yeast (*Saccharomyces cerevisiae*) contain 45 to 55% protein in the total cell, which could be considered protein supplements in animal diets [155–158] (Table 6). Yeasts are rich in Lys but insufficient in sulfur AA, therefore additional Met must be considered in feed formulation [155]. Yeasts would be utilized by intestinal microbiota and produce SCFA, which are energy feeds for the health of intestinal epithelial cells [159]. The overall impact of supplementing yeast protein supplements may reduce ADG by 0.16% and ADFI by 1.86% but improve FE by 1.89% in pigs and broilers (Table 7). However, studies demonstrated that supplementing yeasts at a certain level and replacing conventional protein supplements had positive effects on growth performance, nutrient digestibility, and intestinal morphology in nursery pigs [156,160–162]. The reason for enhanced growth performance is due to β -glucans and mannoprotein from the cell wall, which are beneficial to animal's health [41,158]. However, some studies demonstrated that supplementing yeasts and replacing SBM and fish meal in the diets did not affect intestinal health in nursery pigs regarding the immune response and liver biomarkers [66,159].

Table 6. Composition of crude protein (CP) and amino acids (AA-to-lysine ratio) of single-cell protein from yeasts and microalgae.

				Yeas	st					Micro	oalgae		
		Torul	a Yeast		aromyces visiae		rowia lytica		desmus p.	Chlor	ella sp.	Nannochloropsis oceanica	
CP, %		49.1		44.2		43.5		31.2		47.7		38.2	
	Arg	2.39	72%	2.29	73%	1.81	55%	1.50	94%	3.88	123%	1.99	88%
	His	0.89	27%	1.05	34%	0.95	29%	0.50	31%	0.92	29%	0.64	28%
	Ile	2.16	66%	1.92	62%	1.99	61%	1.10	69%	1.87	60%	1.50	66%
	Leu	3.16	96%	2.99	96%	3.10	94%	2.30	144%	3.58	114%	2.90	128%
Essential	Lys	3.30	-	3.11	-	3.28	-	1.60	-	3.15	100%	2.27	-
AA, %	Met	0.58	18%	0.73	23%	0.72	22%	0.50	31%	0.84	27%	0.57	25%
	Phe	1.92	58%	1.82	58%	1.54	47%	1.30	81%	2.12	67%	1.57	69%
	Thr	2.10	64%	2.19	70%	2.01	61%	1.30	81%	2.63	84%	1.54	68%
	Trp	0.59	18%	0.57	18%	0.65	20%	0.40	25%	0.24	7%	0.49	22%
	Val	2.49	76%	2.24	72%	2.39	73%	1.60	100%	3.44	109%	2.13	94%
	Ala	3.03	92%	2.68	86%	3.63	111%	2.30	144%	1.39	44%	2.22	98%
	Asp	3.98	121%	4.49	144%	3.58	109%	2.70	169%	0.03	1%	2.80	123%
Non-	Cys	0.46	14%	0.54	17%	0.44	13%	0.30	19%	0.42	13%	0.30	13%
	Glu	6.77	205%	6.57	211%	6.07	185%	2.90	181%	2.04	65%	3.34	147%
essential	Gly	1.94	59%	1.75	56%	1.96	60%	1.70	106%	2.43	77%	1.92	85%
AA, %	Pro	1.55	47%	2.10	68%	1.72	52%	2.70	169%	0.94	30%	4.00	176%
	Ser	1.78	54%	2.32	75%	1.82	55%	1.10	69%	0.78	25%	1.21	53%
	Tyr	1.48	45%	1.56	50%	1.50	46%	1.00	63%	1.77	56%	1.20	53%
Reference	s	[137,156	,163,164]	[1	37]	[1	.65]	[1	34]	[166	-168]	[1	69]

Table 7. Single-cell proteins from yeasts and their impacts on growth performance.

Micro-	<u>Seconda</u>		T 1.0/		Improvemer	nt, %	Animal	Doforonco
organism	Species	Product	Level, %	ADG	ADFI	FE	Model	Reference
Yeast	Saccharomyces cerevisiae	Autolyzed yeast	1.25 2.50 5.00	-1.05 -8.99 -12.6	5.56 -0.20 2.69	$1.89 \\ -0.47 \\ -7.55$	Broilers	[136]
Yeast	Saccharomyces cerevisiae	Whole yeast Yeast extract	0.50 0.30	4.32 3.74	-0.33 2.45	5.68 2.84	Broilers	[107]
Yeast	Saccharomyces cerevisiae	Yeast and medium	3.00	-3.61	1.60	-2.27	Nursery pigs	[170]

Micro-	Smaailaa		T 1 0/		Improvemen	ıt, %	Animal	D (
organism	Species	Product	Level, %	ADG	ADFI	FE	Model	Reference
Yeast	Torula yeast	Extracted yeast	4.00 7.00 10.0 15.0	-1.00 -3.00 -4.00 -6.00	-3.00 -3.00 -3.00 -5.00	$-3.00 \\ -1.00 \\ 1.00 \\ 1.00$	Broilers	[149]
Yeast	Torula yeast	Yeast and medium	20.0 4.75	-4.87 1.38	$-0.77 \\ -3.72$	-4.44 5.31	Broilers	[163]
Yeast	Torula yeast	SylPro, Arbiom Inc	10.8 9.00 16.0 23.0	8.76 -3.75 -3.00 -7.87	2.29 -2.93 -7.32 -12.4	6.76 2.38 7.77 2.69	Nursery pigs	[164]
Yeast	Yarrowia lipolytica	Yeast and medium	3.00 6.00	12.4 -1.81	-1.25 -1.63	11.9 -0.63	Nursery pigs	[165]
Yeast	Yarrowia lipolytica	Yeast and medium	3.00	2.27	-1.20	2.14	Nursery pigs	[170]

Table 7. Cont.

3.2.3. Microalgae

Microalgae contain high values of oil and increased protein concentrations after oil extraction [171,172] (Table 6). Therefore, the de-fatted microalgae could be used as protein supplements, and their protein contents vary from 12 to 65% CP based on different microalgae [173]. The overall impact of supplementing microalgal protein supplements improves ADG by 2.24% and FE by 0.44% but reduces ADFI by 0.13% in pigs and broilers (Table 8). Dietary *Arthrospira platensis* of 15%, replacing SBM, is beneficial to growth and health due to enhanced activities of digestive enzymes and nutrient utilization [174]. This microalga could improve meat quality by changing the FA composition and increasing flavor, while the color of meat is more yellow due to the high amount of zeaxanthin in the microalgae [174,175]. However, the growth performance in pigs fed *Arthrospira platensis* did not change, whereas improvements were seen in regard to oxidative stress in muscles and meat quality [176,177]. In contrast, the use of microorganisms *Desmodesmus* sp. and *Nannochloropsis oceanica* in pig and broiler diets promoted protein and FA synthesis by increased expression of the mammalian target of rapamycin (mTOR) and acetyl CoA carboxylase [134,178,179].

Table 8. Single-cell proteins from microalgae and their impacts on growth performance.

Micro-	Smaatias	D 1 (I1 0/	I	mprovemer	nt, %	Animal	D (
organism	Species	Product	Level, %	ADG	ADFI	FE	Model	Reference
Microalgae	Arthrospira platensis	Spirulina, Sopropeche	15.0	-11.5	-2.42	-10.1	Broilers	[174]
Microalgae	Arthrospira platensis	Spirulina powder, Sopropeche	10.0	-12.4	-3.71	-9.46	Nursery pigs	[180]
Microalgae	Chlorella sp.	Pure, whole	7.50 15.0	0.21 -2.07	2.32 -0.51	$-2.50 \\ -1.87$	Broilers	[166]
Microalgae	Chlorella sp.	Allmicroalgae, Natural Products	5.00	12.6	4.30	7.43	Finishing pigs	[167]
Microalgae	Chlorella sp.	Allmicroalgae, Natural Products	10.0	5.44	-3.48	3.14	Broilers	[168]
Microalgae	Desmodesmus sp.	DGM, Cellana	10.0 15.0	-11.4 5.21	-9.73 8.56	$-1.64 \\ 16.42$	Nursery pigs Broilers	[134]

Micro-	Smaatias	D 1 (L	I	mproveme	nt, %	Animal	D (
organism	Species	Product	Level, %	ADG	ADFI	FE	Model	Reference
Microalgae	Desmodesmus sp.	Pure, whole Pure, defatted	5.00 5.00	35.36 20.91	$-0.24 \\ -2.37$	12.24 5.44	Broilers	[179]
Microalgae	Nannochloropsis oceanica	DGA, Cellana	2.00 4.00 8.00 16.0	0.84 -2.39 1.97 -10.4	-1.67 0.83 3.33 -3.33	$1.54 \\ -3.08 \\ 0.00 \\ -6.15$	Broilers	[169]
Microalgae	Staurosira sp.	Pure, defatted	7.50	1.20	6.21	-4.78	Broilers	[51]

Table 8. Cont.

The usage levels of microorganisms should be approached cautiously, as increasing the dose may cause negative effects on growth and health in animals due to the low digestibility of the cell wall of microorganisms and over-reaction to bioactive components [139]. The thick cell wall could be broken down by various technologies, including autolysis and hydrolysis [181,182]. Each technique may reduce palatability and growth performance in animals due to the change in nutrient composition [181,183] and affect nutritional values [7].

In summary, microorganisms such as SCPs are beneficial to animals' growth, health, and meat quality. However, SCP may cause some problems with the usage levels and technology of production [184,185], which need to be considered during feed formulation. Therefore, further studies are needed to discuss the appropriate use of single-cell proteins in animals.

4. Single-Cell Oil (SCO)

4.1. Introduction of SCO

Dietary lipids provide critical energy in feeds and essential fatty acids (EFA), increase nutrient absorption, and reduce feed dust [137]. For energy feeds, vegetable oils and animal fats have been used for over 35 years around the world [186]. Common lipid sources in pig diets are vegetable oils, animal fats, and animal-vegetable fat blends. Fats of animal origin, including poultry fat, tallow, and lard, have been used for a long time due to their higher digestibility in pigs [187,188]. The production of animal fats has increased in recent years and supplied the food industry, animal industry, and diesel production with approximately 3000 million pounds in 2019, but the price also increased from 20 cents/pound to 30 cents/pound from 2006 to 2019 in the U.S [189]. In addition, the European Union has become concerned with the use of animal fat regarding animal health due to disease, bovine spongiform encephalopathy, and chemical contaminations [190]. Vegetable oils, including soybean oil, corn oil, palm oil, and coconut oil, supplemented in diets may enhance higher amounts of long-chain n-3 poly-unsaturated fatty acids (PUFA) in carcasses [191,192]. Although vegetable oils are popularly utilized in various areas including the biodiesel and food industries, the production of vegetable oils competes for land with humans and emits thousands of tons of CO_2 [193]. Therefore, the rise in costs of conventional oils as energy feeds may be substituted with SCOs, such as Lipomyces starkeyi, Yarrowia lipolytica, and Schizochytrium species [170,194,195].

