

**Effects of dietary organic acids and nature identical compounds on growth, immune
parameters and gut microbiota of European sea bass**

Serena Busti¹, Barbara Rossi², Enrico Volpe¹, Sara Ciulli¹, Andrea Piva^{1,2}, Federica D'Amico³,
Matteo Soverini³, Marco Candela³, Pier Paolo Gatta¹, Alessio Bonaldo^{1*}, Ester Grilli¹, Luca Parma¹

¹Department of Veterinary Medical Sciences, University of Bologna, Via Tolara di Sopra 50, 40064
Ozzano Emilia, Bologna, Italy;

²Vetagro S.p.A., Via Porro 2, 42124, Reggio Emilia, Italy;

³Unit of Microbial Ecology of Health, Department of Pharmacy and Biotechnology, University of
Bologna, Via Belmeloro 6, 40126 Bologna, Italy

*Corresponding author: alessio.bonaldo@unibo.it

SUPPLEMENTARY INFORMATION

Supplementary Table S1. Mean relative abundance (%) \pm SD (n=3) of bacterial phyla and genera detected in the hindgut content of European sea bass fed different diets at 71 days (pre-suboptimal rearing condition), and at 81 days (post-suboptimal rearing condition). D0 = 0 mg Kg⁻¹ organic acids and nature-identical compounds (OA, NIC); D250 = 250 mg Kg⁻¹ OA, NIC; D500 = 500 mg Kg⁻¹ OA, NIC; D1000 = 1000 mg Kg⁻¹ OA, NIC. Only taxa with relative abundance >0.05% in at least 1 samples were included. Taxa which represent <0.05% abundance were grouped “Other”.

Pre- suboptimal rearing condition									Post-suboptimal rearing condition					
Diet	D0		D250		D500		D1000		D0		D1000			
Phylum	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD
<i>Firmicutes</i>	69.4	3.6	78	10.8	78.4	3.9	77.7	4.5	<i>Firmicutes</i>	54.6	3.2	54.2	1.4	
<i>Actinobacteria</i>	12.3	1.9	9.1	1.6	10.8	0.5	10.5	1.3	<i>Proteobacteria</i>	13.5	3.8	18.4	0.8	
<i>Cyanobacteria</i>	12.2	2.8	7.3	8.2	6.3	4.6	3.4	1.2	<i>Fusobacteria</i>	7.7	6.0	9.9	2.2	
<i>Proteobacteria</i>	5	0.8	4.7	1.7	3.5	1.4	6	4.0	<i>Bacteroidetes</i>	13	3.7	8.3	1.3	
<i>Acidobacteria</i>	0.4	0.2	0.2	0.2	0.3	0.3	0.8	1.1	<i>Actinobacteria</i>	6.5	1.5	5.6	0.3	
<i>Bacteroidetes</i>	0.4	0.2	0.4	0.3	0.3	0.1	0.4	0.3	<i>Cyanobacteria</i>	0.3	0.2	0.9	0.8	
<i>Chloroflexi</i>	0	0.0	0	0.0	0	0.0	0.4	0.6	<i>Verrucomicrobia</i>	2.4	1.2	0.6	0.5	
<i>TM7</i>	0.1	0.1	0.1	0.1	0.2	0.1	0.3	0.3	<i>Chlamydiae</i>	0.1	0.1	0.3	0.3	
<i>Planctomycetes</i>	0	0.0	0	0.0	0	0.0	0.1	0.2	<i>Acidobacteria</i>	0.1	0.1	0.2	0.1	
<i>WPS-2</i>	0	0.0	0	0.0	0	0.0	0.1	0.2	<i>Chloroflexi</i>	0.1	0.2	0.1	0.2	
<i>Verrucomicrobia</i>	0.1	0.0	0	0.0	0	0.0	0.1	0.1	<i>Planctomycetes</i>	0.1	0.1	0.1	0.1	
<i>Other</i>	0.1	0.1	0.1	0.0	0.1	0.0	0.2	0.2	<i>Other</i>	1.3	1.2	1.2	1.3	
Diet	D0		D250		D500		D1000		D0		D1000			
Genus	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD
<i>Lactobacillus</i>	25.2	0.8	23.1	3.4	27.6	1.5	23.2	4.6	<i>Enterobacteriaceae</i>	8.3	5.4	11.5	2.0	
<i>Leuconostocaceae</i>	12.4	3.2	21.3	4.2	17.7	0.9	14.5	7.0	<i>Eubacterium</i>	7.9	5.9	10.5	3.3	
<i>Streptococcus</i>	16.3	3.9	16.1	3.1	14.7	4.0	13.2	7.0	<i>Cetobacterium</i>	7.7	5.9	9.8	2.2	
<i>Streptophyta</i>	12.2	2.8	7.3	8.2	6.3	4.6	3.2	1.6	<i>Bacillus</i>	3.7	2.5	5.5	1.7	
<i>Bacillus</i>	3.0	2.4	3.4	1.9	4.4	3.2	7.4	6.8	<i>Clostridium</i>	3.4	1.2	4.4	0.2	
<i>Corynebacterium</i>	8.8	1.2	5.6	1.2	7.2	0.5	6.6	0.8	<i>Erysipelotrichaceae</i>	3.0	1.7	4.2	0.9	

