

**Supplementary Table 1.** Information stated in the inclusion and exclusion criteria

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Inclusion criteria

- 1 Subjects who agree to participate in this study with a written informed consent.
- 2 Subjects who age is  $\geq 30$  years-old and  $< 70$  years-old.
- 3 Subjects who ALT level is  $\geq 25$  U/L and  $< 100$  U/L (men),  $\geq 17$  U/L and  $< 100$  U/L (women).
- 4 Subjects who BMI is  $\geq 23$  kg/m<sup>2</sup> and  $< 30$  kg/m<sup>2</sup>.

Exclusion criteria

- 1 Subjects who are under physician's advice, treatment, and/or medication for, hyperuricemia and/or dyslipidemia.
  - 2 Subjects with hepatic diseases except NAFLD (such as NASH, hepatitis C, hepatitis B, autoimmune hepatitis, liver cirrhosis and liver cancer etc.).
  - 3 Subjects who UA level is not less than the reference value ( $\geq 7.1$  mg/dL).
  - 4 Subjects who AST level is  $\geq 100$  U/L.
  - 5 Subjects who use pacemaker or defibrillator.
  - 6 Subjects with serious cerebrovascular, cardiac, hepatic, renal, gastrointestinal diseases, and/or affected with infectious diseases requiring reports to the authorities.
  - 7 Subjects with major surgical history relevant to the digestive system such as gastrectomy, gastrorrhaphy, enterectomy, etc.
  - 8 Subjects with unusually high and/or low blood pressure and/or abnormal hematological data.
  - 9 Subjects with severe anaemia.
  - 10 Pre- or post-menopausal women complaining of obvious physical changes.
  - 11 Subjects who are at risk of having allergic reactions to drugs or foods, especially salmon and/or milt.
  - 12 Subjects who regularly take medicine, functional foods, and/or supplements which would affect liver function.
  - 13 Subjects who regularly take medicine, functional foods, and/or supplements which would affect lipid metabolism.
  - 14 Heavy smokers, alcohol addicts or subjects with disordered lifestyle.
  - 15 Subjects who donated either 400 mL whole blood within 16 weeks (women), 12 weeks (men), 200 mL whole blood within 4 weeks (men and women), or blood components within 2 weeks (men and women), prior to the current study.
  - 16 Pregnant or lactating women or women who expect to be pregnant during this study.
  - 17 Subjects who currently participate in other clinical trials, or participated within the last 4 weeks prior to the current study.
  - 18 Any other medical and/or health reasons unfavorable to participation in the current study, as judged by the principal investigator.
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**Supplementary Table 2.** Basic information on the test food and placebo

	Placebo	Test food	Method
Description	Beige tablet	Beige tablet	
Weight (mg/tablet)	300	300	
Dextrin (mg/tablet)	290	107	
DNA (mg/tablet)	-	133	
Raw material	Fine silicon dioxide, calcium stearate, cacao pigment, caramel	Fine silicon dioxide, calcium stearate	
Nutrition facts			
Moisture (g/tablet)	0.016	0.017	Loss-on-drying method
Protein (g/tablet)	ND	0.156	Kjeldahl method
Lipid (g/tablet)	0.003	0.004	Acid hydrolysis method
Carbohydrate (g/tablet)	0.276	0.071	Carbohydrate = tablet - moisture - protein - lipid - ash
Ash (g/tablet)	0.006	0.053	Direct ashing method
Energy (kcal/tablet)	1.128	0.942	-
Sodium (mg/tablet)	0.014	9.700	Atomic absorption spectrophotometry

**Supplementary Table 3.** Effect of SM DNA on primary endpoints obtained after subgroup analysis based on BMI  $\geq 25 \text{ kg/m}^2$