The advantages of SCO include decreased land usage, lower cost, and a shorter life cycle for large-scale production. Apart from their high protein content, SCO also has valuable fatty acids (Table 9). Microbial oils contain over 20% of lipid content and valuable polyunsaturated fatty acids (PUFA), which enhance immunity in young animals [196,197]. Fatty acid composition from different dietary lipids may affect animal growth performance [198], energy digestibility [198–200], intestinal health [201], and meat quality [202,203]. In fish diets, fish need high n-3 PUFA, including EPA and DHA, so plant oils and fish oils are the main energy feeds for fish [204]. Some studies demonstrated that SCOs replacing conventional lipids in diets can be practically used in aquaculture due to similar fatty acid composition and no adverse effects on fish growth and quality [205–207].

	Poultry Fat	Soybean Oil	Yarrowia lipolytica	Schizochytrium sp.	Crypthecodinium cohnii	
ME, kcal/kg	8364	8574	-	-	-	
Total saturated, %	28.7	14.2	19.4	36.5	-	
Total unsaturated, %	64.8	81.0	80.6	62.4	-	
FA, %				-		
C 14:0	0.9	0.1	0.3	11.0	16.0	
C 16:0	21.6	10.3	10.7	38.5	25.0	
C 16:1	5.7	0.2	1.5	18.5	0.4	
C 18:0	6.0	3.8	6.6	1.10	-	
C 18:1	37.4	22.8	8.8	3.15	16.0	
C 18:2	19.5	51.0	22.9	-	0.5	
C 18:3	1.0	6.8	2.3	-	0.4	
C 20:0	-	-	0.7	-	-	
C 20:1	1.1	0.2	0.2	0.60	-	
C 20:4	0.1	0.0	4.0	-	-	
C 20:5	0.0	0.0	30.2	1.65	0.1	
C 22:1	0.0	0.0	0.9	0.10	-	
C 22:5	0.0	0.0	0.9	12.9	-	
C 22:6	0.0	0.0	-	24.0	39.0	
References	[137]	[137]	[206]	[208,209]	[207]	

Table 9. Fatty acid composition (% of total lipids) in different sources of lipid supplements.

4.2. Application of SCO

Within SCOs, oleaginous yeasts have been involved in various biotechnological applications [210,211] (Table 10). Yarrowia lipolytica has the ability to produce valuable protein, lipids, lipolytic enzymes, and organic acids, which have been widely used in the food industry [212]. Yarrowia lipolytica is not only an alternative protein supplement in animal diets but also a lipid source containing 20% lipids in the cell [206]. When 3% dried Yarrowia lipolytica was used as a protein supplement, replacing soybean meal, it improved ADG and feed efficiency [165]. However, 6% Yarrowia lipolytica, with its high lipid content, resulted in diarrhea in piglets, as well as a reduction in the growth performance of nursery pigs [165]. In addition, Cheng et al. (2022) [213] demonstrated that 1.5% Yarrowia lipolytica used as energy feeds, replacing poultry fat, maintained intestinal health and growth performance in nursery pigs, while the thick cell wall may reduce nutrient digestibility when supplementing 3% Yarrowia lipolytica. Hatlen et al. (2012) [206] reported that 10 to 30% Yarrowia lipolytica supplemented in fish diets improved feed efficiency and protein and energy retention; however, protein digestibility and energy digestibility were reduced due to the indigestible yeast cell wall. The result may indicate that the lysis of yeast cell walls may be required to release nutrients and increase nutrient digestibility in diets. Berge et al. (2013) [214] reported that disrupted Yarrowia lipolytica released more lipids from the cell and improved nutrient digestibility. Another oleaginous yeast, Lipomyces starkeyi, is a feasible replacement for vegetable oils in fish without adverse effects on fish growth and meat quality [205].

Table 10. Single-cell oils from microorganisms and their impacts on growth performance.

Microorganism	Species	Level, %	Improvement, %				
			ADG	ADFI	FE	— Animal Model	Reference
Yeast	Yarrowia lipolytica	1.50 3.00	15.2 4.64	-5.08 -3.23	20.0 7.27	Nursery pigs	[213]
Microalgae	Aurantiochytrium acetophilum	1.00 2.00 4.00	$-2.65 \\ -4.05 \\ -12.7$	6.42 1.83 1.83	$-8.18 \\ -6.21 \\ -14.7$	Broilers	[215]
Microalgae	Schizochytrium sp.	3.12	-0.23	-3.69	4.35	Nursery pigs	[216]
Microalgae	Schizochytrium sp.	3.60	-1.92	-0.95	-0.86	Growing-finishing pigs	[217]
Microalgae	Schizochytrium sp.	3.70	-0.93	-3.03	3.45	Growing-finishing pigs	[218]
Microalgae	Schizochytrium sp.	0.25 0.50	4.17 4.17	0.65 1.18	3.38 3.80	Growing-finishing pigs	[116]

Microalgae is high in n-3 PUFA, especially EPA (20:5n-3) and DHA (22:6n-3), so it is effective for use in young fish [219]. Harel et al. (2002) [220] reported that adding microalgae *Crypthecodinium* sp. to replace fish oil in aquacultural diets demonstrated similar growth performance compared with a commercial control diet. Due to the high arachidonic acid proportion in microalgae, *Crypthecodinium* sp. improved the hatching rate of eggs [220] and reduced mortality during the larval stage of fish [207,221]. Supplementing *Schizochytrium* sp. not only improved growth performance but also enhanced intestinal health in nursery pigs and meat quality in growing to finishing pigs based on functional FA [116,216]. However, microalgae as energy feed are not competitive compared to other sources of oils due to the price of production and animal feasibility and acceptability [14]. Furthermore, microalgae can accumulate heavy metals, which may cause animal health problems, so it should be used cautiously to prevent toxic effects [173,194]. Even though SCOs are not as common as animal fats and plant oils supplemented in pig and broiler diets, SCOs may be promising alternative energy feeds based on their valuable FA for animal health and growth.

5. Conclusions

The production of selected microorganisms from fermentation is one of the sustainable solutions for the environmental challenges of animal agriculture. Selected microorganisms with nutritional and functional roles in improving the growth and health of young animals provide enhanced production efficiency and profits in animal agriculture. From the review, the use of selected microorganisms as feedstuffs and feed additives enhanced growth by 1.83%, feed intake by 0.24%, and feed efficiency by 1.46% in nursery pigs and broilers. Selected microorganisms, based on their properties, can reduce the use of traditional protein supplements, energy feeds, and functional feed additives. Collectively, selected microorganisms can be promising alternatives as nutritional and functional feedstuffs in diets for nursery pigs and broilers.

Author Contributions: Conceptualization, Y.-C.C. and S.W.K.; methodology, S.W.K.; formal analysis, Y.-C.C.; investigation, Y.-C.C.; resources, S.W.K.; data curation, Y.-C.C. and S.W.K.; writing—original draft preparation, Y.-C.C. and S.W.K.; writing—review and editing, Y.-C.C. and S.W.K.; supervision, S.W.K.; project administration, S.W.K.; funding acquisition, S.W.K. All authors have read and agreed to the published version of the manuscript.

Funding: This research was funded by the North Carolina Agricultural Foundation.

Institutional Review Board Statement: Not applicable.

Informed Consent Statement: Not applicable.

Data Availability Statement: Not applicable.

Conflicts of Interest: The authors declare no conflict of interest.

References

- Pomar, C.; Remus, A. Precision Pig Feeding: A Breakthrough toward Sustainability. *Anim. Front.* 2019, 9, 52–59. [CrossRef] [PubMed]
- Hong, K.-J.; Lee, C.-H.; Kim, S.W. Aspergillus Oryzae GB-107 Fermentation Improves Nutritional Quality of Food Soybeans and Feed Soybean Meals. J. Med. Food 2004, 7, 430–435. [CrossRef] [PubMed]
- Deng, Z.; Duarte, M.E.; Jang, K.B.; Kim, S.W. Soy Protein Concentrate Replacing Animal Protein Supplements and Its Impacts on Intestinal Immune Status, Intestinal Oxidative Stress Status, Nutrient Digestibility, Mucosa-Associated Microbiota, and Growth Performance of Nursery Pigs. J. Anim. Sci. 2022, 100, 1–76. [CrossRef]
- 4. Kim, S.W. Bio-Fermentation Technology to Improve Efficiency of Swine Nutrition. *Asian-Australas. J. Anim. Sci.* 2010, 23, 825–832. [CrossRef]
- Babel, W.; Pöhland, H.-D.; Soyez, K. Single Cell Proteins. In Ullmann's Encyclopedia of Industrial Chemistry; Wiley-VCH Verlag GmbH & Co. KGaA: Weinheim, Germany, 2000; pp. 437–452. ISBN 9781482279047.
- 6. Giec, A.; Skupin, J. Single Cell Protein as Food and Feed. Food/Nahrung 1988, 32, 219–229. [CrossRef]
- Ugalde, U.O.; Castrillo, J.I. Single Cell Proteins from Fungi and Yeasts. In *Applied Mycology and Biotechnology*; Elsevier: Amsterdam, The Netherlands, 2002; Volume 2, pp. 123–149.