<i>Leuconostoc</i>	4.5	0.3	5.3	0.6	4.9	1.2	4.6	2.6	<i>Bacteroidaceae</i>	3.1	2.6	3.8	1.3
<i>Bacillaceae</i>	1.3	0.6	1.8	1.0	1.9	0.8	3.7	3.5	<i>Coriobacteriaceae</i>	3.0	1.7	3.8	0.6
<i>Geobacillus</i>	0.6	0.5	0.9	0.4	1.2	0.6	2.1	2.4	<i>Lachnospiraceae</i>	3.8	0.1	3.1	0.0
<i>mitochondria</i>	2.9	1.2	2.6	2.6	1.9	0.9	1.8	0.7	<i>Mogibacteriaceae</i>	2.0	1.6	3.0	0.8
<i>Lactococcus</i>	1.3	0.2	1.7	0.4	1.6	0.3	1.5	0.6	<i>Bacillaceae</i>	2.2	1.6	2.7	1.0
<i>Bifidobacteriaceae</i>	1.4	0.9	1.5	0.3	1.5	0.1	1.5	1.0	<i>Peptostreptococcaceae</i>	1.7	1.1	2.4	0.5
<i>Clostridium</i>	0.4	0.2	0.5	0.5	0.8	0.6	0.9	0.7	<i>Ruminococcaceae</i>	6.2	3.7	1.8	1.1
<i>Bifidobacterium</i>	0.6	0.3	0.6	0.0	0.6	0.2	0.7	0.2	<i>Geobacillus</i>	1.4	1.0	1.8	0.1
<i>Bradyrhizobium</i>	0.1	0.1	0.2	0.1	0.2	0.2	0.7	0.9	<i>Dorea</i>	1.4	0.4	1.7	0.6
<i>Delftia</i>	0.2	0.2	0.6	0.4	0.2	0.2	0.6	0.6	<i>Sediminibacterium</i>	1.1	0.9	1.5	0.7
<i>Peptostreptococcaceae</i>	0.2	0.1	0.3	0.4	0.1	0.1	0.5	0.6	<i>Caloramator</i>	1.1	0.8	1.4	0.3
<i>Caulobacteraceae</i>	0.1	0.0	0.1	0.1	0.1	0.1	0.5	0.7	<i>Carnobacterium</i>	0.9	0.8	1.3	0.8
<i>Faecalibacterium</i>	0.1	0.1	0.0	0.1	0.1	0.1	0.4	0.5	<i>Clostridiales</i>	1.5	1.0	1.2	0.9
<i>Sphingobium</i>	0.0	0.0	0.2	0.2	0.2	0.1	0.4	0.6	<i>Bacteroides</i>	5.4	3.8	1.2	0.9
<i>Rhodococcus</i>	0.2	0.1	0.2	0.1	0.3	0.1	0.4	0.2	<i>Delftia</i>	0.6	0.5	1.0	0.6
<i>Lactobacillales</i>	0.6	0.1	0.4	0.0	0.3	0.1	0.4	0.3	<i>Sphingobium</i>	0.7	0.6	1.0	0.5
<i>Sporanaerobacter</i>	0.2	0.1	0.2	0.1	0.2	0.2	0.4	0.4	<i>Anaerobacillus</i>	0.6	0.6	0.8	0.3
<i>Blautia</i>	0.2	0.2	0.1	0.1	0.1	0.1	0.4	0.4	<i>Lachnospira</i>	1.0	0.8	0.8	1.0
<i>Solibacterales</i>	0.2	0.1	0.1	0.1	0.2	0.2	0.4	0.5	<i>Streptophyta</i>	0.2	0.1	0.7	0.8
<i>Sphingomonas</i>	0.1	0.1	0.2	0.2	0.1	0.1	0.4	0.5	<i>Faecalibacterium</i>	2.1	2.6	0.7	0.5
<i>Ellin6513</i>	0.1	0.1	0.1	0.1	0.1	0.0	0.4	0.6	<i>Caulobacteraceae</i>	0.5	0.4	0.6	0.2
<i>Ruminococcus</i>	0.1	0.0	0.2	0.1	0.1	0.0	0.4	0.0	<i>Akkermansia</i>	2.3	1.2	0.6	0.5
<i>Lachnospiraceae</i>	0.2	0.1	0.2	0.2	0.2	0.1	0.3	0.3	<i>Methylobacterium</i>	0.4	0.3	0.5	0.3
<i>Lactobacillaceae</i>	0.3	0.1	0.2	0.1	0.3	0.2	0.3	0.3	<i>Eubacteriaceae;Other</i>	0.4	0.2	0.5	0.3
<i>Clostridiaceae</i>	0.0	0.1	0.0	0.0	0.0	0.0	0.3	0.3	<i>Vagococcus</i>	0.4	0.3	0.5	0.4
<i>Conexibacteraceae</i>	0.0	0.1	0.1	0.1	0.0	0.0	0.3	0.2	<i>Epulopiscium</i>	0.3	0.2	0.5	0.2
<i>MLE1-12</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.2	0.4	<i>Lactobacillus</i>	0.8	0.6	0.4	0.5
<i>Bacillales</i>	0.1	0.0	0.1	0.2	0.2	0.1	0.2	0.3	<i>Ralstonia</i>	0.2	0.2	0.4	0.5
<i>Sporosarcina</i>	0.1	0.1	0.1	0.0	0.2	0.1	0.2	0.2	<i>Rikenellaceae</i>	1.7	0.5	0.4	0.6
<i>Thermogemmatisporaceae</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.2	0.3	<i>Myroides</i>	0.2	0.2	0.4	0.1
<i>Solibacillus</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.2	0.4	<i>Streptococcus</i>	0.7	0.6	0.4	0.4