	BMI $\geq 25 \text{ kg/m}^2$										BMI $< 25 \text{ kg/m}^2$														
	Placebo (n = 10)					SM DNA (n = 12)					Group	Period effect	P-value*			Placebo (n = 11)					Group	Period effect	P-value*		
	Placebo		SM DNA			Treatment effect		Interaction	Placebo			SM DNA			Treatment effect		Interaction	P-value*							
ALT (U/L)																									
Baseline	37.20	$\pm$	14.34	39.83	$\pm$	15.36	0.684	0.307	0.631	0.488		35.73	$\pm$	11.61	32.00	$\pm$	8.75	0.405	0.014	#	0.280	0.824			
$\Delta 4\text{wk}$	5.10	$\pm$	20.57	4.50	$\pm$	22.36	0.949					0.82	$\pm$	7.65	3.45	$\pm$	7.75	0.432							
$\Delta 8\text{wk}$	4.50	$\pm$	9.65	-2.67	$\pm$	13.52	0.176					-2.36	$\pm$	8.99	0.27	$\pm$	6.78	0.447							
$\Delta 12\text{wk}$	0.60	$\pm$	8.93	-0.42	$\pm$	18.13	0.873					-3.82	$\pm$	4.87	0.27	$\pm$	8.43	0.179							
AST (U/L)																									
Baseline	26.40	$\pm$	7.35	28.25	$\pm$	7.45	0.566	0.165	0.639	0.260		27.91	$\pm$	5.59	25.00	$\pm$	3.35	0.154	0.017	#	0.066	+	0.778		
$\Delta 4\text{wk}$	0.60	$\pm$	7.59	4.33	$\pm$	11.28	0.384					-0.91	$\pm$	5.01	1.09	$\pm$	2.55	0.252							
$\Delta 8\text{wk}$	0.70	$\pm$	5.25	-1.00	$\pm$	5.53	0.471					-2.91	$\pm$	4.18	-0.09	$\pm$	2.34	0.065	+						
$\Delta 12\text{wk}$	-1.20	$\pm$	4.42	0.17	$\pm$	5.56	0.537					-3.00	$\pm$	2.57	-1.18	$\pm$	2.48	0.107							
$\gamma$ -GTP (U/L)																									
Baseline	64.60	$\pm$	85.99	73.33	$\pm$	47.10	0.765	0.030	#	0.249	0.519		61.00	$\pm$	49.76	73.55	$\pm$	62.50	0.608	0.004	#	0.143	0.892		
$\Delta 4\text{wk}$	35.70	$\pm$	71.47	9.58	$\pm$	45.13	0.309					-3.18	$\pm$	12.69	8.64	$\pm$	23.08	0.152							
$\Delta 8\text{wk}$	11.50	$\pm$	27.73	-9.33	$\pm$	32.75	0.127					-10.55	$\pm$	19.78	-0.73	$\pm$	14.57	0.200							
$\Delta 12\text{wk}$	2.50	$\pm$	8.36	-6.42	$\pm$	48.15	0.571					-18.45	$\pm$	27.29	-8.91	$\pm$	13.94	0.318							
ALP (U/L)																									
Baseline	238.00	$\pm$	59.47	238.92	$\pm$	94.85	0.979	0.206	0.729	0.755		229.36	$\pm$	56.28	232.55	$\pm$	64.88	0.903	0.422		0.606	0.801			
$\Delta 4\text{wk}$	17.20	$\pm$	39.81	6.92	$\pm$	56.17	0.633					4.09	$\pm$	14.00	6.82	$\pm$	28.85	0.781							
$\Delta 8\text{wk}$	1.80	$\pm$	21.56	-1.00	$\pm$	41.80	0.850					-2.27	$\pm$	19.27	-0.91	$\pm$	25.66	0.889							
$\Delta 12\text{wk}$	0.00	$\pm$	19.64	-2.33	$\pm$	43.85	0.878					-3.82	$\pm$	24.40	3.55	$\pm$	24.70	0.490							
LDH (U/L)																									
Baseline	173.00	$\pm$	31.85	181.75	$\pm$	23.02	0.464	0.071	+	0.781	0.437		196.73	$\pm$	32.71	174.55	$\pm$	14.46	0.059	+	0.018	#	0.202	0.600	
$\Delta 4\text{wk}$	8.00	$\pm$	14.12	9.58	$\pm$	11.53	0.775					0.27	$\pm$	11.11	5.09	$\pm$	12.32	0.347							
$\Delta 8\text{wk}$	3.90	$\pm$	13.49	-2.42	$\pm$	16.11	0.337					-10.09	$\pm$	10.90	-2.36	$\pm$	6.80	0.060	+						
$\Delta 12\text{wk}$	2.10	$\pm$	7.62	3.67	$\pm$	12.38	0.731					2.45	$\pm$	16.51	4.00	$\pm$	13.74	0.814							
ChE (U/L)																									
Baseline	395.10	$\pm$	62.65	409.92	$\pm$	46.86	0.533	0.041	#	0.747	0.511		398.27	$\pm$	80.04	422.64	$\pm$	70.32	0.457	0.541		0.877	0.751		
$\Delta 4\text{wk}$	17.80	$\pm$	24.99	11.58	$\pm$	17.40	0.500					6.64	$\pm$	35.87	8.27	$\pm$	35.21	0.915							
$\Delta 8\text{wk}$	14.90	$\pm$	16.32	10.25	$\pm$	20.67	0.571					5.55	$\pm$	29.91	4.18	$\pm$	19.10	0.900							
$\Delta 12\text{wk}$	1.40	$\pm$	28.08	4.42	$\pm$	24.45	0.790					4.45	$\pm$	22.36	-0.64	$\pm$	22.93	0.604							
AST/ALT ratio																									
Baseline	0.75	$\pm$	0.20	0.78	$\pm$	0.25	0.748	0.970	0.777	0.418		0.83	$\pm$	0.19	0.81	$\pm$	0.15	0.846	0.220		0.699	0.245			
$\Delta 4\text{wk}$	-0.04	$\pm$	0.20	-0.01	$\pm$	0.22	0.777					-0.07	$\pm$	0.19	-0.05	$\pm$	0.12	0.759							
$\Delta 8\text{wk}$	-0.06	$\pm$	0.13	0.00	$\pm$	0.16	0.387					-0.05	$\pm$	0.12	0.02	$\pm$	0.11	0.206							
$\Delta 12\text{wk}$	-0.02	$\pm$	0.08	-0.04	$\pm$	0.22	0.755					0.02	$\pm$	0.07	-0.02	$\pm$	0.13	0.391							
TIBC (μg/dL)																									
Baseline	338.50	$\pm$	23.21	323.50	$\pm$	40.92	0.317	0.006	#	0.139	0.022	#	320.36	$\pm$	45.01	350.27	$\pm$	48.34	0.149	0.081	+	0.392	0.257		
$\Delta 4\text{wk}$	12.90	$\pm$	11.88	8.75	$\pm$	15.00	0.487					9.91	$\pm$	25.56	5.00	$\pm$	27.04	0.666							
$\Delta 8\text{wk}$	8.80	$\pm$	13.69	13.75	$\pm$	9.28	0.326					10.45	$\pm$	19.77	-2.64	$\pm$	14.24	0.090	+						
$\Delta 12\text{wk}$	-7.30	$\pm$	13.51	8.83	$\pm$	12.60	0.009	#				2.27	$\pm$	21.90	-1.45	$\pm$	14.40	0.642							
UIBC (μg/dL)																									
Baseline	208.50	$\pm$	50.62	205.42	$\pm$	57.20	0.896	0.376	0.993	0.702		207.00	$\pm$	63.30	226.64	$\pm$	70.92	0.501	0.835		0.993	0.264			
$\Delta 4\text{wk}$	23.50	$\pm$	75.22	19.50	$\pm$	44.28	0.878					9.18	$\pm$	28.90	18.36	$\pm$	60.08	0.653							
$\Delta 8\text{wk}$	1.70	$\pm$	60.91	11.92	$\pm$	41.30	0.645					23.82	$\pm$	39.96	9.27	$\pm$	51.64	0.469							
$\Delta 12\text{wk}$	11.00	$\pm$	37.96	5.25	$\pm$	49.65	0.767					9.55	$\pm$	39.68	14.45	$\pm$	47.63	0.796							
Ferritin (ng/mL)																									
Baseline	150.87	$\pm$	97.97	179.27	$\pm$	106.56	0.526	0.036	#	0.723	0.723		93.40	$\pm$	52.65	149.37	$\pm$	75.67	0.058	+	0.621	0.136	0.136		
$\Delta 12\text{wk}$	-17.49	$\pm$	19.40	-12.67	$\pm$	38.51	0.723					-9.15	$\pm$	20.06	4.68	$\pm$	21.64	0.136							