- Rounsevell, M.D.A.; Ewert, F.; Reginster, I.; Leemans, R.; Carter, T.R. Future Scenarios of European Agricultural Land Use. Agric. Ecosyst. Environ. 2005, 107, 117–135. [CrossRef]
- 9. Kim, S.W.; Easter, R.A. Nutritional Value of Fish Meals in the Diet for Young Pigs. J. Anim. Sci. 2001, 79, 1829. [CrossRef]
- Weaver, A.C.; Campbell, J.M.; Crenshaw, J.D.; Polo, J.; Kim, S.W. Efficacy of Dietary Spray Dried Plasma Protein to Mitigate the Negative Effects on Performance of Pigs Fed Diets with Corn Naturally Contaminated with Multiple Mycotoxins. *J. Anim. Sci.* 2014, 92, 3878–3886. [CrossRef]
- 11. U.S. Department of Agriculture. Agricultural Statistics 2019; Bernan Press: Washington, DC, USA, 2019; ISBN 978-1-64143-466-9.
- 12. Kim, S.W.; Less, J.F.; Wang, L.; Yan, T.; Kiron, V.; Kaushik, S.J.; Lei, X.G. Meeting Global Feed Protein Demand: Challenge, Opportunity, and Strategy. *Annu. Rev. Anim. Biosci.* **2019**, *7*, 221–243. [CrossRef]
- Kim, S.W.; Hansen, J.A. Diet Formulation and Feeding Programs. In Sustainable Swine Nutrition; Wiley: Oxford, UK, 2013; pp. 215–227. ISBN 9780813805344.
- 14. Bharathiraja, B.; Sridharan, S.; Sowmya, V.; Yuvaraj, D.; Praveenkumar, R. Microbial Oil—A Plausible Alternate Resource for Food and Fuel Application. *Bioresour. Technol.* 2017, 233, 423–432. [CrossRef]
- Kwak, W.S.; Kang, J.S. Effect of Feeding Food Waste-Broiler Litter and Bakery by-Product Mixture to Pigs. *Bioresour. Technol.* 2006, 97, 243–249. [CrossRef]
- 16. Almeida, F.N.; Petersen, G.I.; Stein, H.H. Digestibility of Amino Acids in Corn, Corn Coproducts, and Bakery Meal Fed to Growing Pigs. J. Anim. Sci. 2011, 89, 4109–4115. [CrossRef]
- 17. Rojas, O.J.; Liu, Y.; Stein, H.H. Phosphorus Digestibility and Concentration of Digestible and Metabolizable Energy in Corn, Corn Coproducts, and Bakery Meal Fed to Growing Pigs. *J. Anim. Sci.* **2013**, *91*, 5326–5335. [CrossRef]
- Luciano, A.; Espinosa, C.D.; Pinotti, L.; Stein, H.H. Standardized Total Tract Digestibility of Phosphorus in Bakery Meal Fed to Pigs and Effects of Bakery Meal on Growth Performance of Weanling Pigs. *Anim. Feed Sci. Technol.* 2022, 284, 115148. [CrossRef]
- Guo, J.Y.; Phillips, C.E.; Coffey, M.T.; Kim, S.W. Efficacy of a Supplemental Candy Coproduct as an Alternative Carbohydrate Source to Lactose on Growth Performance of Newly Weaned Pigs in a Commercial Farm Condition. *J. Anim. Sci.* 2015, 93, 5304–5312. [CrossRef]
- Neumann, C.; Velten, S.; Liebert, F. N Balance Studies Emphasize the Superior Protein Quality of Pig Diets at High Inclusion Level of Algae Meal (*Spirulina platensis*) or Insect Meal (*Hermetia illucens*) When Adequate Amino Acid Dupplementation Is Ensured. *Animals* 2018, *8*, 172. [CrossRef]
- Allegretti, G.; Talamini, E.; Schmidt, V.; Bogorni, P.C.; Ortega, E. Insect as Feed: An Emergy Assessment of Insect Meal as a Sustainable Protein Source for the Brazilian Poultry Industry. J. Clean. Prod. 2018, 171, 403–412. [CrossRef]
- 22. DiGiacomo, K.; Leury, B.J. Review: Insect Meal: A Future Source of Protein Feed for Pigs? Animal 2019, 13, 3022–3030. [CrossRef]
- 23. Hogg, S. Essential Microbiology, 2nd ed.; John Wiley & Sons, Inc.: Hoboken, NJ, USA, 2013; ISBN 978-1-119-97890-9.
- 24. Johannsen, L. Biological Properties of Bacterial Peptidoglycan. Apmis 1993, 101, 337–344. [CrossRef]
- 25. Pazos, M.; Peters, K. Peptidoglycan. In Subcellular Biochemistry; Springer: Berlin/Heidelberg, Germany, 2019; pp. 127–168.
- Van Amersfoort, E.S.; Kuiper, J. Receptors, Mediators, and Mechanisms Involved in Bacterial Sepsis and Septic Shock. In Endotoxins; CRC Press: Boca Raton, FL, USA, 2007; pp. 403–426. ISBN 9781420020595.
- 27. Duarte, M.E.; Tyus, J.; Kim, S.W. Synbiotic Effects of Enzyme and Probiotics on Intestinal Health and Growth of Newly Weaned Pigs Challenged with Enterotoxigenic F18+Escherichia Coli. *Front. Vet. Sci.* **2020**, *7*, 7–573. [CrossRef]
- 28. Bandara, T. Alternative Feed Ingredients in Aquaculture: Opportunities and Challenges. J. Entomol. Zool. Stud. 2018, 6, 3087–3094.
- 29. Sun, Y.; Duarte, M.E.; Kim, S.W. Dietary Inclusion of Multispecies Probiotics to Reduce the Severity of Post-Weaning Diarrhea Caused by Escherichia Coli F18+ in Pigs. *Anim. Nutr.* **2021**, *7*, 326–333. [CrossRef] [PubMed]
- Brown, A.J.; Goldsworthy, S.M.; Barnes, A.A.; Eilert, M.M.; Tcheang, L.; Daniels, D.; Muir, A.I.; Wigglesworth, M.J.; Kinghorn, I.; Fraser, N.J.; et al. The Orphan G Protein-Coupled Receptors GPR41 and GPR43 Are Activated by Propionate and Other Short Chain Carboxylic Acids. J. Biol. Chem. 2003, 278, 11312–11319. [CrossRef] [PubMed]
- Lukaszczyk, M.; Pradhan, B.; Remaut, H. Bacterial Cell Walls and Membranes; Kuhn, A., Ed.; Subcellular Biochemistry; Springer International Publishing: Cham, Switzerland, 2019; Volume 92, ISBN 978-3-030-18767-5.
- 32. Morphology, G. Yeast Cell Architecture and Functions. In Yeast; Wiley: Weinheim, Germany, 2012; pp. 5–24. ISBN 9783527332526.
- 33. Pacheco, M.T.B.; Caballero-Cordoba, G.M.; Sgarbieri, V.C. Composition and Nutritive Value of Yeast Biomass and Yeast Protein Concentrates. *J. Nutr. Sci. Vitaminol.* **1997**, *43*, 601–612. [CrossRef] [PubMed]
- Becker, W.; Richmond, A. Handbook of Microalgal Culture; Richmond, A., Ed.; Blackwell Publishing Ltd.: Oxford, UK, 2003; ISBN 9780470995280.
- de Jesus Raposo, M.; de Morais, A.; de Morais, R. Emergent Sources of Prebiotics: Seaweeds and Microalgae. Mar. Drugs 2016, 14, 27. [CrossRef]
- 36. Kay, R.A.; Barton, L.L. Microalgae as Food and Supplement. Crit. Rev. Food Sci. Nutr. 1991, 30, 555–573. [CrossRef]
- 37. Koch, A.L. *The Bacteria: Their Origin, Structure, Function and Antibiosis;* Springer: Dordrecht, The Netherlands, 2006; ISBN 978-1-4020-6625-2.
- Kang, S.-S.; Sim, J.-R.; Yun, C.-H.; Han, S.H. Lipoteichoic Acids as a Major Virulence Factor Causing Inflammatory Responses via Toll-like Receptor. Arch. Pharm. Res. 2016, 39, 1519–1529. [CrossRef]
- Stewart, G.G. Energy Metabolism by the Yeast Cell. In *Brewing and Distilling Yeasts*; Springer International Publishing: Cham, Switzerland, 2017; pp. 77–107.