<i>Ruminococcaceae</i>	0.1	0.1	0.1	0.1	0.3	0.0	0.2	0.2	<i>Blautia</i>	1.3	1.7	0.4	0.2
<i>Curtobacterium</i>	0.3	0.2	0.3	0.1	0.2	0.1	0.2	0.1	<i>Ruminococcus</i>	1.1	0.5	0.4	0.0
<i>Virgibacillus</i>	0.1	0.1	0.1	0.0	0.0	0.0	0.2	0.2	<i>Roseburia</i>	0.5	0.4	0.4	0.1
<i>TM7-1</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.2	0.3	<i>Bifidobacterium</i>	1.8	2.8	0.3	0.1
<i>Enterococcus</i>	0.1	0.1	0.1	0.1	0.2	0.1	0.2	0.2	<i>Clostridiaceae</i>	0.3	0.1	0.3	0.1
<i>Anaerobacillus</i>	0.0	0.0	0.1	0.1	0.1	0.1	0.2	0.3	<i>Eggerthella</i>	0.3	0.2	0.3	0.0
<i>Ellin329</i>	0.0	0.1	0.0	0.0	0.0	0.0	0.2	0.3	<i>Collinsella</i>	0.5	0.5	0.3	0.2
<i>Sediminibacterium</i>	0.2	0.1	0.1	0.2	0.1	0.1	0.1	0.2	<i>Enhydrobacter</i>	0.1	0.1	0.3	0.4
<i>Carnobacterium</i>	0.1	0.1	0.1	0.1	0.1	0.0	0.1	0.0	<i>Prevotella</i>	0.2	0.2	0.3	0.4
<i>Methylobacterium</i>	0.0	0.0	0.1	0.1	0.0	0.0	0.1	0.2	<i>Parabacteroides</i>	0.7	0.5	0.3	0.5
<i>Ralstonia</i>	0.1	0.0	0.1	0.1	0.0	0.0	0.1	0.2	<i>Bradyrhizobium</i>	0.2	0.2	0.3	0.0
<i>WPS-2</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.2	<i>Corynebacterium</i>	0.3	0.4	0.3	0.4
<i>Tissierella_Soehngenia</i>	0.1	0.1	0.0	0.0	0.0	0.0	0.1	0.2	<i>Bacillales</i>	0.2	0.2	0.3	0.1
<i>Collinsella</i>	0.1	0.1	0.0	0.0	0.2	0.3	0.1	0.1	<i>Lysinibacillus</i>	0.2	0.2	0.2	0.1
<i>Bacteroides</i>	0.1	0.1	0.1	0.0	0.1	0.1	0.1	0.1	<i>Phascolarctobacterium</i>	0.4	0.4	0.2	0.2
<i>Planctomyces</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.2	<i>Paenibacillus</i>	0.1	0.2	0.2	0.1
<i>Ochrobactrum</i>	0.0	0.0	0.1	0.1	0.1	0.0	0.1	0.1	<i>Coprococcus</i>	0.5	0.4	0.2	0.2
<i>Staphylococcus</i>	0.2	0.1	0.3	0.2	0.1	0.0	0.1	0.1	<i>MLE1-12</i>	0.1	0.1	0.2	0.1
<i>Pediococcus</i>	0.3	0.1	0.2	0.0	0.0	0.0	0.1	0.1	<i>Oscillospira</i>	0.3	0.3	0.2	0.2
<i>Peptostreptococcus</i>	0.1	0.2	0.0	0.0	0.0	0.0	0.1	0.1	<i>Anaerostipes</i>	0.1	0.2	0.2	0.2
<i>Ureibacillus</i>	0.0	0.0	0.0	0.1	0.1	0.1	0.1	0.0	<i>Sphingomonas</i>	0.1	0.1	0.2	0.2
<i>Enterobacteriaceae</i>	0.1	0.2	0.1	0.1	0.0	0.0	0.1	0.2	<i>Acinetobacter</i>	0.1	0.1	0.2	0.0
<i>JG30-KF-CM45</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.2	<i>Serratia</i>	0.1	0.1	0.2	0.2
<i>Thermicanaceae</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.2	<i>mitochondria</i>	0.1	0.0	0.2	0.1
<i>Rs-045</i>	0.1	0.1	0.1	0.1	0.2	0.1	0.1	0.1	<i>Anaerovorax</i>	0.1	0.1	0.2	0.1
<i>Sinobacteraceae</i>	0.0	0.0	0.0	0.0	0.1	0.1	0.1	0.1	<i>Staphylococcus</i>	0.1	0.2	0.2	0.2
<i>Brevibacillus</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Ochrobactrum</i>	0.0	0.0	0.2	0.1
<i>Erysipelotrichaceae</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Parachlamydia</i>	0.1	0.0	0.1	0.1
<i>Pseudomonas</i>	0.0	0.0	0.0	0.1	0.0	0.0	0.1	0.1	<i>Thermogemmatisporaceae</i>	0.1	0.2	0.1	0.2
<i>Acinetobacter</i>	0.0	0.0	0.1	0.1	0.0	0.0	0.1	0.1	<i>Morganella</i>	0.0	0.0	0.1	0.1
<i>Clostridiales</i>	0.1	0.0	0.0	0.0	0.2	0.1	0.1	0.1	<i>Halomonas</i>	0.0	0.1	0.1	0.1
<i>Bradyrhizobiaceae</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Ellin329</i>	0.1	0.1	0.1	0.1