L/S ratio																					
Baseline	1.09	±	0.22	0.97	±	0.33	0.317	0.232	0.031	#	0.031	#	1.13	±	0.21	1.07	±	0.21	0.491	0.038	
Δ12wk	-0.03	±	0.12	0.11	±	0.16	0.031	#					0.04	±	0.05	0.05	±	0.11	0.755	#	
TP (g/dL)																					
Baseline	7.37	±	0.35	7.21	±	0.24	0.209	0.071	†	0.246	0.015	#	7.26	±	0.34	7.57	±	0.37	0.056	†	
Δ4wk	0.07	±	0.29	0.08	±	0.24	0.909						-0.07	±	0.29	-0.05	±	0.35	0.843		
Δ8wk	0.07	±	0.31	0.09	±	0.19	0.848						-0.06	±	0.28	-0.11	±	0.25	0.693		
Δ12wk	-0.16	±	0.20	0.12	±	0.21	0.005	#					0.00	±	0.24	0.01	±	0.18	0.920		
Alb (g/dL)																					
Baseline	4.50	±	0.27	4.50	±	0.20	1.000	0.117		0.630	0.003	#	4.57	±	0.26	4.61	±	0.21	0.724	0.310	
Δ4wk	0.15	±	0.19	0.03	±	0.17	0.115						0.02	±	0.19	0.07	±	0.21	0.529		
Δ8wk	0.19	±	0.25	0.10	±	0.15	0.329						0.01	±	0.23	0.02	±	0.17	0.915		
Δ12wk	0.01	±	0.17	0.13	±	0.13	0.065	†					0.05	±	0.23	0.09	±	0.16	0.604		
A/G ratio																					
Baseline	1.60	±	0.25	1.67	±	0.13	0.396	0.002	#	0.066	†	0.057	†	1.71	±	0.18	1.58	±	0.22	0.143	0.472
Δ4wk	0.09	±	0.07	-0.03	±	0.08	0.002	#					0.07	±	0.13	0.11	±	0.10	0.416		
Δ8wk	0.13	±	0.07	0.05	±	0.14	0.131						0.05	±	0.12	0.09	±	0.09	0.405		
Δ12wk	0.09	±	0.09	0.06	±	0.11	0.488						0.05	±	0.16	0.07	±	0.09	0.693		
TNF-α (pg/mL)																					
Baseline	2.67	±	0.58	2.68	±	0.50	0.970	<0.001	#	0.307	0.307		2.23	±	0.41	2.55	±	1.01	0.349	<0.001	
Δ12wk	0.60	±	0.43	0.40	±	0.47	0.307						0.40	±	0.34	0.65	±	0.50	0.197	#	
IL-1β (pg/mL)																					
Baseline	0.47	±	0.18	0.46	±	0.18	0.930	<0.001	#	0.841	0.841		0.40	±	0.13	0.64	±	0.43	0.095	†	
Δ12wk	-0.28	±	0.07	-0.27	±	0.11	0.841						-0.28	±	0.05	-0.25	±	0.17	0.624		
IL-6 (pg/mL)																					
Baseline	1.53	±	0.74	1.70	±	0.88	0.643	<0.001	#	0.581	0.581		1.48	±	0.90	2.22	±	1.21	0.123	<0.001	
Δ12wk	-0.47	±	0.34	-0.56	±	0.39	0.581						-0.47	±	0.52	-0.83	±	0.64	0.162	#	
Body weight (kg)																					
Baseline	71.85	±	7.46	73.63	±	7.86	0.594	0.124		0.157	0.256		63.95	±	8.66	65.63	±	5.30	0.591	0.001	
Δ4wk	0.94	±	1.91	0.37	±	1.00	0.377						0.26	±	0.76	-0.02	±	0.61	0.349		
Δ8wk	0.48	±	1.45	-0.07	±	1.11	0.327						0.09	±	0.85	-0.28	±	1.05	0.373		
Δ12wk	0.88	±	1.62	-0.33	±	1.07	0.049	#					0.02	±	1.03	-1.12	±	1.05	0.019	#	
BMI (kg/m <sup>2</sup> )																					
Baseline	26.49	±	1.21	27.29	±	1.30	0.152	0.112		0.143	0.215		23.99	±	0.90	23.84	±	0.80	0.675	0.002	
Δ4wk	0.35	±	0.67	0.14	±	0.37	0.365						0.09	±	0.30	-0.03	±	0.22	0.306		
Δ8wk	0.19	±	0.53	-0.03	±	0.41	0.298						0.05	±	0.31	-0.09	±	0.40	0.380		
Δ12wk	0.34	±	0.65	-0.13	±	0.38	0.049	#					0.01	±	0.39	-0.40	±	0.37	0.021	#	
Body fat percentage (%)																					
Baseline	31.46	±	6.74	31.81	±	6.10	0.900	0.026	#	0.497	0.770		30.51	±	6.49	26.18	±	5.87	0.116	<0.001	
Δ4wk	0.42	±	0.55	0.23	±	0.98	0.598						0.89	±	1.04	0.65	±	0.72	0.542		
Δ8wk	0.60	±	0.67	0.25	±	1.20	0.422						0.40	±	1.39	0.46	±	0.83	0.897		
Δ12wk	-0.03	±	0.90	-0.18	±	0.97	0.707						0.31	±	1.26	-0.25	±	1.05	0.277		
Adiponectin (μg/mL)																					
Baseline	8.81	±	3.24	6.89	±	2.08	0.127	0.037	#	0.174	0.174		10.41	±	5.25	8.17	±	2.72	0.229	0.132	
Δ12wk	-0.59	±	0.78	-0.13	±	0.73	0.174						-0.71	±	1.23	-0.01	±	0.88	0.142	#	
Leptin (ng/mL)																					
Baseline	19.53	±	11.36	22.48	±	16.95	0.645	0.354		0.340	0.340		15.27	±	7.49	12.01	±	4.95	0.242	<0.001	
Δ12wk	0.05	±	10.92	-3.26	±	3.99	0.340						-0.41	±	2.12	-2.71	±	1.24	0.007	#	