- 40. Stewart, G.G. The Structure and Function of the Yeast Cell Wall, Plasma Membrane and Periplasm. In *Brewing and Distilling Yeasts;* Springer International Publishing: Cham, Switzerland, 2017; pp. 55–75. ISBN 9783319691244.
- 41. Shurson, G.C. Yeast and Yeast Derivatives in Feed Additives and Ingredients: Sources, Characteristics, Animal Responses, and Quantification Methods. *Anim. Feed Sci. Technol.* **2018**, 235, 60–76. [CrossRef]
- 42. Kim, S.W.; Holanda, D.M.; Gao, X.; Park, I.; Yiannikouris, A. Efficacy of a Yeast Cell Wall Extract to Mitigate the Effect of Naturally Co-Occurring Mycotoxins Contaminating Feed Ingredients Fed to Young Pigs: Impact on Gut Health, Microbiome, and Growth. *Toxins* 2019, *11*, 633. [CrossRef]
- 43. Shen, Y.B.; Piao, X.S.; Kim, S.W.; Wang, L.; Liu, P.; Yoon, I.; Zhen, Y.G. Effects of Yeast Culture Supplementation on Growth Performance, Intestinal Health, and Immune Response of Nursery Pigs. J. Anim. Sci. 2009, 87, 2614–2624. [CrossRef]
- LeMieux, F.M.; Southern, L.L.; Bidner, T.D. Effect of Mannan Oligosaccharides on Growth Performance of Weanling Pigs1. J. Anim. Sci. 2003, 81, 2482–2487. [CrossRef]
- Lipke, P.N.; Ovalle, R. Cell Wall Architecture in Yeast: New Structure and New Challenges. J. Bacteriol. 1998, 180, 3735–3740. [CrossRef]
- Kogan, G.; Pajtinka, M.; Babincova, M.; Miadokova, E.; Rauko, P.; Slamenova, D.; Korolenko, T.A. Yeast Cell Wall Polysaccharides as Antioxidants and Antimutagens: Can They Fight Cancer? *Neoplasma* 2008, 55, 387–393.
- 47. Baudelet, P.-H.; Ricochon, G.; Linder, M.; Muniglia, L. A New Insight into Cell Walls of Chlorophyta. *Algal Res.* 2017, 25, 333–371. [CrossRef]
- 48. Da Silva, J.C.; Lombardi, A.T. Chlorophylls in Microalgae: Occurrence, Distribution, and Biosynthesis. In *Pigments from Microalgae Handbook*; Springer International Publishing: Cham, Switzerland, 2020; pp. 1–18. ISBN 9783030509705.
- 49. Bold, H.C.; Wynne, M.J. Introduction to the Algae: Structure and Reproduction, 2nd ed.Prentice-Hall: Englewood Cliffs, NJ, USA, 1985; ISBN 9780134777467.
- 50. Metting, F.B. Biodiversity and Application of Microalgae. J. Ind. Microbiol. Biotechnol. 1996, 17, 477–489. [CrossRef]
- 51. Austic, R.E.; Mustafa, A.; Jung, B.; Gatrell, S.; Lei, X.G. Potential and Limitation of a New Defatted Diatom Microalgal Biomass in Replacing Soybean Meal and Corn in Diets for Broiler Chickens. *J. Agric. Food Chem.* **2013**, *61*, 7341–7348. [CrossRef]
- 52. Kim, J.; Magnuson, A.; Tao, L.; Barcus, M.; Lei, X.G. Potential of Combining Flaxseed Oil and Microalgal Biomass in Producing Eggs-Enriched with n 3 Fatty Acids for Meeting Human Needs. *Algal Res.* **2016**, *17*, 31–37. [CrossRef]
- 53. Yi, Z.; Xu, M.; Di, X.; Brynjolfsson, S.; Fu, W. Exploring Valuable Lipids in Diatoms. Front. Mar. Sci. 2017, 4, 17. [CrossRef]
- Velazquez-Lucio, J.; Rodríguez-Jasso, R.M.; Colla, L.M.; Sáenz-Galindo, A.; Cervantes-Cisneros, D.E.; Aguilar, C.N.; Fernandes, B.D.; Ruiz, H.A. Microalgal Biomass Pretreatment for Bioethanol Production: A Review. *Biofuel Res. J.* 2018, 5, 780–791. [CrossRef]
- 55. Pôjo, V.; Tavares, T.; Malcata, F.X. Processing Methodologies of Wet Microalga Biomass toward Oil Separation: An Overview. *Molecules* **2021**, *26*, 641. [CrossRef]
- 56. Kihlberg, R. The Microbe as a Source of Food. Annu. Rev. Microbiol. 1972, 26, 427–466. [CrossRef] [PubMed]
- 57. Bozell, J.J.; Petersen, G.R. Technology Development for the Production of Biobased Products from Biorefinery Carbohydrates—the US Department of Energy's "Top 10" Revisited. *Green Chem.* **2010**, *12*, 539. [CrossRef]
- Babson, D.M.; Held, M.; Schmidt-Dannert, C. Designer Microbial Ecosystems—Toward Biosynthesis with Engineered Microbial Consortia. In *Natural Products*; John Wiley & Sons, Inc.: Hoboken, NJ, USA, 2014; pp. 23–38.
- Namioka, S.; Sasaki, T.; Maede, Y. Immunopotentiation of the Small Intestine of Weaning Piglets by Peptidoglycan Derived from Bifidobacterium Thermophilum. *Bifidobact. Microflora* 1991, 10, 1–9. [CrossRef]
- 60. Kogan, G.; Kocher, A. Role of Yeast Cell Wall Polysaccharides in Pig Nutrition and Health Protection. *Livest. Sci.* 2007, 109, 161–165. [CrossRef]
- 61. Puri, M.; Thyagarajan, T.; Gupta, A.; Barrow, C.J. Omega-3 Fatty Acids Produced from Microalgae. In *Hb25_Springer Handbook of Marine Biotechnology*; Springer: Berlin/Heidelberg, Germany, 2015; pp. 1043–1057.
- 62. Salzman, N.H. Microbiota–Immune System Interaction: An Uneasy Alliance. Curr. Opin. Microbiol. 2011, 14, 99–105. [CrossRef]
- Lallès, J.-P.; Bosi, P.; Smidt, H.; Stokes, C.R. Nutritional Management of Gut Health in Pigs around Weaning. Proc. Nutr. Soc. 2007, 66, 260–268. [CrossRef]
- 64. Duarte, M.E.; Kim, S.W. Intestinal Microbiota and Its Interaction to Intestinal Health in Nursery Pigs. *Anim. Nutr.* **2022**, *8*, 169–184. [CrossRef]
- Furbeyre, H.; van Milgen, J.; Mener, T.; Gloaguen, M.; Labussière, E. Effects of Dietary Supplementation with Freshwater Microalgae on Growth Performance, Nutrient Digestibility and Gut Health in Weaned Piglets. *Animal* 2017, *11*, 183–192. [CrossRef]
- Iakhno, S.; Umu, Ö.C.O.; Håkenåsen, I.M.; Åkesson, C.P.; Mydland, L.T.; Press, C.M.; Sørum, H.; Øverland, M. Effect of Cyberlindnera Jadinii Yeast as a Protein Source on Intestinal Microbiota and Butyrate Levels in Post-Weaning Piglets. *Anim. Microbiome* 2020, 2, 13. [CrossRef]
- 67. Broom, L.J.; Miller, H.M.; Kerr, K.G.; Knapp, J.S. Effects of Zinc Oxide and Enterococcus Faecium SF68 Dietary Supplementation on the Performance, Intestinal Microbiota and Immune Status of Weaned Piglets. *Res. Vet. Sci.* 2006, *80*, 45–54. [CrossRef]
- Monger, X.C.; Gilbert, A.-A.; Saucier, L.; Vincent, A.T. Antibiotic Resistance: From Pig to Meat. *Antibiotics* 2021, 10, 1209. [CrossRef] [PubMed]
- 69. Butaye, P.; Devriese, L.A.; Haesebrouck, F. Antimicrobial Growth Promoters Used in Animal Feed: Effects of Less Well Known Antibiotics on Gram-Positive Bacteria. *Clin. Microbiol. Rev.* **2003**, *16*, 175–188. [CrossRef]

- 70. Kenny, M.; Smidt, H.; Mengheri, E.; Miller, B. Probiotics—Do They Have a Role in the Pig Industry? *Animal* 2011, *5*, 462–470. [CrossRef]
- Teame, T.; Wang, A.; Xie, M.; Zhang, Z.; Yang, Y.; Ding, Q.; Gao, C.; Olsen, R.E.; Ran, C.; Zhou, Z. Paraprobiotics and Postbiotics of Probiotic Lactobacilli, Their Positive Effects on the Host and Action Mechanisms: A Review. *Front. Nutr.* 2020, 7, 344. [CrossRef]
- De Marzi, M.C.; Todone, M.; Ganem, M.B.; Wang, Q.; Mariuzza, R.A.; Fernández, M.M.; Malchiodi, E.L. Peptidoglycan Recognition Protein-Peptidoglycan Complexes Increase Monocyte/Macrophage Activation and Enhance the Inflammatory Response. *Immunology* 2015, 145, 429–442. [CrossRef]
- Kim, K.W.; Kang, S.S.; Woo, S.J.; Park, O.J.; Ahn, K.B.; Song, K.D.; Lee, H.K.; Yun, C.H.; Han, S.H. Lipoteichoic Acid of Probiotic Lactobacillus Plantarum Attenuates Poly I: C-Induced IL-8 Production in Porcine Intestinal Epithelial Cells. *Front. Microbiol.* 2017, *8*, 1827. [CrossRef]
- 74. Irazoki, O.; Hernandez, S.B.; Cava, F. Peptidoglycan Muropeptides: Release, Perception, and Functions as Signaling Molecules. *Front. Microbiol.* **2019**, *10*, 500. [CrossRef]
- 75. Sorbara, M.T.; Philpott, D.J. Peptidoglycan: A Critical Activator of the Mammalian Immune System during Infection and Homeostasis. *Immunol. Rev.* 2011, 243, 40–60. [CrossRef] [PubMed]
- Peters, M.; Guidato, P.M.; Peters, K.; Megger, D.A.; Sitek, B.; Classen, B.; Heise, E.M.; Bufe, A. Allergy-Protective Arabinogalactan Modulates Human Dendritic Cells via C-Type Lectins and Inhibition of NF-KB. J. Immunol. 2016, 196, 1626–1635. [CrossRef] [PubMed]
- Caruso, A.; Flamminio, G.; Folghera, S.; Peroni, L.; Foresti, I.; Balsari, A.; Turano, A. Expression of Activation Markers on Peripheral-Blood Lymphocytes Following Oral Administration of Bacillus Subtilis Spores. *Int. J. Immunopharmacol.* 1993, 15, 87–92. [CrossRef]
- Poulsen, A.-S.R.; de Jonge, N.; Nielsen, J.L.; Højberg, O.; Lauridsen, C.; Cutting, S.M.; Canibe, N. Impact of Bacillus Spp. Spores and Gentamicin on the Gastrointestinal Microbiota of Suckling and Newly Weaned Piglets. *PLoS ONE* 2018, 13, e0207382. [CrossRef] [PubMed]
- Suda, Y.; Villena, J.; Takahashi, Y.; Hosoya, S.; Tomosada, Y.; Tsukida, K.; Shimazu, T.; Aso, H.; Tohno, M.; Ishida, M.; et al. Immunobiotic Lactobacillus Jensenii as Immune-Health Promoting Factor to Improve Growth Performance and Productivity in Post-Weaning Pigs. *BMC Immunol.* 2014, 15, 24. [CrossRef]
- Taras, D.; Vahjen, W.; Macha, M.; Simon, O. Response of Performance Characteristics and Fecal Consistency to Long-Lasting Dietary Supplementation with the Probiotic Strain Bacillus Cereus Var. Toyoi to Sows and Piglets. *Arch. Anim. Nutr.* 2005, 59, 405–417. [CrossRef]
- 81. He, Y.; Jinno, C.; Kim, K.; Wu, Z.; Tan, B.; Li, X.; Whelan, R.; Liu, Y. Dietary *Bacillus* Spp. Enhanced Growth and Disease Resistance of Weaned Pigs by Modulating Intestinal Microbiota and Systemic Immunity. *J. Anim. Sci. Biotechnol.* **2020**, *11*, 101. [CrossRef]
- Pessione, E. Lactic Acid Bacteria Contribution to Gut Microbiota Complexity: Lights and Shadows. Front. Cell. Infect. Microbiol. 2012, 2, 86. [CrossRef]
- Taras, D.; Vahjen, W.; Macha, M.; Simon, O. Performance, Diarrhea Incidence, and Occurrence of Escherichia Coli Virulence Genes during Long-Term Administration of a Probiotic Enterococcus Faecium Strain to Sows and Piglets. J. Anim. Sci. 2006, 84, 608–617. [CrossRef]
- 84. Pollmann, M.; Nordhoff, M.; Pospischil, A.; Tedin, K.; Wieler, L.H. Effects of a Probiotic Strain of Enterococcus Faecium on the Rate of Natural Chlamydia Infection in Swine. *Infect. Immun.* **2005**, *73*, 4346–4353. [CrossRef]
- Jørgensen, J.N.; Laguna, J.S.; Millán, C.; Casabuena, O.; Gracia, M.I. Effects of a Bacillus-Based Probiotic and Dietary Energy Content on the Performance and Nutrient Digestibility of Wean to Finish Pigs. Anim. Feed Sci. Technol. 2016, 221, 54–61. [CrossRef]
- Prenafeta-Boldú, F.X.; Fernández, B.; Viñas, M.; Lizardo, R.; Brufau, J.; Owusu-Asiedu, A.; Walsh, M.C.; Awati, A. Effect of Bacillus Spp. Direct-Fed Microbial on Slurry Characteristics and Gaseous Emissions in Growing Pigs Fed with High Fibre-Based Diets. *Animal* 2017, *11*, 209–218. [CrossRef]
- Lee, S.H.; Ingale, S.L.; Kim, J.S.; Kim, K.H.; Lokhande, A.; Kim, E.K.; Kwon, I.K.; Kim, Y.H.; Chae, B.J. Effects of Dietary Supplementation with Bacillus Subtilis LS 1–2 Fermentation Biomass on Growth Performance, Nutrient Digestibility, Cecal Microbiota and Intestinal Morphology of Weanling Pig. *Anim. Feed Sci. Technol.* 2014, 188, 102–110. [CrossRef]
- Sonia, T.A.; Ji, H.; Hong-Seok, M.; Chul-Ju, Y. Evaluation of Lactobacillus and Bacillus-Based Probiotics as Alternatives to Antibiotics in Enteric Microbial Challenged Weaned Piglets. *Afr. J. Microbiol. Res.* 2014, *8*, 96–104. [CrossRef]
- 89. Zhao, X.; Guo, Y.; Guo, S.; Tan, J. Effects of *Clostridium Butyricum* and *Enterococcus Faecium* on Growth Performance, Lipid Metabolism, and Cecal Microbiota of Broiler Chickens. *Appl. Microbiol. Biotechnol.* **2013**, *97*, 6477–6488. [CrossRef] [PubMed]
- Mohana Devi, S.; Kim, I. Effect of Medium Chain Fatty Acids (MCFA) and Probiotic (*Enterococcus faecium*) Supplementation on the Growth Performance, Digestibility and Blood Profiles in Weanling Pigs. *Vet. Med.* 2014, 59, 527–535. [CrossRef]
- Xu, X.; Duarte, M.E.; Kim, S.W. Postbiotic Effects of Lactobacillus Fermentate on Intestinal Health, Mucosa-Associated Microbiota, and Growth Efficiency of Nursery Pigs Challenged with F18+ *Escherichia coli*. J. Anim. Sci. 2022, 100, skac210. [CrossRef] [PubMed]
- 92. Veizaj-Delia, E.; Piu, T.; Lekaj, P.; Tafaj, M. Using Combined Probiotic to Improve Growth Performance of Weaned Piglets on Extensive Farm Conditions. *Livest. Sci.* 2010, 134, 249–251. [CrossRef]
- 93. Huang, C.; Qiao, S.; Li, D.; Piao, X.; Ren, J. Effects of Lactobacilli on the Performance, Diarrhea Incidence, VFA Concentration and Gastrointestinal Microbial Flora of Weaning Pigs. *Asian-Australas. J. Anim. Sci.* **2004**, *17*, 401–409. [CrossRef]