<i>Lysinibacillus</i>	0.0	0.1	0.1	0.1	0.1	0.1	0.1	0.0	<i>Stenotrophomonas</i>	0.1	0.1	0.1	0.1
<i>Saccharopolyspora</i>	0.0	0.0	0.1	0.1	0.1	0.0	0.1	0.1	<i>Jeotgalicoccus</i>	0.1	0.1	0.1	0.2
<i>Megamonas</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Allobaculum</i>	0.1	0.1	0.1	0.1
<i>Aneurinibacillus</i>	0.1	0.1	0.0	0.0	0.0	0.0	0.1	0.0	<i>Ureibacillus</i>	0.1	0.1	0.1	0.0
<i>Oxalobacteraceae</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Pseudomonas</i>	0.1	0.2	0.1	0.1
<i>Akkermansia</i>	0.0	0.1	0.0	0.0	0.0	0.0	0.1	0.1	<i>Leuconostocaceae</i>	0.1	0.0	0.1	0.1
<i>Agrobacterium</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Fusobacterium</i>	0.1	0.1	0.1	0.0
<i>Kurthia</i>	0.3	0.1	0.1	0.1	0.2	0.1	0.1	0.1	<i>Parachlamydiaceae</i>	0.0	0.0	0.1	0.1
<i>Myroides</i>	0.0	0.0	0.1	0.0	0.1	0.1	0.1	0.0	<i>Christensenellaceae</i>	0.3	0.0	0.1	0.1
<i>Microbacterium</i>	0.1	0.0	0.0	0.0	0.1	0.0	0.1	0.1	<i>Solibacterales</i>	0.1	0.1	0.1	0.1
<i>Micrococcaceae</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Sutterella</i>	0.4	0.4	0.1	0.1
<i>Micrococcus</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Natronobacillus</i>	0.0	0.0	0.1	0.1
<i>Mycetocola</i>	0.1	0.1	0.1	0.1	0.1	0.0	0.1	0.0	<i>Sinobacteraceae</i>	0.0	0.0	0.1	0.1
<i>Halomonas</i>	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1	<i>Dietzia</i>	0.1	0.1	0.1	0.1
<i>Coriobacteriaceae</i>	0.1	0.1	0.1	0.2	0.1	0.0	0.1	0.1	<i>Paracoccus</i>	0.0	0.1	0.1	0.1
<i>Ruminococcus</i>	0.0	0.0	0.1	0.1	0.0	0.0	0.1	0.0	<i>Rhodococcus</i>	0.1	0.1	0.1	0.0
<i>Frigoribacterium</i>	0.1	0.0	0.0	0.0	0.0	0.0	0.0	0.1	<i>Lachnospiraceae;Other</i>	0.0	0.1	0.1	0.0
<i>Streptomyces</i>	0.1	0.1	0.0	0.1	0.0	0.0	0.0	0.1	<i>Dechloromonas</i>	0.0	0.0	0.1	0.1
<i>Tepidimicrobium</i>	0.0	0.0	0.1	0.0	0.0	0.0	0.0	0.1	<i>Aneurinibacillus</i>	0.1	0.1	0.1	0.1
<i>Peptoniphilus</i>	0.1	0.2	0.0	0.1	0.0	0.0	0.0	0.0	<i>RF32</i>	0.1	0.1	0.1	0.1
<i>Vagococcus</i>	0.1	0.0	0.1	0.1	0.0	0.0	0.0	0.0	<i>Enterococcus</i>	0.0	0.0	0.1	0.0
<i>Brevibacterium</i>	0.0	0.0	0.0	0.1	0.1	0.1	0.0	0.0	<i>Prevotella</i>	0.0	0.2	0.1	0.4
<i>Tissierellaceae</i>	0.1	0.1	0.0	0.0	0.0	0.0	0.0	0.0	<i>MND1</i>	0.0	0.0	0.1	0.1
<i>Paracoccus</i>	0.3	0.4	0.1	0.1	0.0	0.1	0.0	0.0	<i>Barnesiellaceae</i>	0.2	0.2	0.1	0.1
<i>Enhydrobacter</i>	0.3	0.5	0.0	0.0	0.0	0.0	0.0	0.0	<i>Sporanaerobacter</i>	0.1	0.2	0.1	0.1
<i>Planococcaceae</i>	0.0	0.0	0.1	0.0	0.0	0.0	0.0	0.0	<i>Actinomyces</i>	0.1	0.1	0.1	0.1
<i>Paraprevotella</i>	0.0	0.0	0.1	0.2	0.0	0.0	0.0	0.0	<i>Ellin6513</i>	0.0	0.0	0.1	0.0
<i>Roseomonas</i>	0.2	0.3	0.0	0.0	0.0	0.0	0.0	0.0	<i>AF12</i>	0.0	0.0	0.1	0.1
<i>Unassigned</i>	0.1	0.1	0.1	0.0	0.1	0.0	0.2	0.2	<i>Rubellimicrobium</i>	0.0	0.0	0.1	0.1
									<i>Chryseobacterium</i>	0.0	0.1	0.1	0.1
									<i>Rhodocyclaceae</i>	0.0	0.0	0.1	0.1
									<i>Tepidimicrobium</i>	0.1	0.1	0.1	0.1

