

Table S1. Primers used in this study

| Primer | Accession No. | Sequence (5'-3') | Amplicon size (bp) |
|-----------|----------------|----------------------------|--------------------|
| tlr4-F | | ACAGATCACCTGGACAGCAAGA | |
| tlr4-R | AY388400.1 | CGGCTTGAAAGTCCCGCAT | 153 |
| myd88-F | | GGGACTGACACCTGAGACCTT | |
| myd88-R | DQ100359.1 | CGATAAGCTCACTGGCGATGG | 176 |
| nfkb-F | | AAGGAATCATGGCTGGCGAG | |
| nfkb-R | DQ066717 | GTAAGCAAGGCCATCAACTGC | 156 |
| il1β-F | | CGGCAGCTCCATAAACACCTTC | |
| il1β-R | BC098597 | CGCTCGGTGTCTTCCTGT | 108 |
| il6-F | | CAGCTGCAGGTGAGAGACG | |
| il6-R | NM_001261449 | GCAGGCGTCGATCATCACG | 132 |
| tnfa-F | | GCTTTCTGAATCCTACGGAGGC | |
| tnfa-R | AY427649.1 | CGTGTCTGTGCCAGTCTG | 155 |
| sod1-F | | AGACCTGGTAATGTGACCGC | |
| sod1-R | NM_131294.1 | GCCACCCTCCCCAAGT | 145 |
| sod2-F | | TGCAGAGTCGGATATGTTGGAG | |
| sod2-R | NM_199976.1 | GTGAGGCTCAAGTGCACCAT | 123 |
| hsp90-F | | CCAAACTGGACAGCGGCAA | |
| hsp90-R | L35587.1 | AGATCGGCTTGGTCATCCA | 106 |
| cat-F | | GATCGCTGTCCGCTTTC | |
| cat-R | NM_130912.2 | CAGTTGCCCTCATCGGTGT | 105 |
| gpx-F | | TGGCGTCGCTTGAGGC | |
| gpx-R | AY216589.1 | TCCTGGTGCCCGAACTG | 121 |
| muc2.2-F | | ACACGCTCAAGTAATCGCACAGTC | |
| muc2.2-R | XM 002667543 | TCAGCGAGTGTGGCTCACTT | 137 |
| mmp9-F | | CCAACATTAAAGATGCCCTGATGTAT | |
| mmp9-R | NM 213123.1 | CCC AGTGGTGGTCCGTGGTTGA | 146 |
| β-def-1-F | | TGTCTTGTGTAAGCATTGCAC | |
| β-def-1-R | NM 001081553.1 | ACACACTCCTGTCTGCAAACACC | 100 |
| β-actin-F | | AGGCTCCCTGAATCCAAA | |
| β-actin-R | AF057040.1 | GTCACACCATCACCAAGAGTCC | 160 |

Table S2. Relative abundance of zebrafish gut microbiome in phylum level

| Phylum ^{a)} | Relative abundance (%) | | | |
|----------------------|------------------------|-----------|------------|-----------|
| | Ctrl. | Otc | Smx/Tmp | Ery |
| Fusobacteria | 57.11±2.6 | 55.72±4.3 | 52.09±6.4 | 52.83±4.6 |
| Proteobacteria | 18.89±2.5 | 22.02±2.7 | 27.77±10.5 | 36.02±3.7 |
| Firmicutes | 13.26±1.5 | 11.91±2.1 | 13.78±5.7 | 5.01±2.3 |
| Bacteroidetes | 10.25±0.9 | 8.28±3.5 | 5.61±1.6 | 5.55±2.8 |
| Actinobacteria | 0.36±0.1 | 1.63±1.2 | 0.40±0.3 | 0.10±0.0 |
| Planctomycetes | 0.02±0.0 | 0.13±0.1 | 0.16±0.1 | 0.19±0.1 |
| Verrucomicrobia | 0.01±0.0 | 0.08±0.1 | 0.12±0.0 | 0.25±0.2 |
| Tenericutes | 0.09±0.1 | 0.22±0.4 | 0.02±0.0 | 0.001±0.0 |
| Chlamydiae | 0.00±0.0 | 0.001±0.0 | 0.03±0.0 | 0.03±0.0 |
| Armatimonadetes | 0.00±0.0 | 0.00±0.0 | 0.02±0.0 | 0.00±0.0 |
| Deinococcus-Thermus | 0.003±0.0 | 0.00±0.0 | 0.006±0.0 | 0.00±0.0 |
| Acidobacteria | 0.00±0.0 | 0.001±0.0 | 0.002±0.0 | 0.002±0.0 |
| Spirochaetes | 0.00±0.0 | 0.00±0.0 | 0.005±0.0 | 0.00±0.0 |
| Cyanobacteria | 0.00±0.0 | 0.00±0.0 | 0.00±0.0 | 0.004±0.0 |
| Other | 0.00±0.0 | 0.00±0.0 | 0.00±0.0 | 0.007±0.0 |