- de Graaff, P.; Govers, C.; Wichers, H.J.; Debets, R. Consumption of β-Glucans to Spice up T Cell Treatment of Tumors: A Review. Expert Opin. Biol. Ther. 2018, 18, 1023–1040. [CrossRef]
- 95. Gallois, M.; Rothkötter, H.J.; Bailey, M.; Stokes, C.R.; Oswald, I.P. Natural Alternatives to In-Feed Antibiotics in Pig Production: Can Immunomodulators Play a Role? *Animal* **2009**, *3*, 1644–1661. [CrossRef]
- 96. Akira, S.; Uematsu, S.; Takeuchi, O. Pathogen Recognition and Innate Immunity. Cell 2006, 124, 783-801. [CrossRef]
- 97. Li, T.-H.; Liu, L.; Hou, Y.-Y.; Shen, S.-N.; Wang, T.-T. C-Type Lectin Receptor-Mediated Immune Recognition and Response of the Microbiota in the Gut. *Gastroenterol. Rep.* 2019, *7*, 312–321. [CrossRef]
- Anwar, M.I.; Muhammad, F.; Awais, M.M.; Akhtar, M. A Review of β-Glucans as a Growth Promoter and Antibiotic Alternative against Enteric Pathogens in Poultry. *World's Poult. Sci. J.* 2017, 73, 651–661. [CrossRef]
- Li, J.; Li, D.F.; Xing, J.J.; Cheng, Z.B.; Lai, C.H. Effects of β-Glucan Extracted from Saccharomyces Cerevisiae on Growth Performance, and Immunological and Somatotropic Responses of Pigs Challenged with Escherichia Coli Lipopolysaccharide. J. Anim. Sci. 2006, 84, 2374–2381. [CrossRef]
- Price, K.L.; Totty, H.R.; Lee, H.B.; Utt, M.D.; Fitzner, G.E.; Yoon, I.; Ponder, M.A.; Escobar, J. Use of Saccharomyces Cerevisiae Fermentation Product on Growth Performance and Microbiota of Weaned Pigs during Salmonella Infection. *J. Anim. Sci.* 2010, *88*, 3896–3908. [CrossRef]
- Kim, S.W.; Brandherm, M.; Freeland, M.; Newton, B.; Cook, D.; Yoon, I. Effects of Yeast Culture Supplementation to Gestation and Lactation Diets on Growth of Nursing Piglets. *Asian-Australas. J. Anim. Sci.* 2008, 21, 1011–1014. [CrossRef]
- Kim, S.W.; Brandherm, M.; Newton, B.; Cook, D.R.; Yoon, I.; Fitzner, G. Effect of Supplementing Saccharomyces Cerevisiae Fermentation Product in Sow Diets on Reproductive Performance in a Commercial Environment. *Can. J. Anim. Sci.* 2010, 90, 229–232. [CrossRef]
- 103. Shen, Y.B.; Carroll, J.A.; Yoon, I.; Mateo, R.D.; Kim, S.W. Effects of Supplementing *Saccharomyces Cerevisiae* Fermentation Product in Sow Diets on Performance of Sows and Nursing Piglets. *J. Anim. Sci.* **2011**, *89*, 2462–2471. [CrossRef]
- Lu, H.; Wilcock, P.; Adeola, O.; Ajuwon, K.M. Effect of Live Yeast Supplementation to Gestating Sows and Nursery Piglets on Postweaning Growth Performance and Nutrient Digestibility. J. Anim. Sci. 2019, 97, 2534–2540. [CrossRef]
- Davis, M.E.; Maxwell, C.V.; Brown, D.C.; de Rodas, B.Z.; Johnson, Z.B.; Kegley, E.B.; Hellwig, D.H.; Dvorak, R.A. Effect of Dietary Mannan Oligosaccharides and(or) Pharmacological Additions of Copper Sulfate on Growth Performance and Immunocompetence of Weanling and Growing/Finishing Pigs. J. Anim. Sci. 2002, 80, 2887–2894. [CrossRef]
- 106. Onwurah, F. Effect of Baker's Yeast (Saccharomyces cerevisiae) Inclusion in Feed and in Drinking Water on Performance of Broiler Birds. Br. J. Appl. Sci. Technol. 2014, 4, 144–151. [CrossRef]
- 107. Zhang, A.W.; Lee, B.D.; Lee, S.K.; Lee, K.W.; An, G.H.; Song, K.B.; Lee, C.H. Effects of Yeast (*Saccharomyces cerevisiae*) Cell Components on Growth Performance, Meat Quality, and Ileal Mucosa Development of Broiler Chicks. *Poult. Sci.* 2005, 84, 1015–1021. [CrossRef]
- 108. Sauerwein, H.; Schmitz, S.; Hiss, S. Effects of a Dietary Application of a Yeast Cell Wall Extract on Innate and Acquired Immunity, on Oxidative Status and Growth Performance in Weanling Piglets and on the Ileal Epithelium in Fattened Pigs. J. Anim. Physiol. Anim. Nutr. 2007, 91, 369–380. [CrossRef]
- 109. Mathew, A.G.; Chattin, S.E.; Robbins, C.M.; Golden, D.A. Effects of a Direct-Fed Yeast Culture on Enteric Microbial Populations, Fermentation Acids, and Performance of Weanling Pigs. *J. Anim. Sci.* **1998**, *76*, 2138. [CrossRef] [PubMed]
- Park, J.H.; Lee, S.I.; Kim, I.H. Effect of Dietary Spirulina (*Arthrospira*) Platensis on the Growth Performance, Antioxidant Enzyme Activity, Nutrient Digestibility, Cecal Microflora, Excreta Noxious Gas Emission, and Breast Meat Quality of Broiler Chickens. *Poult. Sci.* 2018, 97, 2451–2459. [CrossRef] [PubMed]
- Moran, C.A.; Morlacchini, M.; Keegan, J.D.; Fusconi, G. Dietary Supplementation of Finishing Pigs with the Docosahexaenoic Acid-Rich Microalgae, *Aurantiochytrium Limacinum*: Effects on Performance, Carcass Characteristics and Tissue Fatty Acid Profile. *Asian-Australas. J. Anim. Sci.* 2018, 31, 712–720. [CrossRef] [PubMed]
- Waldenstedt, L.; Inborr, J.; Hansson, I.; Elwinger, K. Effects of Astaxanthin-Rich Algal Meal (*Haematococcus pluvalis*) on Growth Performance, Caecal Campylobacter and Clostridial Counts and Tissue Astaxanthin Concentration of Broiler Chickens. *Anim. Feed Sci. Technol.* 2003, 108, 119–132. [CrossRef]
- 113. Kibria, S.; Kim, I.H. Impacts of Dietary Microalgae (*Schizochytrium* JB5) on Growth Performance, Blood Profiles, Apparent Total Tract Digestibility, and Ileal Nutrient Digestibility in Weaning Pigs. *J. Sci. Food Agric.* **2019**, *99*, 6084–6088. [CrossRef]
- 114. Raja, R.; Hemaiswarya, S.; Kumar, N.A.; Sridhar, S.; Rengasamy, R. A Perspective on the Biotechnological Potential of Microalgae. *Crit. Rev. Microbiol.* **2008**, *34*, 77–88. [CrossRef]
- 115. Mason, R.P.; Libby, P.; Bhatt, D.L. Emerging Mechanisms of Cardiovascular Protection for the Omega-3 Fatty Acid Eicosapentaenoic Acid. *Arterioscler. Thromb. Vasc. Biol.* 2020, 40, 1135–1147. [CrossRef]
- 116. Sardi, L.; Martelli, G.; Lambertini, L.; Parisini, P.; Mordenti, A. Effects of a Dietary Supplement of DHA-Rich Marine Algae on Italian Heavy Pig Production Parameters. *Livest. Sci.* 2006, 103, 95–103. [CrossRef]
- 117. Baňoch, T.; Svoboda, M.; Kuta, J.; Saláková, A.; Fajt, Z. The Effect of Iodine from Iodine-Enriched Alga Chlorella Spp. on the Pork Iodine Content and Meat Quality in Finisher Pigs. *Acta Vet. Brno* **2012**, *81*, 339–346. [CrossRef]
- 118. Pourkarimi, S.; Hallajisani, A.; Alizadehdakhel, A.; Nouralishahi, A.; Golzary, A. Factors Affecting Production of Beta-Carotene from Dunaliella Salina Microalgae. *Biocatal. Agric. Biotechnol.* **2020**, *29*, 101771. [CrossRef]

