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## Supplementary information

2 **Table S1.** Outcome measures by treatment group and within participants taking metformin.

<b>Parameter</b>	<b>Time-point</b>	<b>All Participants</b>		<b>Participants on metformin</b>	
		<b>Probiotic (N=30)</b>	<b>Placebo (N=30)</b>	<b>Probiotic (N=14)</b>	<b>Placebo (N=14)</b>
Weight (kg)	<i>Baseline</i>	100.1±20.4	101.7±21.9	97.6±21.9	101.0±20.6
	<i>Week 12</i>	97.5±21.2	100.8±22.8	91.7±23.9	99.5±22.2
BMI (kg/m <sup>2</sup> )	<i>Baseline</i>	35.5±6.20	36.3±7.5	35.0±6.1	35.6±8.2
	<i>Week 12</i>	34.5±6.3	36.0±7.8	32.3±6.5	35.1±8.8
Waist (cm)	<i>Baseline</i>	114.7±12.9	114.3±14.7	113.7±13.6	113.4±12.8
	<i>Week 12</i>	114.3±16.0	116.0±17.1	109.4±18.0	116.1±16.2
Waist:Hip	<i>Baseline</i>	0.97±0.06	0.95±0.07	1.0±0.1	1.0±0.1
	<i>Week 12</i>	0.97±0.06	0.98±0.72	0.9±0.1	1.0±0.1
Body fat (%)	<i>Baseline</i>	37.7±8.2	41.8±7.9	39.5±9.1	40.3±7.8
	<i>Week 12</i>	38.1±7.8	40.9±8.4	37.3±8.5	38.9±8.7
Systolic BP (mmHg)	<i>Baseline</i>	133.0±10.8	127.8±12.5	134.5±14.4	130.4±12.3
	<i>Week 12</i>	128.5±12.5	127.0±13.1	129.0±17.9	127.5±11.3
Diastolic BP (mmHg)	<i>Baseline</i>	79.8±7.3	81.6±6.3	79.4±6.5	83.7±5.8
	<i>Week 12</i>	78.5±5.8	81.6±7.3	78.0±6.6	82.6±6.1
TC (mmol/L)	<i>Baseline</i>	4.9±1.1	5.4±1.3	4.7±1.0	5.1±1.1
	<i>Week 12</i>	4.9±1.1	5.3±1.4	4.4±0.9	4.7±0.8
HDL-c (mmol/L)	<i>Baseline</i>	1.2±0.3	1.3±0.2	1.3±0.3	1.3±0.2
	<i>Week 12</i>	1.2±0.3	1.2±0.2	1.2±0.3	1.2±0.2
LDL-c (mmol/L)	<i>Baseline</i>	2.8±1.1	3.2±1.0	2.7±0.9	3.0±1.1
	<i>Week 12</i>	2.9±1.1	3.2±1.1	2.4±1.0	2.7±0.8
Triglycerides (mmol/L)	<i>Baseline</i>	1.8±0.9	1.9±1.2	1.7±0.8	1.6±0.8
	<i>Week 12</i>	1.7±0.7	2.1±1.3	1.6±0.9	1.6±0.6
FFA (μmol/L)	<i>Baseline</i>	608.5±296.4	647.8±234.9	687.1±345.8	651.1±248.4
	<i>Week 12</i>	618.0±219.1	642.0±183.2	714.6±195.2	621.6±229.3
hs-CRP (mg/L)	<i>Baseline</i>	3.9±4.1	3.0±2.4	4.7±4.1	5.5±9.1
	<i>Week 12</i>	4.1±4.7	4.3±6.2	5.4±5.8	5.5±10.3
LPS (EU/mL)	<i>Baseline</i>	0.5±0.2	0.6±0.3	0.5±0.3	0.5±0.3
	<i>Week 12</i>	0.5 + 0.2	0.5±0.2	0.5±0.2	0.5±0.2
Zonulin (mg/dL)	<i>Baseline</i>	203.6±171.9	207.8±172.9	291.0±211.3	233.9±239.0
	<i>Week 12</i>	166.7 ± 112.6	136.3±79.9	197.9±119.3 <sup>a</sup>	151.6±71.8

3 Data are means ± SD or median (IQR). Difference between baseline and endpoint. P value obtained  
4 from repeated measure ANOVA<sup>a</sup> or Wilcoxon matched-pairs signed rank test<sup>b</sup> for the within-group  
5 comparisons. \**p* < 0.05. BMI: body mass index; BP: blood pressure; TC: total cholesterol; HDL-c: high-  
6 density lipoprotein cholesterol; LDL: low-density lipoprotein cholesterol; FFA: free fatty acids; hs-CRP:  
7 high-sensitive C-reactive protein; LPS: lipopolysaccharide.