Values are presented as the mean ± SD.

\*P-values were determined by independent sample t-tests; †: P < 0.10, #: P < 0.05.

**Supplementary Table 4.** Effects of SM DNA on vital signs and blood parameters

	Placebo (n = 22–25)		SM DNA (n = 24)		P-value*		
Systolic blood pressure (mmHg)							
Baseline	129.96	±	17.08	132.13	±	18.05	0.668
Δ4wk	0.39	±	10.50	2.17	±	9.48	0.545
Δ8wk	-1.48	±	10.00	0.46	±	8.81	0.484
Δ12wk	-4.36	±	10.05	-0.83	±	9.11	0.218
Diastolic blood pressure (mmHg)							
Baseline	81.24	±	11.26	85.63	±	10.95	0.174
Δ4wk	-1.96	±	7.32	1.42	±	5.77	0.085†
Δ8wk	-3.70	±	4.92	0.17	±	6.38	0.025#
Δ12wk	-3.50	±	6.86	-0.33	±	5.08	0.080†
Pulse rate (times/min)							
Baseline	74.96	±	14.31	74.79	±	10.49	0.963
Δ4wk	-5.96	±	8.74	-2.17	±	8.58	0.141
Δ8wk	-1.35	±	9.45	-1.79	±	10.98	0.883
Δ12wk	-4.59	±	9.94	-1.63	±	8.27	0.276
WBC ( $\times 10^3/\mu\text{L}$ )							
Baseline	5.89	±	1.25	5.33	±	1.03	0.090†
Δ4wk	-0.31	±	0.67	-0.13	±	1.11	0.509
Δ8wk	0.32	±	0.80	-0.04	±	1.03	0.185
Δ12wk	-0.09	±	0.79	0.30	±	1.26	0.219
RBC ( $\times 10^4/\mu\text{L}$ )							
Baseline	488.00	±	55.65	484.33	±	39.04	0.791
Δ4wk	4.91	±	16.71	5.79	±	18.49	0.865
Δ8wk	12.35	±	18.04	11.38	±	12.57	0.832
Δ12wk	7.55	±	18.75	10.33	±	17.08	0.600
Hb (g/dL)							
Baseline	14.58	±	1.69	14.85	±	1.25	0.536
Δ4wk	0.14	±	0.43	0.10	±	0.55	0.765
Δ8wk	0.42	±	0.58	0.34	±	0.41	0.586
Δ12wk	0.19	±	0.62	0.27	±	0.47	0.600
Ht (%)							
Baseline	44.45	±	4.82	44.94	±	3.94	0.702
Δ4wk	0.93	±	1.68	0.97	±	1.81	0.931
Δ8wk	2.09	±	1.88	2.03	±	1.24	0.902
Δ12wk	0.52	±	1.83	0.78	±	1.32	0.585
Plt ( $\times 10^3/\mu\text{L}$ )							