- 119. Brito, A.d.F.; Silva, A.S.; de Oliveira, C.V.C.; de Souza, A.A.; Ferreira, P.B.; de Souza, I.L.L.; da Cunha Araujo, L.C.; da Silva Félix, G.; de Souza Sampaio, R.; Tavares, R.L.; et al. Spirulina Platensis Prevents Oxidative Stress and Inflammation Promoted by Strength Training in Rats: Dose-Response Relation Study. *Sci. Rep.* 2020, *10*, 6382. [CrossRef]
- 120. Hussain, T.; Tan, B.; Yin, Y.; Blachier, F.; Tossou, M.C.B.; Rahu, N. Oxidative Stress and Inflammation: What Polyphenols Can Do for Us? *Oxid. Med. Cell. Longev.* 2016, 2016, 7432797. [CrossRef]
- Kiros, T.G.; Derakhshani, H.; Pinloche, E.; D'Inca, R.; Marshall, J.; Auclair, E.; Khafipour, E.; Van Kessel, A. Effect of Live Yeast Saccharomyces Cerevisiae (Actisaf Sc 47) Supplementation on the Performance and Hindgut Microbiota Composition of Weanling Pigs. Sci. Rep. 2018, 8, 5315. [CrossRef]
- 122. den Besten, G.; van Eunen, K.; Groen, A.K.; Venema, K.; Reijngoud, D.-J.; Bakker, B.M. The Role of Short-Chain Fatty Acids in the Interplay between Diet, Gut Microbiota, and Host Energy Metabolism. *J. Lipid Res.* **2013**, *54*, 2325–2340. [CrossRef]
- 123. Jiao, A.; Diao, H.; Yu, B.; He, J.; Yu, J.; Zheng, P.; Luo, Y.; Luo, J.; Wang, Q.; Wang, H.; et al. Infusion of Short Chain Fatty Acids in the Ileum Improves the Carcass Traits, Meat Quality and Lipid Metabolism of Growing Pigs. *Anim. Nutr.* 2021, 7, 94–100. [CrossRef]
- 124. Lum, K.K.; Kim, J.; Lei, X.G. Dual Potential of Microalgae as a Sustainable Biofuel Feedstock and Animal Feed. J. Anim. Sci. Biotechnol. 2013, 4, 53. [CrossRef]
- 125. Taliercio, E.; Kim, S.W. Epitopes from Two Soybean Glycinin Subunits Are Antigenic in Pigs. J. Sci. Food Agric. 2013, 93, 2927–2932. [CrossRef]
- 126. Bosi, P.; Han, I.K.; Jung, H.J.; Heo, K.N.; Perini, S.; Castellazzi, A.M.; Casini, L.; Creston, D.; Gremokolini, C. Effect of Different Spray Dried Plasmas on Growth, Ileal Digestibility, Nutrient Deposition, Immunity and Health of Early-Weaned Pigs Challenged with E. Coli K88. Asian-Australas. J. Anim. Sci. 2001, 14, 1138–1143. [CrossRef]
- 127. Olsen, R.L.; Hasan, M.R. A Limited Supply of Fishmeal: Impact on Future Increases in Global Aquaculture Production. *Trends Food Sci. Technol.* **2012**, *27*, 120–128. [CrossRef]
- Nasseri, A.T.; Rasoul-Ami, S.; Morowvat, M.H.; Ghasemi, Y. Single Cell Protein: Production and Process. Am. J. Food Technol. 2011, 6, 103–116. [CrossRef]
- 129. Becker, P.M. Single Cell Proteins in Diets for Weanling Pigs; Animal Sciences Group: Wageningen, The Netherlands, 2014.
- Gatrell, S.; Lum, K.; Kim, J.; Lei, X.G. Nonruminant Nutrition Symposium: Potential of Defatted Microalgae from the Biofuel Industry as an Ingredient to Replace Corn and Soybean Meal in Swine and Poultry Diets. J. Anim. Sci. 2014, 92, 1306–1314. [CrossRef] [PubMed]
- Øverland, M.; Skrede, A.; Matre, T. Bacterial Protein Grown on Natural Gas as Feed for Pigs. *Acta Agric. Scand. Sect. A Anim. Sci.* 2001, 51, 97–106. [CrossRef]
- An, B.-K.; Choi, Y.-I.; Kang, C.-W.; Lee, K.-W. Effects of Dietary *Corynebacterium Ammoniagenes*-Derived Single Cell Protein on Growth Performance, Blood and Tibia Bone Characteristics, and Meat Quality of Broiler Chickens. *J. Anim. Feed Sci.* 2018, 27, 140–147. [CrossRef]
- Cruz, A.; Sterten, H.; Steinhoff, F.S.; Mydland, L.T.; Øverland, M. Cyberlindnera Jadinii Yeast as a Protein Source for Broiler Chickens: Effects on Growth Performance and Digestive Function from Hatching to 30 Days of Age. *Poult. Sci.* 2020, 99, 3168–3178. [CrossRef]
- Ekmay, R.; Gatrell, S.; Lum, K.; Kim, J.; Lei, X.G. Nutritional and Metabolic Impacts of a Defatted Green Marine Microalgal (Desmodesmus sp.) Biomass in Diets for Weanling Pigs and Broiler Chickens. J. Agric. Food Chem. 2014, 62, 9783–9791. [CrossRef]
- Øverland, M.; Kjos, N.P.; Olsen, E.; Skrede, A. Changes in Fatty Acid Composition and Improved Sensory Quality of Backfat and Meat of Pigs Fed Bacterial Protein Meal. *Meat Sci.* 2005, 71, 719–729. [CrossRef]
- 136. Moniruzzaman, M.; Mollah, M. Autolyzed Saccharomyces Cerevisiae as a Single Cell Protein for Broiler Diet. *Bangladesh J. Anim. Sci.* **2019**, *48*, 1–8. [CrossRef]
- 137. NRC. Nutrient Requirements of Swine, 11th ed.; National Academies Press: Washington, DC, USA, 2012; ISBN 978-0-309-22423-9.
- Zhang, H.Y.; Piao, X.S.; Li, P.; Yi, J.Q.; Zhang, Q.; Li, Q.Y.; Liu, J.D.; Wang, G.Q. Effects of Single Cell Protein Replacing Fish Meal in Diet on Growth Performance, Nutrient Digestibility and Intestinal Morphology in Weaned Pigs. *Asian-Australas. J. Anim. Sci.* 2013, 26, 1320–1328. [CrossRef]
- 139. Cheng, Y.-C.; Duarte, M.E.; Kim, S.W. Nutritional and Functional Values of Lysed *Corynebacterium Glutamicum* Cell Mass for Intestinal Health and Growth of Nursery Pigs. J. Anim. Sci. 2021, 99, skab331. [CrossRef]
- 140. D'Mello, J.P.F.; Acamovic, T. Evaluation of Methanol-grown Bacteria as a Source of Protein and Energy for Young Chicks. *Br. Poult. Sci.* **1976**, *17*, 393–401. [CrossRef]
- 141. Braude, R.; Rhodes, D.N. Pruteen, a New Source of Protein for Growing Pigs. II. Feeding Trial: Growth Rate, Feed Utilization and Carcass and Meat Quality. *Livest. Prod. Sci.* **1977**, *4*, 91–100. [CrossRef]
- 142. Schøyen, H.F.; Svihus, B.; Storebakken, T.; Skrede, A. Bacterial Protein Meal Produced on Natural Gas Replacing Soybean Meal or Fish Meal in Broiler Chicken Diets. *Arch. Anim. Nutr.* 2007, *61*, 276–291. [CrossRef]
- Øverland, M.; Tauson, A.-H.; Shearer, K.; Skrede, A. Evaluation of Methane-Utilising Bacteria Products as Feed Ingredients for Monogastric Animals. Arch. Anim. Nutr. 2010, 64, 171–189. [CrossRef]
- 144. Skrede, A.; Berge, G.; Storebakken, T.; Herstad, O.; Aarstad, K.; Sundstøl, F. Digestibility of Bacterial Protein Grown on Natural Gas in Mink, Pigs, Chicken and Atlantic Salmon. *Anim. Feed Sci. Technol.* **1998**, *76*, 103–116. [CrossRef]