<i>Pseudoramibacter_Eubacterium</i>	0.1	0.0	0.1	0.0
<i>Pseudomonadaceae</i>	0.0	0.0	0.1	0.0
<i>Conexibacteraceae</i>	0.0	0.0	0.1	0.1
<i>Pirellulaceae</i>	0.0	0.0	0.1	0.1
<i>Butyricimonas</i>	0.0	0.0	0.1	0.0
<i>Mycobacterium</i>	0.0	0.0	0.1	0.1
<i>Paludibacter</i>	0.1	0.0	0.0	0.0
<i>Thermoactinomyces</i>	0.1	0.1	0.0	0.0
<i>Ruminococcus</i>	0.3	0.5	0.0	0.0
<i>Anaerotruncus</i>	0.1	0.1	0.0	0.1
<i>Luteimonas</i>	0.1	0.1	0.0	0.1
<i>Dialister</i>	0.5	0.8	0.0	0.0
<i>Acidaminococcus</i>	0.1	0.1	0.0	0.0
<i>Tissierellaceae</i>	0.1	0.0	0.0	0.0
<i>Turicibacter</i>	0.1	0.1	0.0	0.0
<i>Odoribacter</i>	0.1	0.1	0.0	0.0
<i>Bilophila</i>	0.3	0.5	0.0	0.0
<i>Catenibacterium</i>	0.1	0.2	0.0	0.0
<i>Anaerobiospirillum</i>	0.2	0.3	0.0	0.0
<i>Slackia</i>	0.1	0.2	0.0	0.0
<i>Megamonas</i>	0.1	0.1	0.0	0.0
<i>Other</i>	1.3	1.2	1.2	1.3

Supplementary Table S2. Statistical analysis on gut microbiota detected in the hindgut content of European sea bass fed different diets at 71 days (pre-suboptimal rearing condition), and at 81 days (post-suboptimal rearing condition). All the *P* values were appropriately corrected for multiple comparisons using the Benjamini-Hochberg method. False discovery rate (FDR) ≤ 0.05 was considered as statistically significant. D0= 0 mg Kg⁻¹ organic acids and nature-identical compounds (OA, NIC); D250 = 250 mg Kg⁻¹ OA, NIC; D500 = 500 mg Kg⁻¹ OA, NIC; D1000 = 1000mg Kg⁻¹ OA, NIC. D0vsD250=comparisons between D0 and D250 genera under pre-suboptimal rearing condition; D0vsD500=comparisons between D0 and D500 genera under pre-suboptimal rearing condition; D0vsD1000=comparisons between D0 and D1000 genera under pre-suboptimal rearing condition and post-suboptimal rearing condition; D0vsD0=comparisons between pre-suboptimal rearing condition and post- suboptimal rearing condition D0 genera; D1000vsD1000=comparisons between pre- suboptimal rearing condition and post- suboptimal rearing condition D1000 genera.

Diet	Pre-suboptimal rearing condition			Post-suboptimal rearing condition		Pre-Post suboptimal rearing condition		
				D0 vs D1000	P-value	D0	D1000	
	D0 vs D250	D0 vs D500	D0 vs D1000			vs D0	vs D1000	
Genus	P- value				P-value			P-value
<i>Lactobacillus</i>	0.66	0.08	0.66	<i>Enterobacteriaceae</i>	1.0	<i>Rikenellaceae</i>	0.06	0.08
<i>Leuconostocaceae</i>	0.08	0.08	0.66	<i>Eubacterium</i>	0.7	<i>Myroides</i>	0.18	0.1
<i>Streptococcus</i>	1.0	0.66	1.0	<i>Bacillus</i>	0.7	<i>Acinetobacter</i>	0.06	0.7
<i>Bacillus</i>	1.0	1.0	0.38	<i>Clostridium</i>	0.7	<i>Enterococcus</i>	0.08	0.7
<i>Corynebacterium</i>	0.08	0.19	0.08	<i>Erysipelotrichaceae</i>	0.4	<i>Planctomyces</i>	0.2	0.5
<i>Leuconostoc</i>	0.12	0.66	1.0	<i>Bacteroides</i>	0.2	<i>Paraprevotella</i>	0.64	NA
<i>Bacillaceae</i>	0.66	0.38	0.38	<i>Coriobacteriaceae</i>	1.0	<i>Christensenellaceae</i>	0.08	0.06
<i>Streptophyta</i>	0.66	0.19	0.08	<i>Lachnospiraceae</i>	1.0	<i>Morganella</i>	0.18	0.06
<i>Geobacillus</i>	0.38	0.38	0.38	<i>Mogibacteriaceae</i>	0.4	<i>Akkermansia</i>	0.08	0.08
<i>mitochondria</i>	0.66	0.19	0.38	<i>Bacillaceae</i>	0.4	<i>Ruminococcaceae</i>	0.08	0.1
<i>Lactococcus</i>	0.19	0.38	0.66	<i>Peptostreptococcaceae</i>	0.4	<i>Bacteroides</i>	0.1	0.1
<i>Bifidobacteriaceae</i>	0.82	0.66	1.0	<i>Ruminococcaceae</i>	0.2	<i>Streptophyta</i>	0.1	0.2
<i>Clostridium</i>	1.0	0.38	0.27	<i>Geobacillus</i>	1.0	<i>Sinobacteraceae</i>	0.38	1.0
<i>Bifidobacterium</i>	1.0	1.0	0.66	<i>Dorea</i>	1.0	<i>Propionibacteriaceae</i>	0.08	0.38
<i>Bradyrhizobium</i>	1.0	0.82	0.38	<i>Sediminibacterium</i>	1.0	<i>Lactobacillales</i>	0.08	0.08
<i>Delftia</i>	0.66	1.0	0.38	<i>Caloramator</i>	1.0	<i>Leuconostoc</i>	0.08	0.08
<i>Peptostreptococcaceae</i>	1.0	0.26	0.66	<i>Carnobacterium</i>	0.7	<i>Legionellaceae</i>	1.0	0.5
<i>Caulobacteraceae</i>	0.66	0.82	0.66	<i>Clostridiales</i>	0.7	<i>Natronobacillus</i>	1.0	0.51