	Baseline	26.47	$\pm$	5.79	24.55	$\pm$	4.88	0.218
	$\Delta$ 4wk	-0.08	$\pm$	2.36	0.14	$\pm$	3.88	0.811
	$\Delta$ 8wk	0.16	$\pm$	2.17	0.03	$\pm$	2.31	0.842
	$\Delta$ 12wk	-0.99	$\pm$	1.98	-0.65	$\pm$	2.81	0.635
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BUN (mg/dL)	Baseline	13.65	$\pm$	3.88	14.43	$\pm$	2.98	0.434
	$\Delta$ 4wk	-0.93	$\pm$	3.13	0.10	$\pm$	2.54	0.219
	$\Delta$ 8wk	-0.56	$\pm$	3.35	-0.42	$\pm$	2.77	0.877
	$\Delta$ 12wk	-0.43	$\pm$	2.94	0.05	$\pm$	2.26	0.535
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CRE (mg/dL)	Baseline	0.79	$\pm$	0.17	0.80	$\pm$	0.15	0.756
	$\Delta$ 4wk	0.02	$\pm$	0.05	0.01	$\pm$	0.04	0.601
	$\Delta$ 8wk	0.02	$\pm$	0.06	0.03	$\pm$	0.04	0.777
	$\Delta$ 12wk	0.00	$\pm$	0.04	0.03	$\pm$	0.05	0.037#
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UA (mg/dL)	Baseline	5.36	$\pm$	0.99	5.69	$\pm$	1.41	0.344
	$\Delta$ 4wk	-0.02	$\pm$	0.59	0.08	$\pm$	0.67	0.605
	$\Delta$ 8wk	0.02	$\pm$	0.62	0.02	$\pm$	0.82	0.997
	$\Delta$ 12wk	-0.05	$\pm$	0.69	0.15	$\pm$	0.70	0.334
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T-Cho (mg/dL)	Baseline	217.80	$\pm$	44.00	225.83	$\pm$	35.75	0.488
	$\Delta$ 4wk	8.61	$\pm$	19.62	9.29	$\pm$	21.71	0.911
	$\Delta$ 8wk	12.83	$\pm$	19.01	6.29	$\pm$	18.45	0.238
	$\Delta$ 12wk	7.82	$\pm$	18.05	9.71	$\pm$	22.23	0.754
<hr/>								
HDL-Cho (mg/dL)	Baseline	59.80	$\pm$	12.33	58.83	$\pm$	12.59	0.787
	$\Delta$ 4wk	-0.61	$\pm$	5.37	2.00	$\pm$	7.00	0.160
	$\Delta$ 8wk	-0.09	$\pm$	6.46	0.29	$\pm$	4.74	0.819
	$\Delta$ 12wk	0.73	$\pm$	6.30	3.00	$\pm$	4.24	0.155
<hr/>								
LDL-Cho (mg/dL)	Baseline	140.04	$\pm$	44.27	145.58	$\pm$	31.43	0.617
	$\Delta$ 4wk	5.61	$\pm$	21.54	2.92	$\pm$	19.01	0.651
	$\Delta$ 8wk	5.17	$\pm$	18.22	0.50	$\pm$	13.78	0.325
	$\Delta$ 12wk	3.55	$\pm$	16.29	4.42	$\pm$	16.92	0.860
<hr/>								
TG (mg/dL)	Baseline	117.80	$\pm$	64.86	133.33	$\pm$	66.88	0.413
	$\Delta$ 4wk	9.00	$\pm$	54.93	15.63	$\pm$	83.20	0.750
	$\Delta$ 8wk	-2.00	$\pm$	37.72	-6.38	$\pm$	44.33	0.718
	$\Delta$ 12wk	6.45	$\pm$	54.81	0.79	$\pm$	53.30	0.724

Fasting blood glucose (mg/dL)				
Baseline	91.92	±	10.68	97.38
Δ4wk	1.35	±	5.10	-0.63
Δ8wk	1.22	±	6.01	-2.04
Δ12wk	2.27	±	3.56	-0.79
				± 11.57
				0.093†
				0.231
				0.057†
				0.037#
HbA1c (%)				
Baseline	5.61	±	0.36	5.61
Δ4wk	-0.06	±	0.11	0.00
Δ8wk	-0.05	±	0.09	-0.02
Δ12wk	-0.10	±	0.10	-0.09
				± 0.43
				0.998
				0.123
				0.441
				0.725
HOMA-IR				
Baseline	1.51	±	0.59	2.01
Δ4wk	0.15	±	0.60	-0.22
Δ8wk	0.03	±	0.49	-0.44
Δ12wk	0.15	±	0.68	-0.12
				± 0.92
				0.026#
				0.103
				0.004#
				0.207

Values are presented as the mean ± SD.

\*P-values are determined by independent sample t-tests; †: P < 0.10, #: P < 0.05.

**Supplementary Table 5.** Effect of SM DNA on primary endpoints obtained after subgroup analysis based on the sex of the participants

