



Public Access to Summary Statistics for Genome-wide Association Studies of Body Mass Index, Weight, and Height Among Healthy Japanese Individuals: The Japanese Consortium of Genetic Epidemiology Studies

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Recently, using the Mendelian randomization approach, we found that a genetically predicted high body mass index (BMI) was associated with an increased risk of colorectal cancer (CRC) in Japanese populations, strongly supporting the notion that high BMI influences the risk of CRC.¹ In the study, we utilized data for single nucleotide polymorphism (SNP)-BMI associations from genome-wide association studies (GWAS) among 36,303 participants in the Japanese Consortium of Genetic Epidemiology studies (J-CGE). We also conducted GWAS in the J-CGE for SNP-height and SNP-weight associations. The J-CGE consists of data analyzed among healthy participants from the Tohoku Medical Megabank Community-Based Cohort (TMM) Study, the Japan Public Health Centre-based Prospective (JPHC) Study, the Japan Multi-Institutional Collaborative Cohort (J-MICC) Study, and the Hospital-based Epidemiologic Research Program at Aichi Cancer Center (HERPACC) Study (Table 1). In each study, BMI, body weight, and height were assessed at the baseline through health check-ups (83%) or self-administered questionnaires. Protocols for data harmonization, quality control of GWAS data, imputation, and statistical analysis were established in advance. For BMI, height, and body weight, after inverse normal conversion and without conversion (ie, crude values), a GWAS was conducted separately for each study and each sex, with age (continuous variable), age squared (continuous variable), and 10 to 20 principal components as covariates and anthropometric indices as outcomes, using multiple regression analysis. A metaanalysis was then conducted using a fixed-effects model for SNPs with the imputation quality above a certain level (INFO score ≥ 0.5).² After quality control of the sample, the JPHC (3,525 men and 6,767 women), J-MICC (6,334 men and 7,736 women), TMM (3,123 men and 5,778 women), and HERPACC (1,502 men and 1,540 women) were combined to analyze 36,305 participants (Table 1). All the studies that contributed to J-CGE were approved by their respective institutional review boards.

The results of the BMI and height GWAS were largely consistent with those reported by a previous large-scale Japanese GWAS among individuals with diseases.^{3,4} We have made the genome-wide summary data for SNP-BMI associations publicly available; SNP-height, and SNP-weight associations will also be made available.⁵ After excluding SNPs with minor allele frequency of <0.01, the number of SNPs included in the summary data were approximately 7,000,000. To the best of our knowledge, this is the first example of GWAS summary statistics made publicly available for BMI, height, and weight from a Japanese consortium comprised of a general population of Japanese individuals. The genome-wide summary data for BMI, weight, and height include coefficients, standard errors, and *P*-values, which are provided in the text files. The GWAS data

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Table 1. Characteristics of the studies utilized for the analyses

Study	BMI, kg/m ²				Height, cm				Weight, kg			
	Men		Women		Men		Women		Men		Women	
	N(%)	Mean (SD)	N(%)	Mean (SD)	N(%)	Mean (SD)	N(%)	Mean (SD)	N(%)	Mean (SD)	N(%)	Mean (SD)
JPHC	3,524	23.6	6,766	23.8	3,525	162.1	6,766	150.6	3,525	62.3	6,767	54.1
	(34.2)	(2.9)	(65.8)	(3.2)	(34.3)	(6.3)	(65.7)	(5.5)	(34.2)	(9.2)	(65.8)	(8.0)
J-MICC	6,334	23.8	7,736	22.5	6,334	167.4	7,736	154.8	6,334	66.8	7,736	53.8
	(45.0)	(3.1)	(55.0)	(3.3)	(45.0)	(6.3)	(55.0)	(5.8)	(45.0)	(10.0)	(55.0)	(8.2)
TMM	3,123	24.1	5,778	23.2	3,123	165.1	5,778	153.1	3,123	65.9	5,778	54.4
	(35.1)	(3.2)	(64.9)	(3.7)	(35.1)	(6.5)	(64.9)	(5.9)	(35.1)	(10.2)	(64.9)	(8.9)
HERPACC	1,502	23.5	1,540	22.0	1,502	167.6	1,540	156.1	1,502	66.2	1,540	53.5
	(49.4)	(2.8)	(50.6)	(3.3)	(49.4)	(6.3)	(50.6)	(5.4)	(49.4)	(9.5)	(50.6)	(8.3)

BMI, body mass index; HERPACC, Hospital-based Epidemiologic Research Program at Aichi Cancer Center; J-MICC, Japan Multi-Institutional Collaborative Cohort Study; JPHC, Japan Public Health Centre-based Prospective Study; SD, standard deviation; TMM, Tohoku Medical Megabank Community-Based Cohort.

among healthy Japanese individuals, such as those from the J-CGE Study, would be ideal for etiologic research with BMI, weight, or height as exposures or outcomes. These publicly available data can be used to perform genetic epidemiologic studies, such as Mendelian randomization studies, among Asian populations.

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REFERENCES

- Suzuki S, Goto A, Nakatochi M, et al. Body mass index and colorectal cancer risk: a Mendelian randomization study. *Cancer Sci.* 2021; 112:1579–1588.
- Marchini J, Howie B. Genotype imputation for genome-wide association studies. *Nat Rev Genet*. 2010;11:499–511.
- Akiyama M, Okada Y, Kanai M, et al. Genome-wide association study identifies 112 new loci for body mass index in the Japanese population. *Nat Genet.* 2017;49:1458–1467.
- Akiyama M, Ishigaki K, Sakaue S, et al. Characterizing rare and lowfrequency height-associated variants in the Japanese population. *Nat Commun.* 2019;10:4393.
- Download page for the Japanese Consortium of Genetic Epidemiology studies (J-CGE). https://epi.ncc.go.jp/cgi-bin/cms/public/index. cgi/jcge/download/index.