- 145. Tlusty, M.; Rhyne, A.; Szczebak, J.T.; Bourque, B.; Bowen, J.L.; Burr, G.; Marx, C.J.; Feinberg, L. A Transdisciplinary Approach to the Initial Validation of a Single Cell Protein as an Alternative Protein Source for Use in Aquafeeds. *PeerJ* 2017, 5, e3170. [CrossRef]
- 146. Hellwing, A.L.F.; Tauson, A.-H.; Skrede, A.; Kjos, N.P.; Ahlstrøm, Ø. Bacterial Protein Meal in Diets for Pigs and Minks: Comparative Studies on Protein Turnover Rate and Urinary Excretion of Purine Base Derivatives. Arch. Anim. Nutr. 2007, 61, 425–443. [CrossRef]
- 147. Chen, Y.; Chi, S.; Zhang, S.; Dong, X.; Yang, Q.; Liu, H.; Zhang, W.; Deng, J.; Tan, B.; Xie, S. Replacement of Fish Meal with Methanotroph (*Methylococcus capsulatus*, Bath) Bacteria Meal in the Diets of Pacific White Shrimp (*Litopenaeus vannamei*). *Aquaculture* **2021**, *541*, 736801. [CrossRef]
- 148. Zhang, Q.; Liang, H.; Longshaw, M.; Wang, J.; Ge, X.; Zhu, J.; Li, S.; Ren, M. Effects of Replacing Fishmeal with Methanotroph (*Methylococcus capsulatus*, Bath) Bacteria Meal (FeedKind®) on Growth and Intestinal Health Status of Juvenile Largemouth Bass (*Micropterus salmoides*). Fish Shellfish Immunol. 2022, 122, 298–305. [CrossRef]
- 149. Plavnik, I.; Bornstein, S.; Hurwitz, S. Evaluation of Methanol-grown Bacteria and Hydrocarbon-grown Yeast as Sources of Protein for Poultry: Studies with Young Chicks. *Br. Poult. Sci.* **1981**, *22*, 123–140. [CrossRef]
- Sahm, H.; Eggeling, L. New Ubiquitous Translocators: Amino Acid Export by Corynebacterium Glutamicum and Escherichia Coli. Arch. Microbiol. 2003, 180, 155–160. [CrossRef] [PubMed]
- 151. Leuchtenberger, W.; Huthmacher, K.; Drauz, K. Biotechnological Production of Amino Acids and Derivatives: Current Status and Prospects. *Appl. Microbiol. Biotechnol.* **2005**, *69*, 1–8. [CrossRef] [PubMed]
- Yang, F.; Wang, A.; Zeng, X.; Hou, C.; Liu, H.; Qiao, S. Lactobacillus Reuteri I5007 Modulates Tight Junction Protein Expression in IPEC-J2 Cells with LPS Stimulation and in Newborn Piglets under Normal Conditions. *BMC Microbiol.* 2015, 15, 32. [CrossRef] [PubMed]
- Wang, J.P.; Kim, J.D.; Kim, J.E.; Kim, I.H. Amino Acid Digestibility of Single Cell Protein from Corynebacterium Ammoniagenes in Growing Pigs. Anim. Feed Sci. Technol. 2013, 180, 111–114. [CrossRef]
- 154. Whittemore, C.T.; Moffat, I.W.; Taylor, A.G. Evaluation by Digestibility, Growth and Slaughter of Microbial Cells as a Source of Protein for Young Pigs. J. Sci. Food Agric. 1976, 27, 1163–1170. [CrossRef]
- 155. Yáñez, E.; Ballester, D.; Fernández, N.; Gattás, V.; Monckeberg, F. Chemical Composition of Candida Utilis and the Biological Quality of the Yeast Protein. *J. Sci. Food Agric.* **1972**, 23, 581–586. [CrossRef]
- 156. Lagos, L.V.; Stein, H.H. Torula Yeast Has Greater Digestibility of Amino Acids and Phosphorus, but Not Energy, Compared with a Commercial Source of Fish Meal Fed to Weanling Pigs. J. Anim. Sci. 2020, 98, skz375. [CrossRef]
- 157. Reed, G.; Nagodawithana, T.W. Yeast Technology; Van Nostrand Reinhold: New York, NY, USA, 1991.
- 158. Chand, N.; Khan, R.U. Replacement of Soybean Meal with Yeast Single Cell Protein in Broiler Ration: The Effect on Performance Traits. *Pak. J. Zool.* 2014, *46*, 1753–1758.
- Lagos, L.; Bekkelund, A.K.; Skugor, A.; Ånestad, R.; Åkesson, C.P.; Press, C.M.; Øverland, M. Cyberlindnera Jadinii Yeast as a Protein Source for Weaned Piglets—Impact on Immune Response and Gut Microbiota. Front. Immunol. 2020, 11, 1924. [CrossRef]
- Cruz, A.; Håkenåsen, I.M.; Skugor, A.; Mydland, L.T.; Åkesson, C.P.; Hellestveit, S.S.; Sørby, R.; Press, C.M.; Øverland, M. Candida Utilis Yeast as a Protein Source for Weaned Piglets: Effects on Growth Performance and Digestive Function. *Livest. Sci.* 2019, 226, 31–39. [CrossRef]
- van Heugten, E.; Funderburke, D.W.; Dorton, K.L. Growth Performance, Nutrient Digestibility, and Fecal Microflora in Weanling Pigs Fed Live Yeast. J. Anim. Sci. 2003, 81, 1004–1012. [CrossRef]
- 162. Kim, B.G.; Liu, Y.; Stein, H.H. Energy Concentration and Phosphorus Digestibility in Yeast Products Produced from the Ethanol Industry, and in Brewers' Yeast, Fish Meal, and Soybean Meal Fed to Growing Pigs1. J. Anim. Sci. 2014, 92, 5476–5484. [CrossRef]
- Valdivie, M.; Compte, X.; Fundora, O. The Utilization of Torula Yeast in Diets for White Leghorn Birds during Growth and Laying Periods. *Anim. Feed Sci. Technol.* 1982, 7, 185–190. [CrossRef]
- Espinosa, C.D.; Lagos, L.V.; Stein, H.H. Effect of Torula Yeast on Growth Performance, Diarrhea Incidence, and Blood Characteristics in Weanling Pigs. J. Anim. Sci. 2020, 98, skaa307. [CrossRef]
- Czech, A.; Smolczyk, A.; Ognik, K.; Kiesz, M. Nutritional Value of Yarrowia Lipolytica Yeast and Its Effect on Growth Performance Indicators in Piglets. *Ann. Anim. Sci.* 2016, 16, 1091–1100. [CrossRef]
- 166. Lipstein, B.; Hurwitz, S. The Nutritional Value of Algae for Poultry. Dried Chlorella in Broiler Diets. *Br. Poult. Sci.* **1980**, *21*, 9–21. [CrossRef]
- 167. Coelho, D.; Pestana, J.; Almeida, J.M.; Alfaia, C.M.; Fontes, C.M.G.A.; Moreira, O.; Prates, J.A.M. A High Dietary Incorporation Level of Chlorella Vulgaris Improves the Nutritional Value of Pork Fat without Impairing the Performance of Finishing Pigs. *Animals* 2020, 10, 2384. [CrossRef]
- 168. Alfaia, C.M.; Pestana, J.M.; Rodrigues, M.; Coelho, D.; Aires, M.J.; Ribeiro, D.M.; Major, V.T.; Martins, C.F.; Santos, H.; Lopes, P.A.; et al. Influence of Dietary Chlorella Vulgaris and Carbohydrate-Active Enzymes on Growth Performance, Meat Quality and Lipid Composition of Broiler Chickens. *Poult. Sci.* 2021, 100, 926–937. [CrossRef]
- Gatrell, S.K.; Derksen, T.J.; O'Neil, E.V.; Lei, X.G. A New Type of Defatted Green Microalgae Exerts Dose-Dependent Nutritional, Metabolic, and Environmental Impacts in Broiler Chicks. J. Appl. Poult. Res. 2017, 26, 358–366. [CrossRef]

- Czech, A.; Smolczyk, A.; Ognik, K.; Wlazło, Ł.; Nowakowicz-Dębek, B.; Kiesz, M. Effect of Dietary Supplementation with Yarrowia Lipolytica or Saccharomyces Cerevisiae Yeast and Probiotic Additives on Haematological Parameters and the Gut Microbiota in Piglets. *Res. Vet. Sci.* 2018, 119, 221–227. [CrossRef]
- 171. Foltz, K.L.; Smith, D.L.; Moritz, J.S. Porcine Feed Intake of Corn–Soybean Based Diets Supplemented with Oil-Extracted Microalgae and Subsequent Performance. *Prof. Anim. Sci.* 2016, *32*, 849–853. [CrossRef]
- 172. Manor, M.L.; Kim, J.; Derksen, T.J.; Schwartz, R.L.; Roneker, C.A.; Bhatnagar, R.S.; Lei, X.G. Defatted Microalgae Serve as a Dual Dietary Source of Highly Bioavailable Iron and Protein in an Anemic Pig Model. *Algal Res.* **2017**, *26*, 409–414. [CrossRef]
- 173. Madeira, M.S.; Cardoso, C.; Lopes, P.A.; Coelho, D.; Afonso, C.; Bandarra, N.M.; Prates, J.A.M. Microalgae as Feed Ingredients for Livestock Production and Meat Quality: A Review. *Livest. Sci.* 2017, 205, 111–121. [CrossRef]
- 174. Pestana, J.M.; Puerta, B.; Santos, H.; Madeira, M.S.; Alfaia, C.M.; Lopes, P.A.; Pinto, R.M.A.; Lemos, J.P.C.; Fontes, C.M.G.A.; Lordelo, M.M.; et al. Impact of Dietary Incorporation of Spirulina (*Arthrospira platensis*) and Exogenous Enzymes on Broiler Performance, Carcass Traits, and Meat Quality. *Poult. Sci.* 2020, 99, 2519–2532. [CrossRef] [PubMed]
- 175. Toyomizu, M.; Sato, K.; Taroda, H.; Kato, T.; Akiba, Y. Effects of Dietary Spirulina on Meat Colour in Muscle of Broiler Chickens. *Br. Poult. Sci.* **2001**, *42*, 197–202. [CrossRef]
- Grinstead, G.S.; Tokach, M.D.; Dritz, S.S.; Goodband, R.D.; Nelssen, J.L. Effects of Spirulina Platensis on Growth Performance of Weanling Pigs. *Anim. Feed Sci. Technol.* 2000, 83, 237–247. [CrossRef]
- 177. El-Bahr, S.; Shousha, S.; Shehab, A.; Khattab, W.; Ahmed-Farid, O.; Sabike, I.; El-Garhy, O.; Albokhadaim, I.; Albosadah, K. Effect of Dietary Microalgae on Growth Performance, Profiles of Amino and Fatty Acids, Antioxidant Status, and Meat Quality of Broiler Chickens. *Animals* 2020, 10, 761. [CrossRef]
- 178. Gatrell, S.K.; Magnuson, A.D.; Barcus, M.; Lei, X.G. Graded Levels of a Defatted Green Microalgae Inclusion in Diets for Broiler Chicks Led to Moderate Up-Regulation of Protein Synthesis Pathway in the Muscle and Liver. *Algal Res.* 2018, 29, 290–296. [CrossRef]
- 179. Sun, T.; Wang, K.; Wyman, B.; Sudibyo, H.; Liu, G.; Beal, C.; Manning, S.; Johnson, Z.I.; Aydemir, T.B.; Tester, J.W.; et al. Supplemental Dietary Full-Fatted and Defatted *Desmodesmus* Sp. Exerted Similar Effects on Growth Performance, Gut Health, and Excreta Hydrothermal Liquefaction of Broiler Chicks. *Algal Res.* **2021**, *54*, 102205. [CrossRef]
- Martins, C.F.; Pestana Assunção, J.; Ribeiro Santos, D.M.; Madeira, M.S.M.D.S.; Alfaia, C.M.R.P.M.; Lopes, P.A.A.B.; Coelho, D.F.M.; Cardoso Lemos, J.P.; Almeida, A.M.; Mestre Prates, J.A.; et al. Effect of Dietary Inclusion of Spirulina on Production Performance, Nutrient Digestibility and Meat Quality Traits in Post-weaning Piglets. *J. Anim. Physiol. Anim. Nutr.* 2021, 105, 247–259. [CrossRef]
- Schøyen, H.F.; Frøyland, J.R.K.; Sahlström, S.; Knutsen, S.H.; Skrede, A. Effects of Autolysis and Hydrolysis of Bacterial Protein Meal Grown on Natural Gas on Chemical Characterization and Amino Acid Digestibility. *Aquaculture* 2005, 248, 27–33. [CrossRef]
- Øverland, M.; Skrede, A. Yeast Derived from Lignocellulosic Biomass as a Sustainable Feed Resource for Use in Aquaculture. J. Sci. Food Agric. 2017, 97, 733–742. [CrossRef]
- Øverland, M.; Schøyen, H.F.; Skrede, A. Growth Performance and Carcase Quality in Broiler Chickens Fed on Bacterial Protein Grown on Natural Gas. Br. Poult. Sci. 2010, 51, 686–695. [CrossRef]
- Øverland, M.; Mydland, L.T.; Skrede, A. Marine Macroalgae as Sources of Protein and Bioactive Compounds in Feed for Monogastric Animals. J. Sci. Food Agric. 2019, 99, 13–24. [CrossRef]
- Ritala, A.; Häkkinen, S.T.; Toivari, M.; Wiebe, M.G. Single Cell Protein—State-of-the-Art, Industrial Landscape and Patents 2001–2016. Front. Microbiol. 2017, 8, 2009. [CrossRef]
- Mielke, T. World Markets for Vegetable Oils and Animal Fats. In *Biokerosene*; Springer: Berlin/Heidelberg, Germany, 2018; pp. 147–188. ISBN 9783662530658.
- Cera, K.R.; Mahan, D.C.; Reinhart, G.A. Weekly Digestibilities of Diets Supplemented with Corn Oil, Lard or Tallow by Weanling Swine. J. Anim. Sci. 1988, 66, 1430. [CrossRef]
- Cera, K.R.; Mahan, D.C.; Reinhart, G.A. Apparent Fat Digestibilities and Performance Responses of Postweaning Swine Fed Diets Supplemented with Coconut Oil, Corn Oil or Tallow. J. Anim. Sci. 1989, 67, 2040. [CrossRef]
- 189. USDA. Oil Crops Yearbook; Economic Research Service, U.S. Department of Agriculture: Washington, DC, USA, 2020.
- European Parliament, C. of the E.U. Regulation 1069/2009. Regul. No 1069/2009 European Parliament and of the Council of 21 October 2009 Lay down Health rules as regards Anim. by-products Derived Products not Intended for Human Consumption repealing Regulation no. 1774/2002. Off. J. Eur. Union 2009, 52, 109.
- Park, S.W.; Seo, S.H.; Chang, M.B.; Shin, I.S.; Paik, I.K. Evaluation of Soybean Oil as a Lipid Source for Pig Diets. *Asian-Australasian J. Anim. Sci.* 2009, 22, 1311–1319. [CrossRef]
- 192. Shurson, G.C.; Kerr, B.J.; Hanson, A.R. Evaluating the Quality of Feed Fats and Oils and Their Effects on Pig Growth Performance. J. Anim. Sci. Biotechnol. 2015, 6, 10. [CrossRef]
- Darvishi, F.; Salmani, N.; Hosseini, B. Biovalorization of Vegetable Oil Refinery Wastewater into Value-added Compounds by Yarrowia Lipolytica. J. Chem. Technol. Biotechnol. 2019, 94, 2961–2968. [CrossRef]
- 194. Abril, R.; Garrett, J.; Zeller, S.G.; Sander, W.J.; Mast, R.W. Safety Assessment of DHA-Rich Microalgae from Schizochytrium Sp. Part V: Target Animal Safety/Toxicity Study in Growing Swine. *Regul. Toxicol. Pharmacol.* **2003**, *37*, 73–82. [CrossRef]
- Alvarez, H.M.; Kalscheuer, R.; Steinbüchel, A. Accumulation and Mobilization of Storage Lipids by Rhodococcus Opacus PD630 and Rhodococcus Ruber NCIMB 40126. *Appl. Microbiol. Biotechnol.* 2000, 54, 218–223. [CrossRef] [PubMed]