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11 **Table S2.** Plasma SCFA concentrations by treatment group within the participants on metformin  
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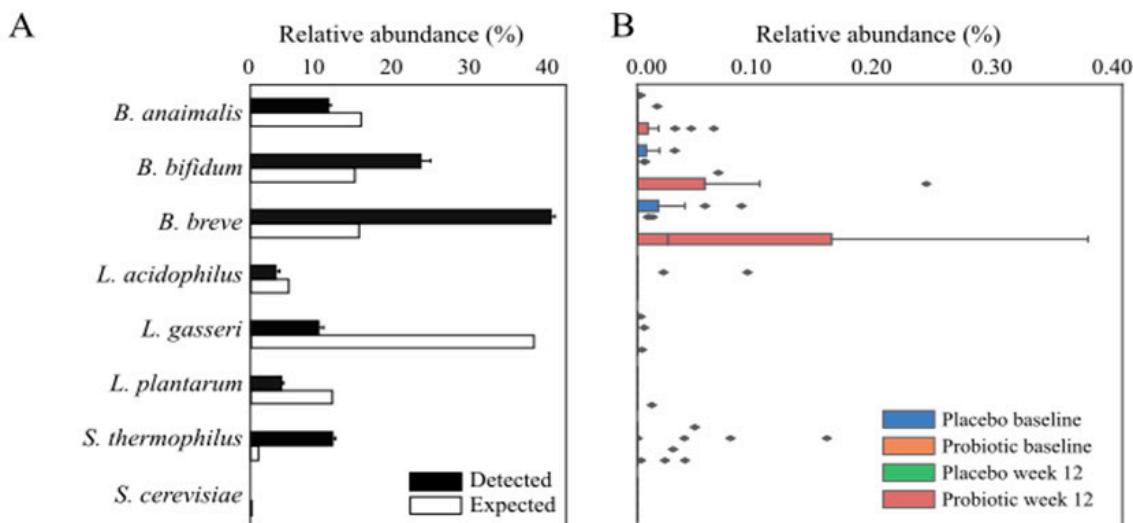
SCFA ( $\mu$ M)	Time-point	All participants		Participants on metformin	
		Probiotic (N=30)	Placebo (N=30)	Probiotic (N=13)	Placebo (N=14)
Propionate	Baseline	60.7 (70.0)	56.6 (85.2)	50.2 (57.4)	56.6 (83.8)
	Week 12	89.2 (79.8)	99.6 (80.1)	85.3 (80.3)	99.6 (81.6)
Isobutyrate	Baseline	6.1 (8.4)	6.0 (11.2)	5.4 (6.9)	6.0 (6.0)
	Week 12	6.9 (17.0)	7.0 (11.7)	6.6 (9.8)	7.0 (7.1)
Butyrate	Baseline	4.3 (4.9)	3.5 (7.3)	3.2 (4.3)	3.5 (5.3)
	Week 12	5.8 (8.3) <sup>a*</sup>	6.5 (7.8)	6.3 (9.1) <sup>a*</sup>	6.9 (4.6)
Isovalerate	Baseline	0.4 (2.2)	0.1 (2.1)	0.0 (1.1)	0.1 (1.6)
	Week 12	0.9 (4.7)	1.2 (3.6)	1.4 (3.0)	1.1 (5.8)
Total	Baseline	72.3 (86.6)	62.9 (111.1)	60.2 (67.3)	62.9 (95.6)
	Week 12	103.3 (113.3)	120.4 (99.3)	100.0 (102.4)	114.5 (98.2)

13 Data are median (IQR). <sup>a</sup>Difference between baseline and endpoint. P value obtained from Wilcoxon  
 14 matched-pairs signed rank test for the within-group comparisons. \* $p < 0.05$

