

A unique profile of gut microbiota associated with metabolic syndrome: a remote island most afflicted by obesity in Japan

Tsugumi Uema¹, Jasmine F. Millman^{1*}, Shiki Okamoto^{1*}, Takehiro Nakamura¹, Kiyoto Yamashiro¹, Moriyuki Uehara¹, Ken-ichiro Honma¹, Minoru Miyazato², Asuka Ashikari³, Seiichi Saito³, Shiro Maeda^{4,5}, Minako Imamura^{4,5}, Hajime Ishida⁶, Masayuki Matsushita⁷, Koshi Nakamura⁸, Hiroaki Masuzaki^{1*}

¹ Division of Endocrinology, Diabetes and Metabolism, Hematology, Rheumatology, (Second Department of Internal Medicine), Graduate School of Medicine, University of the Ryukyus, Okinawa, Japan

² Department of Systems Physiology, Graduate School of Medicine, University of the Ryukyus, Okinawa, Japan

³ Department of Urology, Graduate School of Medicine, University of the Ryukyus, Okinawa, Japan

⁴ Department of Advanced Genomic and Laboratory Medicine, Graduate School of Medicine, University of the Ryukyus, Okinawa, Japan

⁵ Division of Clinical Laboratory and Blood Transfusion, University of the Ryukyus Hospital, Okinawa, Japan

⁶ Department of Human Biology and Anatomy, Graduate School of Medicine, University of the Ryukyus, Okinawa, Japan

⁷ Department of Molecular and Cellular Physiology, Graduate School of Medicine, University of the Ryukyus, Okinawa, Japan

⁸ Department of Public Health and Hygiene, Graduate School of Medicine, University
of the Ryukyus, Okinawa, Japan

*** Co-corresponding authors:**

Hiroaki Masuzaki

207 Uehara, Nishihara, Okinawa, 903-0215, Japan

hiroaki@med.u-ryukyu.ac.jp

Tel: +81-98-895-1146, Fax: +81-98-895-1415

ORCID: 0000-0002-2445-1047

Shiki Okamoto

207 Uehara, Nishihara, Okinawa, 903-0215, Japan

shiki@med.u-ryukyu.ac.jp

Tel: +81-98-895-1146, Fax: +81-98-895-1415

ORCID: 0000-0002-2802-0527

Jasmine Frances Millman

207 Uehara, Nishihara, Okinawa, 903-0215, Japan

Jasmine.f.millman@gmail.com

Tel: +81-98-895-1146, Fax: +81-98-895-1415

ORCID: 0000-0001-7049-3881

Supplementary figure legends

Figure S1 Composition of gut microbiota in NO and OB subjects showing relative abundance at the phylum level.

NO; non-obese (blue) (n = 81), OB; obese (red) (n = 59).

Figure S2 Composition of gut microbiota in MHO and MUO subjects showing relative abundance at the phylum level.

MHO; metabolically healthy obese (light blue) (n = 54) and MUO; metabolically unhealthy obese (orange) (n = 21).

Figure S3 Concentration of plasma short chain fatty acids (SCFAs) in NO and OB subjects. **a-d.** SCFA concentrations of acetate, propionate, butyrate, and valerate. **e.** Total concentration of plasma SCFAs. NO; non-obese (blue) (n = 81) and OB; obese (red) (n = 59). Data are shown as median and 25th – 75th percentiles. Data were analyzed using Mann-Whitney U test to compare groups.

Figure S4 Concentration of plasma SCFAs in MHO and MUO subjects. **a-d.** SCFA concentrations of acetate, propionate, butyrate, and valerate. **e.** Total concentration of plasma SCFAs. MHO; metabolically healthy obese (light blue) (n = 54) and MUO; metabolically unhealthy obese (orange) (n = 21). Data are shown as median and 25th – 75th percentiles. Data were analyzed using Mann-Whitney U test to compare groups.