<i>Faecalibacterium</i>	0.37	0.64	0.66	<i>Bacteroides</i>	0.2	<i>mitochondria</i>	0.08	0.1
<i>Sphingobium</i>	0.64	0.2	0.06	<i>Delftia</i>	1.0	<i>Ellin6513</i>	0.07	1.0
<i>Rhodococcus</i>	1.0	0.66	0.19	<i>Sphingobium</i>	0.7	<i>Coprococcus</i>	0.1	0.06
<i>Lactobacillales</i>	0.08	0.08	0.82	<i>Anaerobacillus</i>	0.4	<i>Mogibacteriaceae</i>	0.08	0.06
<i>Sporanaerobacter</i>	0.66	1.0	0.38	<i>Lachnospira</i>	0.4	<i>Parachlamydia</i>	0.06	0.06
<i>Blautia</i>	1.0	0.38	0.66	<i>Streptophyta</i>	0.2	<i>Lachnospira</i>	0.06	0.06
<i>Solibacterales</i>	0.38	1.0	1.0	<i>Faecalibacterium</i>	1.0	<i>Anaerobacillus</i>	0.08	0.08
<i>Sphingomonas</i>	1.0	0.82	1.0	<i>Caulobacteraceae</i>	0.5	<i>Erysipelotrichaceae</i>	0.1	0.1
<i>Ellin6513</i>	1.0	0.38	0.66	<i>Akkermansia</i>	0.2	<i>Lactobacillus</i>	0.1	0.1
<i>Ruminococcus</i>	0.38	0.66	0.08	<i>Methylobacterium</i>	0.7	<i>Peptostreptococcaceae</i>	0.1	0.1
<i>Lachnospiraceae</i>	1.0	1.0	0.66	<i>Vagococcus</i>	1.0	<i>Parabacteroides</i>	0.08	0.2
<i>Lactobacillaceae</i>	0.38	1.0	1.0	<i>Lactobacillus</i>	0.4	<i>Ochrobactrum</i>	1.0	0.7
<i>Clostridiaceae</i>	1.0	0.66	0.18	<i>Ralstonia</i>	1.0	<i>Ruminococcus</i>	0.1	1.0
<i>Conexibacteraceae</i>	0.64	0.64	0.08	<i>Rikenellaceae</i>	0.1	<i>Ruminococcus</i>	0.08	1.0
<i>Bacillales</i>	1.0	0.27	0.38	<i>Myroides</i>	0.1	<i>Virgibacillus</i>	0.07	0.08
<i>Sporosarcina</i>	0.66	0.65	0.38	<i>Streptococcus</i>	0.5	<i>Curtobacterium</i>	0.1	0.08
<i>Ruminococcaceae</i>	1.0	0.08	1.0	<i>Blautia</i>	0.7	<i>Comamonadaceae</i>	1.0	0.82
<i>Curtobacterium</i>	1.0	0.38	0.66	<i>Ruminococcus</i>	0.4	<i>Adlercreutzia</i>	0.2	1.0
<i>Enterococcus</i>	1.0	0.19	0.82	<i>Roseburia</i>	0.7	<i>Solibacillus</i>	0.2	1.0
<i>Anaerobacillus</i>	0.5	0.08	1.0	<i>Bifidobacterium</i>	1.0	<i>Janthinobacterium</i>	0.64	1.0
<i>Sediminibacterium</i>	1.0	0.51	1.0	<i>Clostridiaceae</i>	0.7	<i>Rubellimicrobium</i>	0.19	0.35
<i>Carnobacterium</i>	0.51	0.82	1.0	<i>Eggerthella</i>	0.7	<i>Dechloromonas</i>	0.65	0.38
<i>Ralstonia</i>	0.82	0.51	1.0	<i>Collinsella</i>	1.0	<i>Megamonas</i>	0.5	0.2
<i>Collinsella</i>	0.5	1.0	0.38	<i>Enhydrobacter</i>	1.0	<i>Brevibacillus</i>	0.65	0.2
<i>Bacteroides</i>	0.38	0.27	1.0	<i>Prevotella</i>	0.2	<i>Peptostreptococcus</i>	0.35	0.2
<i>Ochrobactrum</i>	0.66	0.27	0.66	<i>Parabacteroides</i>	0.4	<i>Pediococcus</i>	0.06	0.35
<i>Staphylococcus</i>	0.38	0.51	0.66	<i>Bradyrhizobium</i>	0.7	<i>Tissierella_Soehngenia</i>	0.06	0.35
<i>Pediococcus</i>	0.35	0.08	0.08	<i>Corynebacterium</i>	1.0	<i>Lactobacillaceae</i>	0.06	0.35
<i>Ureibacillus</i>	0.66	0.66	0.12	<i>Bacillales</i>	0.16	<i>Myxococcales</i>	0.2	0.35
<i>Enterobacteriaceae</i>	1.0	1.0	1.0	<i>Lysinibacillus</i>	1.0	<i>Jonesiaceae</i>	0.5	0.5
<i>Rs-045</i>	0.65	0.19	0.65	<i>Phascolarctobacterium</i>	0.0	<i>Slackia</i>	0.5	0.5
<i>Sinobacteraceae</i>	0.81	0.66	0.35	<i>Paenibacillus</i>	0.66	<i>Trichococcus</i>	0.5	0.5