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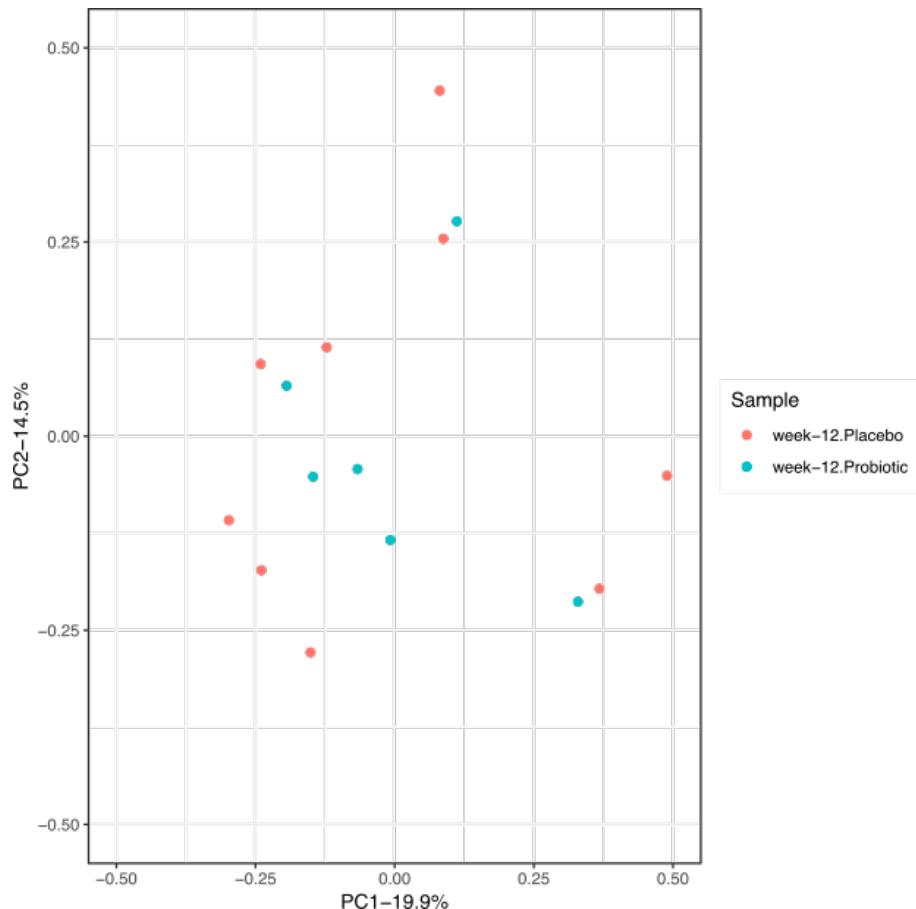


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20 **Figure S1.** Probiotic detection and quality control. A. Comparison of the relative abundance of species  
 21 in the capsule between the target formulation (Expected) and shotgun metagenomic sequencing  
 22 (Detected, n=2). B. Relative abundance of probiotic species found in the study groups at baseline and  
 23 after the 12-week intervention.