	Male					Female					P-value*			
	Placebo (n = 11)			SM DNA (n = 16)		P-value*	Placebo (n = 10)			SM DNA (n = 7)				
ALT (U/L)														
Baseline	39.91	±	14.58	38.75	±	12.61	0.827	32.60	±	9.45	30.00	±	12.60	0.633
Δ4wk	5.00	±	20.00	3.75	±	16.01	0.858	0.50	±	6.54	4.57	±	19.41	0.610
Δ8wk	0.18	±	10.86	-0.38	±	11.67	0.901	1.70	±	8.83	-3.29	±	8.52	0.263
Δ12wk	-0.73	±	8.49	-0.50	±	16.67	0.963	-2.80	±	5.92	0.86	±	4.98	0.202
AST (U/L)														
Baseline	28.27	±	6.00	27.06	±	4.28	0.545	26.00	±	6.86	25.86	±	9.14	0.971
Δ4wk	1.00	±	7.85	3.50	±	6.30	0.368	-1.50	±	3.84	1.14	±	12.27	0.599
Δ8wk	-1.09	±	5.87	0.19	±	3.29	0.523	-1.30	±	4.03	-2.29	±	5.82	0.684
Δ12wk	-1.09	±	3.91	0.13	±	4.80	0.493	-3.30	±	2.98	-1.86	±	2.79	0.330
γ-GTP (U/L)														
Baseline	73.64	±	85.05	83.25	±	60.20	0.733	50.70	±	42.53	51.00	±	25.94	0.987
Δ4wk	30.18	±	69.92	4.44	±	28.40	0.269	-1.00	±	12.99	19.86	±	49.13	0.311
Δ8wk	2.91	±	33.69	-6.25	±	29.40	0.460	-3.30	±	14.24	-2.86	±	14.84	0.951
Δ12wk	-3.91	±	21.94	-10.56	±	41.26	0.592	-13.50	±	23.76	-0.86	±	15.77	0.239
ALP (U/L)														
Baseline	233.91	±	46.84	222.13	±	69.88	0.630	233.00	±	68.27	267.29	±	98.42	0.408
Δ4wk	14.09	±	38.56	2.31	±	31.32	0.390	6.20	±	14.65	17.29	±	67.36	0.682
Δ8wk	-6.73	±	23.07	1.94	±	39.35	0.519	6.70	±	13.82	-7.57	±	18.85	0.091†
Δ12wk	-4.91	±	21.97	2.31	±	39.99	0.592	1.20	±	22.30	-3.71	±	23.39	0.667
LDH (U/L)														
Baseline	186.27	±	33.09	176.50	±	19.34	0.342	184.50	±	36.23	182.43	±	20.16	0.893
Δ4wk	7.00	±	14.99	8.94	±	9.41	0.683	0.60	±	9.89	4.00	±	16.58	0.603
Δ8wk	-5.00	±	14.86	-1.63	±	8.76	0.464	-1.70	±	13.28	-4.14	±	18.86	0.757
Δ12wk	0.45	±	6.01	4.81	±	13.98	0.280	4.30	±	17.67	1.57	±	9.93	0.718
ChE (U/L)														
Baseline	414.55	±	54.17	416.75	±	64.54	0.927	377.20	±	83.53	414.29	±	44.76	0.303
Δ4wk	13.36	±	22.59	13.69	±	28.51	0.975	10.40	±	39.43	1.57	±	21.91	0.601
Δ8wk	5.64	±	18.23	7.44	±	20.89	0.819	14.80	±	29.88	7.14	±	18.32	0.557
Δ12wk	2.18	±	21.58	-0.56	±	25.76	0.774	3.90	±	28.82	7.86	±	16.76	0.750
AST/ALT ratio														
Baseline	0.76	±	0.19	0.75	±	0.19	0.879	0.82	±	0.20	0.91	±	0.21	0.414
Δ4wk	-0.03	±	0.18	-0.01	±	0.16	0.739	-0.08	±	0.20	-0.08	±	0.20	0.985
Δ8wk	-0.04	±	0.13	-0.01	±	0.12	0.466	-0.07	±	0.13	0.04	±	0.18	0.183
Δ12wk	0.00	±	0.07	0.00	±	0.19	0.943	0.00	±	0.09	-0.11	±	0.15	0.093 †

TIBC (μg/dL)																
	Baseline	347.09	±	23.42	337.50	±	50.66	0.515	309.10	±	39.14	333.57	±	34.94	0.205	
Δ4wk		11.45	±	15.87	7.13	±	22.43	0.587	11.20	±	24.34	6.57	±	19.65	0.683	
Δ8wk		11.73	±	11.34	6.06	±	15.99	0.322	7.40	±	21.67	5.57	±	10.67	0.840	
Δ12wk		-5.64	±	13.63	1.69	±	14.46	0.198	1.40	±	23.10	9.00	±	13.09	0.446	
UIBC (μg/dL)																
	Baseline	219.09	±	58.47	202.69	±	69.52	0.527	195.20	±	53.70	245.00	±	35.68	0.005 #	
Δ4wk		19.64	±	71.08	29.38	±	52.03	0.684	12.00	±	32.51	-4.86	±	43.53	0.374	
Δ8wk		14.18	±	59.83	17.19	±	51.22	0.890	12.30	±	42.30	-4.29	±	25.39	0.371	
Δ12wk		10.82	±	38.85	21.25	±	52.43	0.580	9.60	±	38.91	-16.86	±	18.22	0.117	
Ferritin (ng/mL)																
	Baseline	150.76	±	97.11	192.91	±	93.25	0.267	87.77	±	42.57	101.10	±	51.33	0.568	
Δ12wk		-11.46	±	22.86	-3.68	±	38.45	0.553	-14.95	±	16.62	-5.96	±	9.31	0.217	
L/S ratio																
	Baseline	1.13	±	0.19	0.93	±	0.26	0.047	#	1.09	±	0.23	1.20	±	0.22	0.363
Δ12wk		-0.01	±	0.11	0.11	±	0.15	0.034	#	0.01	±	0.07	0.00	±	0.05	0.680
TP (g/dL)																
	Baseline	7.46	±	0.28	7.33	±	0.34	0.295		7.15	±	0.33	7.50	±	0.39	0.065 +
Δ4wk		-0.07	±	0.25	0.02	±	0.26	0.369		0.07	±	0.33	0.03	±	0.40	0.818
Δ8wk		-0.11	±	0.24	0.01	±	0.25	0.246		0.12	±	0.30	-0.03	±	0.23	0.293
Δ12wk		-0.17	±	0.16	0.08	±	0.21	0.002	#	0.03	±	0.26	0.03	±	0.17	0.990
Alb (g/dL)																
	Baseline	4.70	±	0.19	4.57	±	0.22	0.125		4.36	±	0.21	4.51	±	0.20	0.148
Δ4wk		0.02	±	0.19	0.06	±	0.18	0.541		0.15	±	0.19	0.01	±	0.21	0.186
Δ8wk		0.02	±	0.22	0.09	±	0.16	0.353		0.18	±	0.26	0.00	±	0.14	0.120
Δ12wk		-0.05	±	0.14	0.13	±	0.14	0.003	#	0.12	±	0.22	0.07	±	0.15	0.621
A/G ratio																
	Baseline	1.77	±	0.18	1.74	±	0.22	0.684		1.66	±	0.21	1.56	±	0.24	0.400
Δ4wk		-0.04	±	0.12	-0.08	±	0.13	0.523		-0.08	±	0.15	-0.02	±	0.08	0.340
Δ8wk		0.01	±	0.10	-0.02	±	0.13	0.560		0.03	±	0.09	-0.02	±	0.06	0.280
Δ12wk		0.04	±	0.10	0.02	±	0.10	0.626		0.02	±	0.09	0.00	±	0.09	0.668
TNF-α (pg/mL)																
	Baseline	2.56	±	0.55	2.65	±	0.86	0.757		2.31	±	0.51	2.54	±	0.57	0.406
Δ12wk		0.38	±	0.36	0.54	±	0.57	0.424		0.63	±	0.39	0.46	±	0.23	0.343
IL-1β (pg/mL)																
	Baseline	0.41	±	0.17	0.56	±	0.38	0.184		0.46	±	0.15	0.53	±	0.21	0.436
Δ12wk		-0.28	±	0.07	-0.28	±	0.15	0.995		-0.27	±	0.04	-0.21	±	0.11	0.193
IL-6 (pg/mL)																
	Baseline	1.32	±	0.77	1.82	±	1.19	0.235		1.71	±	0.84	2.22	±	0.67	0.194