- Galán, B.; Santos-merino, M.; Nogales, J. Health Consequences of Microbial Interactions with Hydrocarbons, Oils, and Lipids. Goldfine, H., Ed.; Springer International Publishing: Cham, Switzerland, 2020; ISBN 978-3-030-15146-1.
- 197. Thies, F.; Peterson, L.D.; Powell, J.R.; Nebe-von-Caron, G.; Hurst, T.L.; Matthews, K.R.; Newsholme, E.A.; Calder, P.C. Manipulation of the Type of Fat Consumed by Growing Pigs Affects Plasma and Mononuclear Cell Fatty Acid Compositions and Lymphocyte and Phagocyte Functions. *J. Anim. Sci.* **1999**, 77, 137. [CrossRef] [PubMed]
- 198. Jin, C.F.; Kim, J.H.; Han, I.K.; Jung, H.J.; Kwon, C.H. Effects of Various Fat Sources and Lecithin on the Growth Performances and Nutrient Utilization in Pigs Weaned at 21 Days of Age. *Asian-Australas. J. Anim. Sci.* **1998**, *11*, 176–184. [CrossRef]
- Lindblom, S.C.; Dozier, W.A.; Shurson, G.C.; Kerr, B.J. Digestibility of Energy and Lipids and Oxidative Stress in Nursery Pigs Fed Commercially Available Lipids. J. Anim. Sci. 2017, 95, 239. [CrossRef]
- Lauridsen, C.; Bruun Christensen, T.; Halekoh, U.; Krogh Jensen, S. Alternative Fat Sources to Animal Fat for Pigs. *Lipid Technol.* 2007, 19, 156–159. [CrossRef]
- van Heugten, E.; Coffey, M.T.; Spears, J.W. Effects of Immune Challenge, Dietary Energy Density, and Source of Energy on Performance and Immunity in Weanling Pigs. J. Anim. Sci. 1996, 74, 2431. [CrossRef]
- Gatlin, L.A.; See, M.T.; Larick, D.K.; Lin, X.; Odle, J. Conjugated Linoleic Acid in Combination with Supplemental Dietary Fat Alters Pork Fat Quality. J. Nutr. 2002, 132, 3105–3112. [CrossRef]
- Sousa, R.V.; Fialho, E.T.; Lima, J.A.F.; Alvarez-Leite, J.I.; Cortez, W.C.; Ferreira, M.S.S. Effect of Different Oils in Diets for Finishing Pigs: Performance, Carcass Traits and Fatty Acid Profile of the Meat. *Anim. Prod. Sci.* 2010, 50, 863. [CrossRef]
- 204. Pickova, J.; Mørkøre, T. Alternate Oils in Fish Feeds. Eur. J. Lipid Sci. Technol. 2007, 109, 256–263. [CrossRef]
- 205. Blomqvist, J.; Pickova, J.; Tilami, S.K.; Sampels, S.; Mikkelsen, N.; Brandenburg, J.; Sandgren, M.; Passoth, V. Oleaginous Yeast as a Component in Fish Feed. *Sci. Rep.* **2018**, *8*, 15945. [CrossRef]
- Hatlen, B.; Berge, G.M.; Odom, J.M.; Mundheim, H.; Ruyter, B. Growth Performance, Feed Utilisation and Fatty Acid Deposition in Atlantic Salmon, Salmo Salar L., Fed Graded Levels of High-Lipid/High-EPA Yarrowia Lipolytica Biomass. *Aquaculture* 2012, 364–365, 39–47. [CrossRef]
- 207. Eryalçin, K.M.; Ganuza, E.; Atalah, E.; Cruz, M.C.H. Nannochloropsis Gaditana and Crypthecodinium Cohnii, Two Microalgae as Alternative Sources of Essential Fatty Acids in Early Weaning for Gilthead Seabream. Hidrobiologica 2015, 25, 193–202.
- 208. Santigosa, E.; Constant, D.; Prudence, D.; Wahli, T.; Verlhac-Trichet, V. A Novel Marine Algal Oil Containing Both EPA and DHA Is an Effective Source of Omega-3 Fatty Acids for Rainbow Trout Oncorhynchus Mykiss. J. World Aquac. Soc. 2020, 51, 649–665. [CrossRef]
- 209. Barclay, W.; Zeller, S. Nutritional Enhancement of N-3 and n-6 Fatty Acids in Rotifers and Artemia Nauplii by Feeding Spray-Dried *Schizochytrium* sp. *J. World Aquac. Soc.* **1996**, *27*, 314–322. [CrossRef]
- Groenewald, M.; Boekhout, T.; Neuvéglise, C.; Gaillardin, C.; van Dijck, P.W.M.; Wyss, M. Yarrowia Lipolytica: Safety Assessment of an Oleaginous Yeast with a Great Industrial Potential. *Crit. Rev. Microbiol.* 2014, 40, 187–206. [CrossRef]
- 211. Khot, M.; Kamat, S.; Zinjarde, S.; Pant, A.; Chopade, B.; RaviKumar, A. Single Cell Oil of Oleaginous Fungi from the Tropical Mangrove Wetlands as a Potential Feedstock for Biodiesel. *Microb. Cell Fact.* 2012, 11, 71. [CrossRef]
- den Haan, R.; van Rensburg, E.; Rose, S.H.; Görgens, J.F.; van Zyl, W.H. Progress and Challenges in the Engineering of Non-Cellulolytic Microorganisms for Consolidated Bioprocessing. *Curr. Opin. Biotechnol.* 2015, 33, 32–38. [CrossRef]
- Cheng, Y.-C.; Duarte, M.E.; Kim, S.W. Effects of Yarrowia Lipolytica Supplementation on Growth Performance, Intestinal Health and Apparent Ileal Digestibility of Diets Fed to Nursery Pigs. *Anim. Biosci.* 2022, 35, 605–613. [CrossRef]
- 214. Berge, G.M.; Hatlen, B.; Odom, J.M.; Ruyter, B. Physical Treatment of High EPA Yarrowia Lipolytica Biomass Increases the Availability of N-3 Highly Unsaturated Fatty Acids When Fed to Atlantic Salmon. *Aquac. Nutr.* **2013**, *19*, 110–121. [CrossRef]
- 215. Sun, T.; Tolba, S.A.; Magnuson, A.D.; Lei, X.G. Excessive Aurantiochytrium Acetophilum Docosahexaenoic Acid Supplementation Decreases Growth Performance and Breast Muscle Mass of Broiler Chickens. *Algal Res.* **2022**, *63*, 102648. [CrossRef]
- 216. Lee, A.; You, L.; Oh, S.-Y.; Li, Z.; Code, A.; Zhu, C.; Fisher-Heffernan, R.; Regnault, T.; De Lange, C.; Huber, L.-A.; et al. Health Benefits of Supplementing Nursery Pig Diets with Microalgae or Fish Oil. *Animals* 2019, 9, 80. [CrossRef] [PubMed]
- 217. De Tonnac, A.; Labussière, E.; Vincent, A.; Mourot, J. Effect of α -Linolenic Acid and DHA Intake on Lipogenesis and Gene Expression Involved in Fatty Acid Metabolism in Growing-Finishing Pigs. *Br. J. Nutr.* 2016, *116*, 7–18. [CrossRef]
- De Tonnac, A.; Guillevic, M.; Mourot, J. Fatty Acid Composition of Several Muscles and Adipose Tissues of Pigs Fed N-3 PUFA Rich Diets. *Meat Sci.* 2018, 140, 1–8. [CrossRef]
- Patil, V.; Reitan, K.; Mortensen, L.; Källqvist, T.; Olsen, Y.; Vogt, G.; Gislerød, H. Microalgae as a Source of Polyunsaturated Fatty Acids for Aquaculture. *Curr. Top. Plant Biol.* 2005, 6, 57–65.
- Harel, M.; Koven, W.; Lein, I.; Bar, Y.; Behrens, P.; Stubblefield, J.; Zohar, Y.; Place, A.R. Advanced DHA, EPA and ArA Enrichment Materials for Marine Aquaculture Using Single Cell Heterotrophs. *Aquaculture* 2002, 213, 347–362. [CrossRef]
- 221. Atalah, E.; Cruz, C.M.H.; Izquierdo, M.S.; Rosenlund, G.; Caballero, M.J.; Valencia, A.; Robaina, L. Two Microalgae Crypthecodinium Cohnii and Phaeodactylum tricornutum as Alternative Source of Essential Fatty Acids in Starter Feeds for Seabream (Sparus aurata). Aquaculture 2007, 270, 178–185. [CrossRef]