^{a)} Values are presented ad means±SD.

Table S3. Relative abundance of zebrafish gut microbiome in species level

| Species ^{a)} | Relative abundance (%) | | | |
|---|------------------------|-----------|-----------|-----------|
| | Ctrl. | Otc | Smx/Tmp | Ery |
| <i>Cetobacterium somerae</i> | 57.10±2.6 | 55.72±4.3 | 52.08±6.4 | 52.83±4.6 |
| <i>Aeromonas veronii</i> | 10.06±0.6 | 10.02±3.3 | 9.24±1.5 | 25.57±1.4 |
| KM585593 | 9.47±1.0 | 9.23±2.1 | 9.10±4.6 | 3.41±1.6 |
| <i>Cellvibrio fibrivorans</i> | 1.27±0.8 | 1.23±0.3 | 9.63±9.1 | 2.65±3.3 |
| <i>Shewanella xiamenensis</i> | 2.75±0.5 | 4.25±2.1 | 3.88±0.4 | 0.66±0.2 |
| KM585593_g_uc ^{b)} | 3.61±0.3 | 2.04±0.9 | 3.74±2.0 | 1.44±0.7 |
| Bacteroides_uc | 4.36±3.6 | 2.46±1.5 | 1.28±0.6 | 0.06±0.0 |
| Bacteroidaceae_uc | 3.75±2.1 | 0.72±0.6 | 0.30±0.1 | 0.01±0.0 |
| <i>Flavobacterium ginsenosidimutans</i> | 0.07±0.0 | 0.24±0.2 | 0.03±0.0 | 3.30±1.8 |
| <i>Flavobacterium granuli</i> | 0.18±0.0 | 2.13±2.2 | 0.30±0.2 | 0.41±0.2 |
| GQ360021_g_uc | 1.55±0.7 | 0.75±0.2 | 0.56±0.2 | 0.02±0.0 |
| <i>Gemmobacter fontiphilus</i> | 0.75±0.4 | 0.65±0.5 | 0.56±0.2 | 0.72±0.4 |
| Rhodobacteraceae_uc | 0.02±0.0 | 2.13±1.7 | 0.25±0.2 | 0.25±0.3 |
| <i>Flavobacterium succinicans</i> | 0.05±0.0 | 0.73±0.9 | 1.13±0.8 | 0.66±0.5 |
| <i>Aeromonas caviae</i> | 1.60±0.3 | 0.01±0.0 | 0.00±0.0 | 0.004±0.0 |
| <i>Flavobacterium cutihirudinis</i> | 0.002±0.0 | 0.01±0.0 | 1.44±0.8 | 0.01±0.0 |
| <i>Rhodococcus erythropolis</i> | 0.01±0.0 | 1.39±1.1 | 0.01±0.0 | 0.00±0.0 |
| <i>Rhizobium arenae</i> | 0.11±0.1 | 0.003±0.0 | 0.003±0.0 | 1.28±0.2 |
| <i>Kinneretia asaccharophila</i> | 0.96±0.4 | 0.01±0.0 | 0.29±0.1 | 0.03±0.0 |
| <i>Plesiomonas shigelloides</i> | 0.002±0.0 | 0.07±0.1 | 0.06±0.1 | 0.90±1.5 |
| <i>Flavobacterium notoginsengisoli</i> | 0.16±0.1 | 0.37±0.2 | 0.003±0.0 | 0.35±0.3 |
| Hyphomicrobium_uc | 0.002±0.0 | 0.35±0.2 | 0.042±0.3 | 0.08±0.0 |
| EU234324 | 0.001±0.0 | 0.35±0.2 | 0.25±0.1 | 0.19±0.1 |
| <i>Pseudomonas alcaligenes</i> | 0.002±0.0 | 0.00±0.0 | 0.00±0.0 | 0.77±0.6 |
| <i>Bacteroides vulgatus</i> | 0.02±0.0 | 0.61±1.0 | 0.08±0.1 | 0.05±0.0 |

^{a)} Values are presented as means±SD.