<i>Clostridiales</i>	1.0	0.19	0.66	<i>Coprococcus</i>	0.4	<i>Roseomonas</i>	0.5	0.5
<i>Bradyrhizobiaceae</i>	1.0	0.82	0.38	<i>MLE1-12</i>	0.7	<i>Shewanella</i>	0.5	0.5
<i>Lysinibacillus</i>	0.35	0.48	0.5	<i>Oscillospira</i>	1.0	<i>TM7-1</i>	0.5	0.5
<i>Saccharopolyspora</i>	0.82	0.81	0.66	<i>Anaerostipes</i>	0.82	<i>Arthrobacter</i>	0.64	0.5
<i>Oxalobacteraceae</i>	0.82	0.51	1.0	<i>Sphingomonas</i>	0.82	<i>Planococcaceae</i>	NA	0.5
<i>Kurthia</i>	0.08	0.38	0.08	<i>Acinetobacter</i>	0.1	<i>ZB2</i>	NA	0.5
<i>Coriobacteriaceae</i>	0.66	0.65	0.38	<i>mitochondria</i>	0.38	<i>Brevundimonas</i>	NA	0.5
<i>Streptomyces</i>	0.27	0.27	0.27	<i>Staphylococcus</i>	0.66	<i>Rhodoplanes</i>	NA	0.5
<i>Vagococcus</i>	0.82	0.18	0.12	<i>Ochrobactrum</i>	0.4	<i>Phyllobacterium</i>	NA	0.5
<i>Tissierellaceae</i>	0.66	1.0	0.38	<i>Parachlamydia</i>	0.4	<i>Edaphobacter</i>	NA	1.0
<i>Unassigned</i>	0.18	0.26	1.0	<i>Thermogemmatisporaceae</i>	1.0	<i>Devosia</i>	NA	1.0
				<i>Morganella</i>	0.2	<i>Acetobacteraceae</i>	NA	1.0
				<i>Halomonas</i>	0.51	<i>Nocardiodiaceae</i>	0.5	1.0
				<i>Ellin329</i>	1.0	<i>Anaerococcus</i>	0.5	1.0
				<i>Stenotrophomonas</i>	0.51	<i>Agrobacterium</i>	0.5	1.0
				<i>Jeotgalicoccus</i>	1.0	<i>0319-6G20</i>	0.5	1.0
				<i>Ureibacillus</i>	0.51	<i>Rummeliibacillus</i>	0.08	NA
				<i>Pseudomonas</i>	1.0	<i>Solirubrobacterales</i>	0.5	NA
				<i>Leuconostocaceae</i>	1.0	<i>Gemmata</i>	0.5	NA
				<i>Parachlamydiateae</i>	0.4	<i>Gluconacetobacter</i>	0.5	NA
				<i>Christensenellaceae</i>	0.2	<i>Stenotrophomonas</i>	0.35	0.08
				<i>Solibacterales</i>	1.0	<i>Halomonas</i>	0.35	0.38
				<i>Natronobacillus</i>	0.35	<i>Tissierellaceae</i>	1.0	0.64
				<i>Sinobacteraceae</i>	0.2	<i>Ureibacillus</i>	0.7	1.0
				<i>Dietzia</i>	0.82	<i>Caldicoprobacter</i>	0.35	0.19
				<i>Paracoccus</i>	0.7	<i>Conexibacteraceae</i>	0.37	0.1
				<i>Rhodococcus</i>	0.66	<i>Lactococcus</i>	0.08	0.1
				<i>Lachnospiraceae;Other</i>	0.65	<i>SJA-4</i>	0.2	0.5
				<i>Dechloromonas</i>	0.5	<i>Barnesiellaceae</i>	0.2	1.0
				<i>Aneurinibacillus</i>	1.0	<i>Turicibacter</i>	1.0	1.0
				<i>Enterococcus</i>	0.16	<i>Lachnospiraceae;Other</i>	0.5	0.07
				<i>Prevotella</i>	0.2	<i>Sporosarcina</i>	0.38	0.1