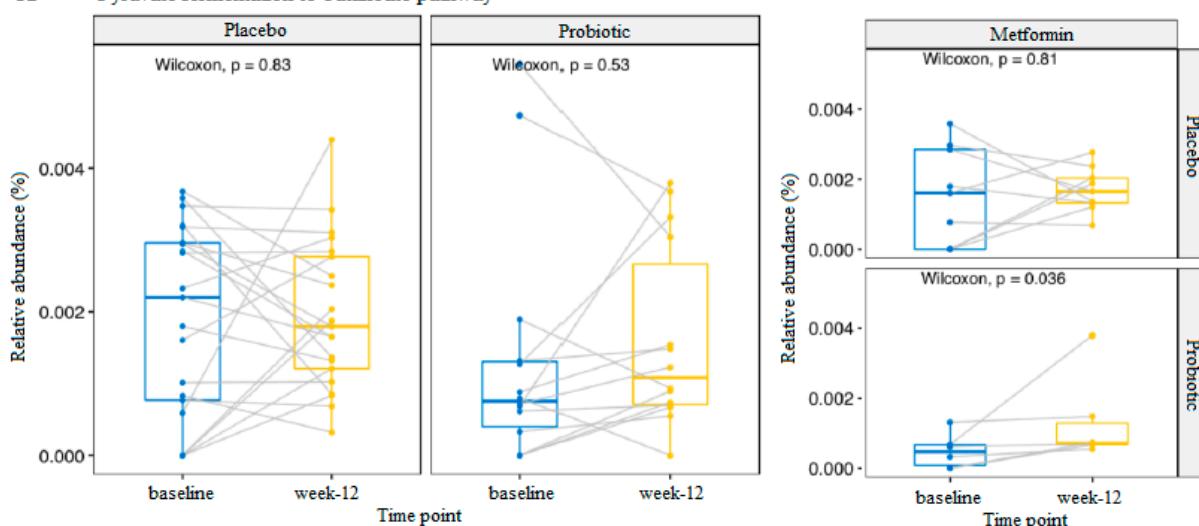
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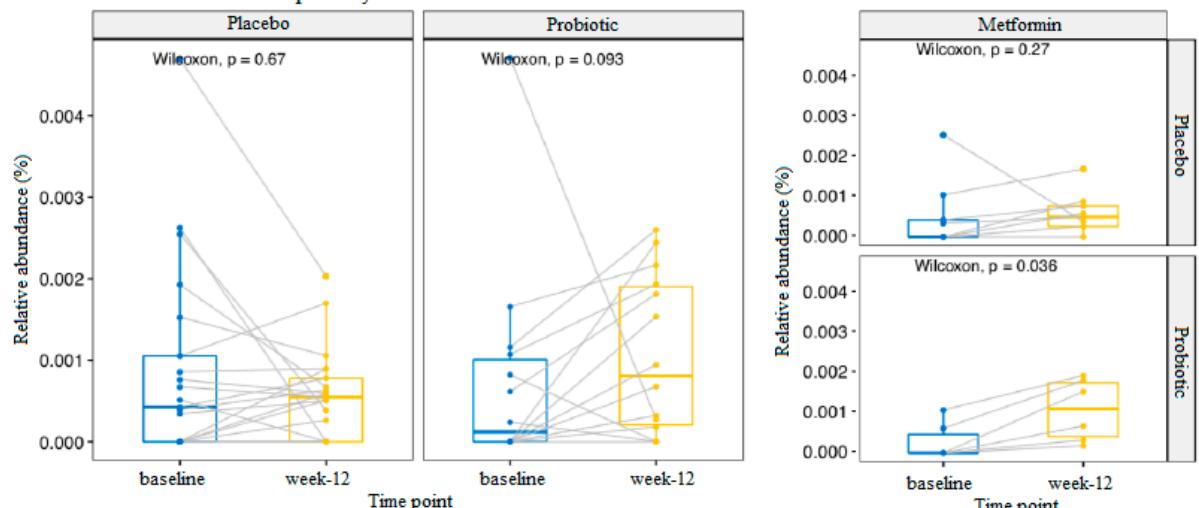
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26 **Figure S2.** PCoA of Bray-Curtis distances between intestinal microbial communities of probiotics and  
27 placebo groups in Metformin treated group after 12 weeks of intervention.

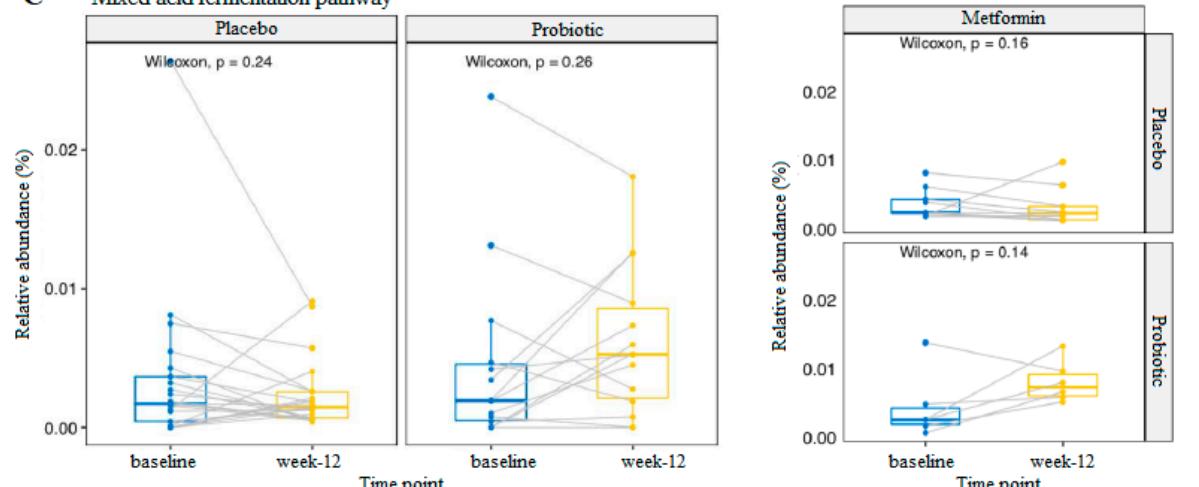
**A Pyruvate fermentation to butanoate pathway**



**B Bifidobacterium shunt pathway**



**C Mixed acid fermentation pathway**



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29 **Figure S3.** Relative abundance of SCFA-producing pathways in the faecal microbiota of participants in  
30 the intervention and metformin groups.  
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