$\Delta$ 12wk	-0.37	$\pm$	0.28	-0.67	$\pm$	0.63	0.108	-0.59	$\pm$	0.54	-0.75	$\pm$	0.25	0.472
<b>Body weight (kg)</b>														
Baseline	73.97	$\pm$	6.95	73.19	$\pm$	6.12	0.761	60.83	$\pm$	4.59	62.06	$\pm$	5.19	0.614
$\Delta$ 4wk	0.87	$\pm$	1.82	0.14	$\pm$	0.85	0.238	0.27	$\pm$	0.80	0.27	$\pm$	0.89	0.997
$\Delta$ 8wk	0.34	$\pm$	1.33	-0.29	$\pm$	1.17	0.210	0.21	$\pm$	1.01	0.10	$\pm$	0.77	0.811
$\Delta$ 12wk	0.34	$\pm$	1.22	-0.92	$\pm$	1.18	0.013	#	$\pm$	1.60	-0.21	$\pm$	0.79	0.276
<b>BMI (kg/m<sup>2</sup>)</b>														
Baseline	25.39	$\pm$	1.91	25.65	$\pm$	2.17	0.752	24.95	$\pm$	1.36	25.61	$\pm$	1.95	0.419
$\Delta$ 4wk	0.31	$\pm$	0.63	0.05	$\pm$	0.31	0.229	0.11	$\pm$	0.33	0.09	$\pm$	0.36	0.889
$\Delta$ 8wk	0.13	$\pm$	0.47	-0.10	$\pm$	0.42	0.203	0.10	$\pm$	0.39	0.04	$\pm$	0.34	0.759
$\Delta$ 12wk	0.12	$\pm$	0.43	-0.33	$\pm$	0.41	0.012	#	$\pm$	0.67	-0.10	$\pm$	0.32	0.260
<b>Body fat percentage (%)</b>														
Baseline	25.99	$\pm$	3.99	25.35	$\pm$	3.14	0.644	36.43	$\pm$	3.40	37.73	$\pm$	2.65	0.412
$\Delta$ 4wk	0.37	$\pm$	0.56	0.25	$\pm$	0.86	0.681	0.99	$\pm$	1.03	0.86	$\pm$	0.80	0.779
$\Delta$ 8wk	0.19	$\pm$	0.75	0.13	$\pm$	1.01	0.870	0.83	$\pm$	1.32	0.86	$\pm$	0.91	0.963
$\Delta$ 12wk	-0.25	$\pm$	0.60	-0.35	$\pm$	1.07	0.791	0.59	$\pm$	1.35	0.10	$\pm$	0.76	0.402
<b>Adiponectin (<math>\mu</math>g/mL)</b>														
Baseline	7.42	$\pm$	2.73	6.88	$\pm$	2.36	0.591	12.10	$\pm$	4.64	8.93	$\pm$	2.13	0.114
$\Delta$ 12wk	-0.24	$\pm$	0.73	-0.33	$\pm$	0.62	0.720	-1.11	$\pm$	1.13	0.51	$\pm$	0.87	0.006
#														
<b>Leptin (ng/mL)</b>														
Baseline	12.07	$\pm$	6.17	12.26	$\pm$	4.53	0.930	23.05	$\pm$	9.46	29.39	$\pm$	19.58	0.386
$\Delta$ 12wk	-1.48	$\pm$	3.68	-2.93	$\pm$	2.08	0.256	1.23	$\pm$	10.25	-3.14	$\pm$	4.61	0.311

Values are presented as the mean  $\pm$  SD.