<i>Barnesiellaceae</i>	0.64	<i>Bacillaceae;Other</i>	0.64	0.82
<i>Sporanaerobacter</i>	1.0	<i>Anoxybacillus</i>	1.0	0.06
<i>Actinomyces</i>	1.0	<i>Rhodococcus</i>	0.1	0.08
<i>Ellin6513</i>	0.38	<i>Paenibacillus</i>	1.0	0.08
<i>Rubellimicrobium</i>	0.5	<i>Bradyrhizobium</i>	1.0	0.66
<i>Chryseobacterium</i>	1.0	<i>Staphylococcus</i>	0.66	1.0
<i>Rhodocyclaceae</i>	0.5	<i>Paracoccus</i>	0.82	0.06
<i>Tepidimicrobium</i>	1.0	<i>Eggerthella</i>	0.2	0.06
<i>Pseudoramibacter_Eubacterium</i>	1.0	<i>Roseburia</i>	0.06	0.08
<i>Pseudomonadaceae</i>	1.0	<i>Eubacterium</i>	0.08	0.08
<i>Conexibacteraceae</i>	0.64	<i>Clostridium</i>	0.1	0.1
<i>Pirellulaceae</i>	1.0	<i>Clostridiales</i>	0.1	0.1
<i>Ruminococcus</i>	0.4	<i>Carnobacterium</i>	0.1	0.1
<i>Acidaminococcus</i>	1.0	<i>Methylobacterium</i>	0.2	0.2
<i>Tissierellaceae</i>	0.51	<i>Sphingobium</i>	0.06	0.4
<i>Turicibacter</i>	0.64	<i>Bacillus</i>	1.0	0.7
<i>Bilophila</i>	1.0	<i>MLE1-12</i>	0.18	0.7
<i>Slackia</i>	0.5	<i>Blautia</i>	0.4	0.7
<i>Megamonas</i>	0.5	<i>Novosphingobium</i>	0.35	0.19
		<i>Erwinia</i>	0.5	0.2
		<i>Peptoniphilus</i>	0.5	1.0
		<i>Phascolarctobacterium</i>	0.35	0.06
		<i>Anaerostipes</i>	0.35	0.2
		<i>Dietzia</i>	0.18	0.64
		<i>Sphingomonas</i>	0.7	1.0
		<i>Enhydrobacter</i>	1.0	0.06
		<i>Pseudoramibacter_Eubacterium</i>	0.06	0.06
		<i>Oscillospira</i>	0.06	0.06
		<i>Vagococcus</i>	0.1	0.08
		<i>Lysinibacillus</i>	0.18	0.08
		<i>Caloramator</i>	0.06	0.08
		<i>Dorea</i>	0.1	0.08

<i>Bifidobacteriaceae</i>	0.08	0.08
<i>Corynebacterium</i>	0.1	0.1
<i>Sporanaerobacter</i>	0.37	0.1
<i>Coriobacteriaceae</i>	0.1	0.1
<i>Lachnospiraceae</i>	0.08	0.1
<i>Leuconostocaceae</i>	0.1	0.1
<i>Bifidobacterium</i>	1.0	0.1
<i>Enterobacteriaceae</i>	0.08	0.1
<i>Streptococcus</i>	0.1	0.1
<i>Sediminibacterium</i>	0.4	0.1
<i>Alkaliphilus</i>	0.64	0.2
<i>Other</i>	0.08	0.2
<i>WPS-2</i>	1.0	0.35
<i>Prevotella</i>	0.06	0.35
<i>Micrococcaceae</i>	0.64	0.35
<i>Leucobacter</i>	0.35	0.35
<i>Microbacterium</i>	0.07	0.35
<i>Microbacteriaceae</i>	0.18	0.35
<i>Collinsella</i>	0.4	0.4
<i>Delftia</i>	0.7	0.4
<i>Solibacterales</i>	0.4	0.4
<i>Ralstonia</i>	0.7	0.4
<i>Veillonella</i>	0.64	0.5
<i>Rothia</i>	1.0	0.5
<i>Propionibacterium</i>	1.0	0.5
<i>Actinomycetales</i>	0.5	0.5
<i>Chryseobacterium</i>	0.5	0.5
<i>Pirellulaceae</i>	0.5	0.5
<i>Kurthia</i>	0.1	0.51
<i>Rhizobiales</i>	1.0	0.51
<i>Tepidimicrobium</i>	0.7	0.64
<i>Ellin329</i>	0.35	0.66

<i>Clostridiaceae</i>	0.08	0.7
<i>Bacillales</i>	0.7	0.7
<i>Geobacillus</i>	0.7	0.7
<i>Faecalibacterium</i>	0.18	0.7
<i>Caulobacteraceae</i>	0.7	0.7
<i>Pseudomonas</i>	1.0	1.0
<i>Bacillaceae</i>	0.7	1.0
<i>Kocuria</i>	0.35	1.0
<i>Oxalobacteraceae</i>	0.49	1.0
<i>Acidaminococcus</i>	0.5	1.0
<i>Bradyrhizobiaceae</i>	0.5	1.0
<i>Phyllobacteriaceae</i>	0.5	1.0
<i>Bilophila</i>	0.5	1.0
<i>Thermogemmatisporaceae</i>	0.82	1.0
<i>Aneurinibacillus</i>	0.7	1.0
<i>Actinomyces</i>	1.0	1.0
<i>Brevibacterium</i>	1.0	1.0
<i>Cloacibacterium</i>	1.0	1.0
<i>Jeotgalicoccus</i>	1.0	1.0
<i>Gemmataceae</i>	1.0	1.0
<i>Mycetocola</i>	0.06	0.06
<i>Rs-045</i>	0.06	0.06
<i>Streptomyces</i>	0.06	0.2
<i>Facklamia</i>	0.5	0.2
<i>Rathayibacter</i>	0.06	0.2
<i>Saccharopolyspora</i>	0.06	0.2
<i>Frigoribacterium</i>	0.06	0.2
<i>Sanguibacter</i>	0.2	0.5
<i>Agrococcus</i>	NA	0.5
<i>JG30-KF-CM45</i>	NA	0.5
<i>Paenibacillaceae</i>	NA	0.5
<i>Erysipelothrix</i>	NA	0.5

	<i>Isosphaeraceae</i>	NA	0.5
	<i>Macrococcus</i>	0.2	0.5
	<i>Williamsia</i>	0.5	0.5
	<i>Photobacterium</i>	0.5	0.5
	<i>Weissella</i>	0.19	NA
	<i>Cellulomonas</i>	0.5	NA
	<i>Proteiniclasticum</i>	0.5	NA
	<i>Peptococcus</i>	0.5	NA