^{b)} ‘uc’ means unclassified.

Table S4. Number of goblet cells per villus

| Number of goblet cells (n = 10) ^{a)} | | | | <i>p</i> value ^{b)} | | |
|---|---------|---------|---------|------------------------------|---------------|-----------|
| Ctrl. | Otc | Smx/Tmp | Ery | Ctrl.-Otc | Ctrl.-Smx/Tmp | Ctrl.-Ery |
| 2.5±1.3 | 3.9±1.9 | 5.6±2.3 | 4.9±1.6 | 0.0696 | 0.0016 | 0.0016 |

^{a)} Values are presented as means±SD.

^{b)} Significance was determined by *t*-test.

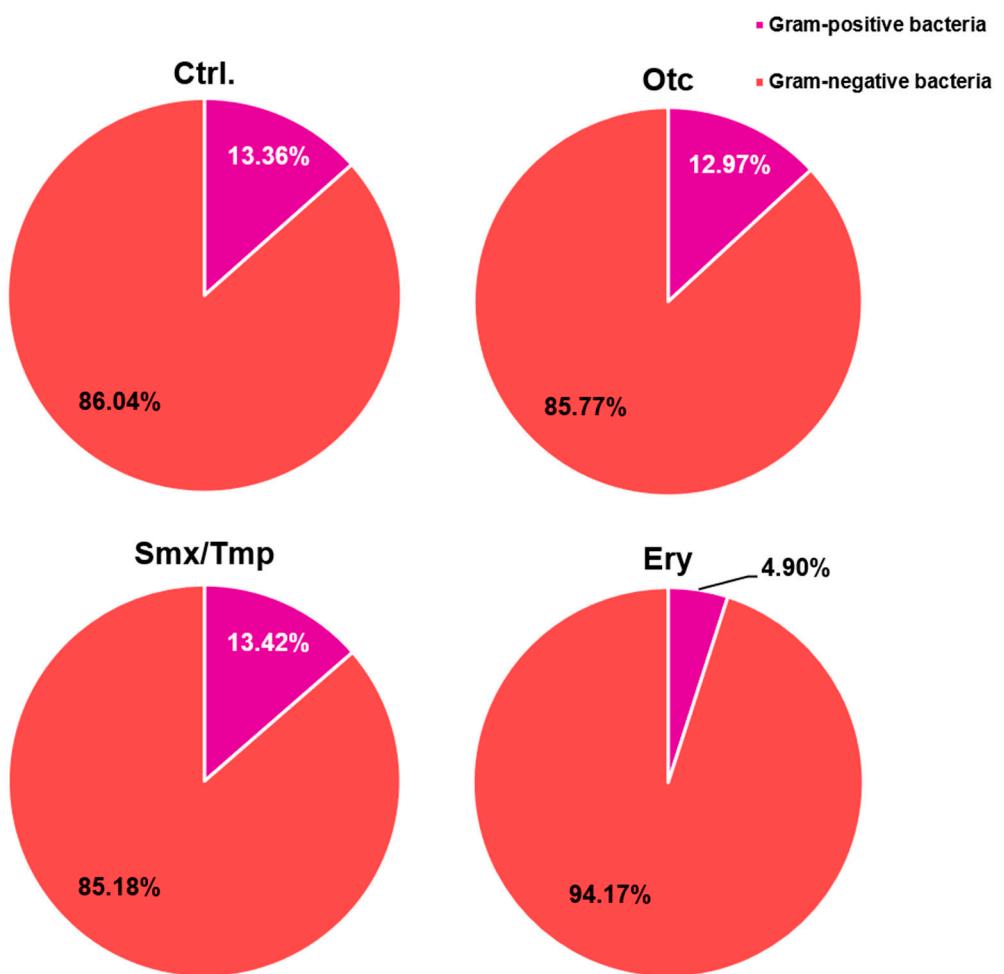


Figure S1. Relative abundance of Gram-positive bacteria and Gram-negative bacteria in zebrafish gut microbiome.