Table S1 Measurement conditions of CE-TOFMS

Figure S1

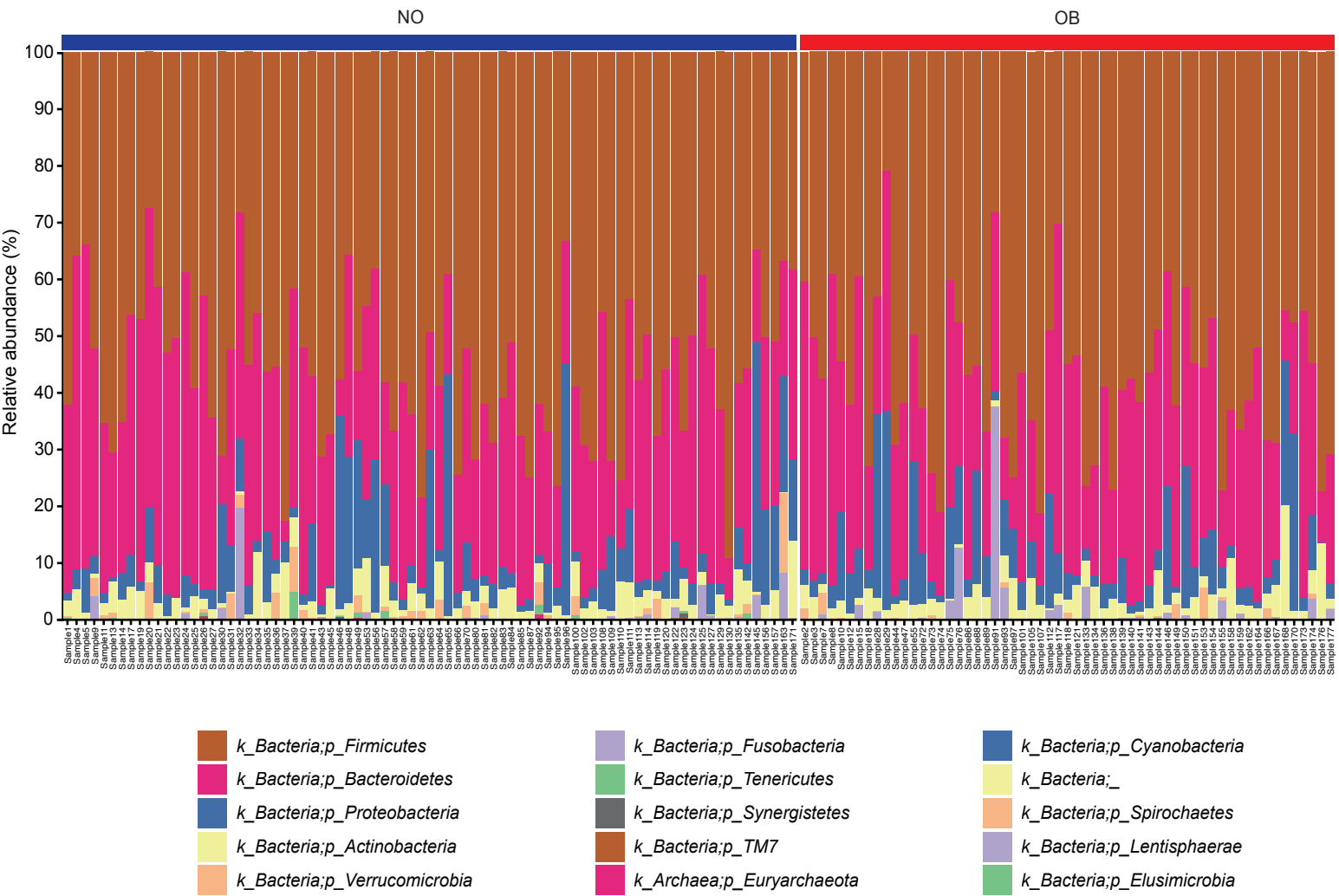


Figure S2

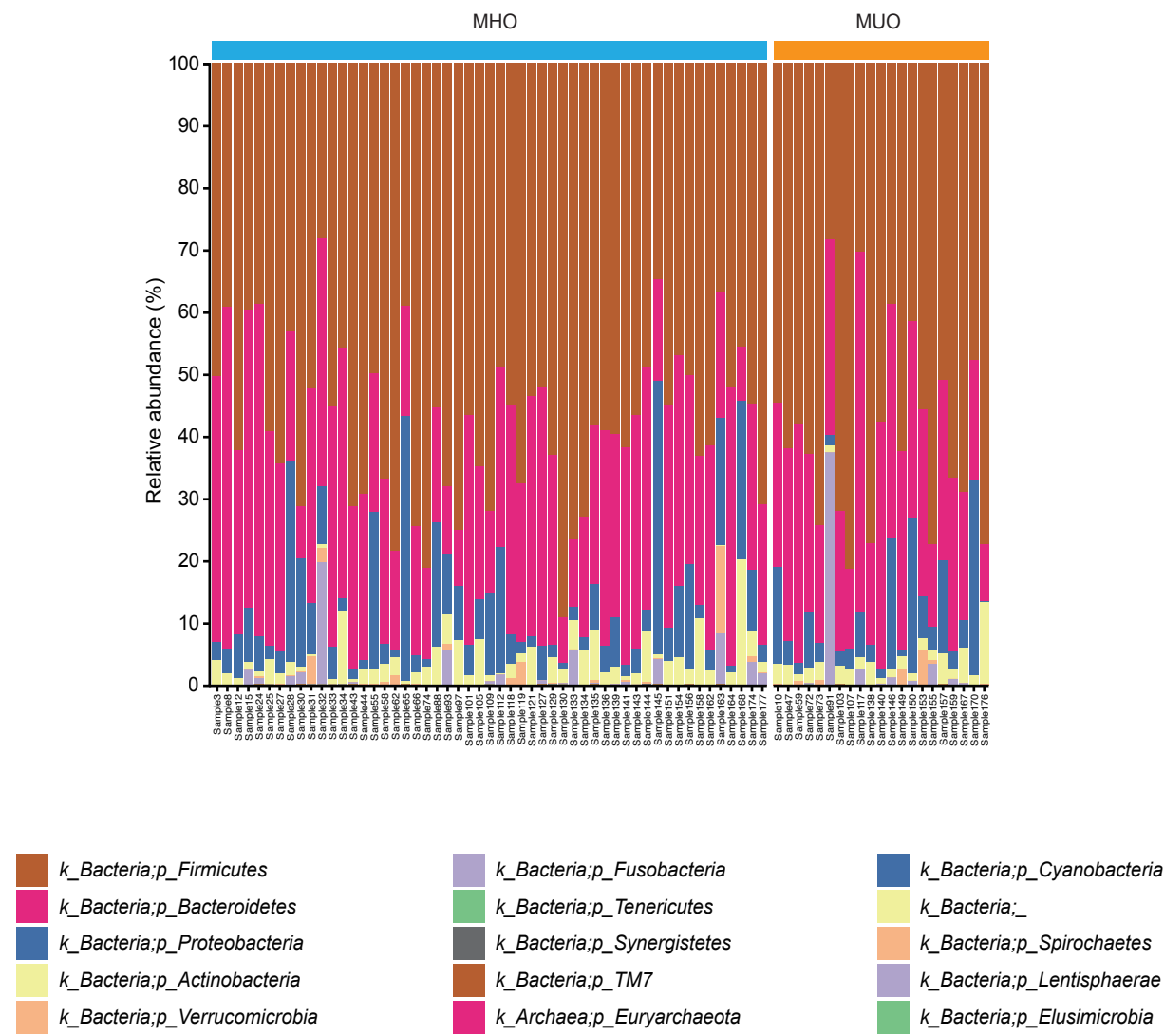


Figure S3

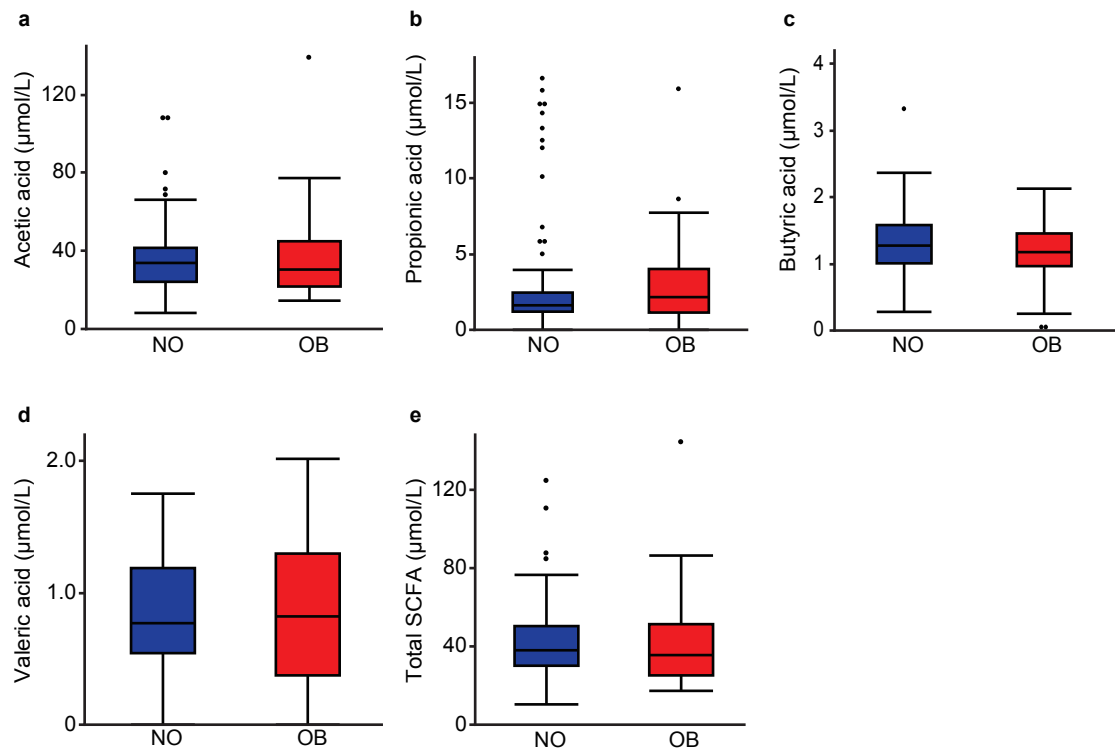


Figure S4

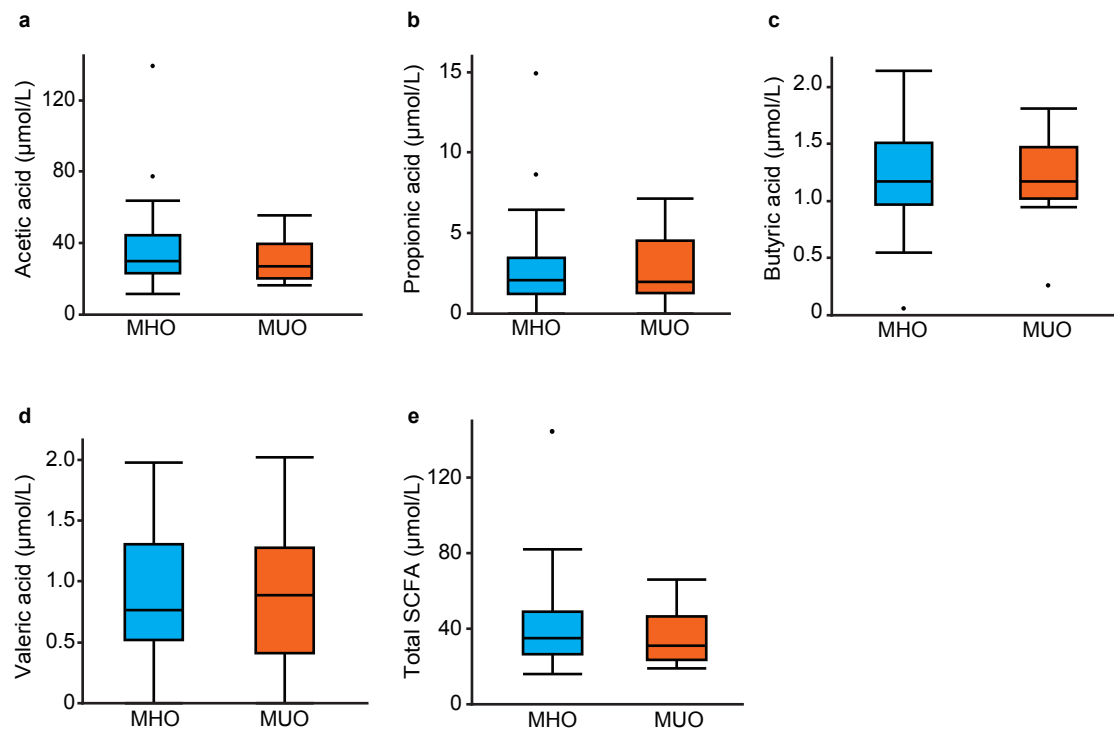


Table S1; Mesuring conditions of CE-TOFMS

	cation	anion
Run buffer	Cation Buffer Solution (p/n :H3301-1001)	Ation Buffer Solution (p/n :I3302-1023)
Rinse buffer	Cation Buffer Solution (p/n :H3301-1001)	Ation Buffer Solution (p/n :I3302-1023)
Sample injection	Pressure injection 50 mbar, 5 sec	Pressure injection 50 mbar, 10 sec
CE voltage	Positive, 30kV	Positive, 30kV
MS ionization	ESI Positive	ESI Negative
MS capillary voltage	4,000V	3,500V
MS scan range	m/z 50-1,000	m/z 50-1,000
Sheath liquid	HMT Sheath Liquid (p/n :H3301-1020)	HMT Sheath Liquid (p/n :H3301-1020)