\*P-values were determined by independent sample t-tests; †: P < 0.10, #: P < 0.05.

**Supplementary Table 6.** Effect of SM DNA on hepatic function markers obtained after subgroup analysis based on the lower ALT values (<51 U/L) participants

	Male						Female							
	Placebo (n = 11)			SM DNA (n = 16)			P-value*	Placebo (n = 10)			SM DNA (n = 7)			P-value*
ALT (U/L)	Baseline	32.38 ± 7.91	35.43 ± 8.36	0.407	30.56 ± 7.32	30.00 ± 12.60	0.920	Δ4wk	7.75 ± 22.70	7.29 ± 11.51	0.958	0.00 ± 6.73	4.57 ± 19.41	0.570
	Δ8wk	3.38 ± 9.16	2.14 ± 7.62	0.753	1.22 ± 9.23	-3.29 ± 8.52	0.329	Δ12wk	0.13 ± 5.41	2.79 ± 11.30	0.466	-2.67 ± 6.26	0.86 ± 4.98	0.231
AST (U/L)	Baseline	25.63 ± 4.50	27.07 ± 4.45	0.478	24.56 ± 5.43	25.86 ± 9.14	0.746	Δ4wk	3.38 ± 7.73	3.57 ± 6.72	0.953	-2.33 ± 2.96	1.14 ± 12.27	0.489
	Δ8wk	1.13 ± 5.03	0.29 ± 3.52	0.684	-1.33 ± 4.27	-2.29 ± 5.82	0.723	Δ12wk	-0.13 ± 2.70	0.57 ± 4.97	0.675	-3.22 ± 3.15	-1.86 ± 2.79	0.375
γ-GTP (U/L)	Baseline	40.63 ± 11.16	64.14 ± 31.85	0.022 #	53.00 ± 44.44	51.00 ± 25.94	0.912	Δ4wk	23.63 ± 63.47	11.50 ± 16.69	0.611	-0.78 ± 13.75	19.86 ± 49.13	0.318
	Δ8wk	10.88 ± 30.96	2.36 ± 11.46	0.475	-3.89 ± 14.97	-2.86 ± 14.84	0.893	Δ12wk	1.63 ± 8.57	0.21 ± 20.24	0.822	-14.78 ± 24.83	-0.86 ± 15.77	0.195
ALP (U/L)	Baseline	220.25 ± 44.88	202.07 ± 45.58	0.378	226.78 ± 69.34	267.29 ± 98.42	0.376	Δ4wk	11.88 ± 25.65	8.50 ± 19.24	0.752	4.67 ± 14.66	17.29 ± 67.36	0.643
	Δ8wk	1.75 ± 18.06	9.21 ± 24.88	0.428	7.00 ± 14.62	-7.57 ± 18.85	0.119	Δ12wk	-0.25 ± 20.82	9.36 ± 23.50	0.335	-1.67 ± 21.62	-3.71 ± 23.39	0.860
LDH (U/L)	Baseline	181.75 ± 30.20	178.07 ± 20.25	0.765	180.89 ± 36.47	182.43 ± 20.16	0.916	Δ4wk	10.50 ± 15.41	8.86 ± 10.08	0.792	-0.11 ± 10.22	4.00 ± 16.58	0.578
	Δ8wk	-1.75 ± 15.96	-1.86 ± 8.98	0.986	-1.89 ± 14.08	-4.14 ± 18.86	0.797	Δ12wk	1.13 ± 5.17	7.21 ± 13.05	0.139	4.56 ± 18.72	1.57 ± 9.93	0.689
ChE (U/L)	Baseline	416.38 ± 62.29	415.50 ± 66.56	0.976	376.11 ± 88.52	414.29 ± 44.76	0.283	Δ4wk	9.38 ± 25.31	12.64 ± 30.27	0.790	9.00 ± 41.55	1.57 ± 21.91	0.653
	Δ8wk	4.25 ± 21.46	7.79 ± 21.28	0.715	11.78 ± 30.03	7.14 ± 18.32	0.709	Δ12wk	0.50 ± 24.85	-0.29 ± 26.79	0.946	0.56 ± 28.43	7.86 ± 16.76	0.533
AST/ALT ratio	Baseline	0.82 ± 0.18	0.79 ± 0.16	0.667	0.83 ± 0.21	0.91 ± 0.21	0.476	Δ4wk	-0.02 ± 0.22	-0.05 ± 0.11	0.710	-0.09 ± 0.21	-0.08 ± 0.20	0.887

$\Delta 8\text{wk}$	-0.05	$\pm$	0.14	-0.03	$\pm$	0.09	0.735	-0.06	$\pm$	0.14	0.04	$\pm$	0.18	0.243
$\Delta 12\text{wk}$	0.01	$\pm$	0.07	-0.03	$\pm$	0.14	0.312	0.00	$\pm$	0.09	-0.11	$\pm$	0.15	0.137
<hr/>														
L/S ratio														
Baseline	1.12	$\pm$	0.21	0.92	$\pm$	0.27	0.071	1.13	$\pm$	0.21	1.20	$\pm$	0.22	0.559
$\Delta 12\text{wk}$	-0.02	$\pm$	0.12	0.11	$\pm$	0.16	0.044	#	0.00	$\pm$	0.07	0.00	$\pm$	0.05
														0.856

Values are presented as the mean  $\pm$  SD.

\*P-values were determined by independent sample *t*-tests; †:  $P < 0.10$ , #:  $P < 0.05$ .