

Total Energy Intake May Be More Associated with Glycemic Control Compared to Each Proportion of Macronutrients in the Korean Diabetic Population

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Background: Major macronutrients for energy intake vary among countries and cultures. Carbohydrates, including rice, are the major component of daily energy intake in Korea. The aim of this study was to examine the association of daily energy intake or each proportion of macronutrients, especially carbohydrates, with glycemic control in diabetic Koreans.

Methods: A total of 334 individuals with diabetes (175 men, age 57.4 ± 0.8 years; 159 women, age 60.9 ± 0.9 years) who participated in the 2005 Korean National Health and Nutrition Examination Survey were examined. Glycemic control was categorized based on concentration of glycated hemoglobin (HbA1c; HbA1c $\leq 6.5\%$; 6.6% to 8.0%; $\geq 8.1\%$). Dietary intake was assessed by using a 24-recall item questionnaire.

Results: High total energy intake was associated with poor glycemic control (HbA1c $\leq 6.5\%$, $1,824 \pm 75$ kcal; 6.6% to 8.0%, $1,990 \pm 57$ kcal; $\geq 8.1\%$, $2,144 \pm 73$ kcal; *P* value for trend = 0.002). Each proportion of protein, fat, or carbohydrate was not associated with glycemic control. Even after adjusting for several parameters, the association of daily energy intake with glycemic control still persisted.

Conclusion: Total energy intake may be more closely related to glycemic control than each proportion of macronutrients in Korean diabetics.

Keywords: Diabetes mellitus; Glycated hemoglobin; Korea; Macronutrient intake

INTRODUCTION

The prevalence of type 2 diabetes mellitus (DM) has increased worldwide, and this trend is expected to continue. The prevalence of diabetes in Korea has explosively increased 6- to 7-fold from 1.5% to almost 10% during the past 30 years [1-4]. Furthermore, the numbers of diabetic patients suffering from various diabetic complications and diabetes-related mortality have rapidly increased over the past few decades [1-3]. Although glycemic control is an important determinant in the development of diabetic complications [5-7], two previous studies reported that only 43.5% and 22.9% of Korean diabetic patients

had HbA1c levels $< 7.0\%$ and $< 6.5\%$, respectively [8,9].

Macronutrients including carbohydrate, protein, and fat have varying effects on blood glucose level. Considering that carbohydrate is the single major dietary component affecting the postprandial glucose level compared to protein and fat, carbohydrate intake is much more important in advanced diabetes, which generally shows marked postprandial hyperglycemia [10]. In far-east Asia, rice is the major staple food and the major source of daily energy intake, which is quite different compared to Western countries. An association between macronutrient intake and glycemic control in Western countries has been conclusively established [11-23]. Total energy

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Received: Oct. 20, 2011; Accepted: Feb. 16, 2012

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intake and total fat intake are closely related with glycemic control [11,12,22,23]. However, the association between carbohydrate intake and glycemic control is unclear [13-21]. While most of the latter studies reported that low carbohydrate intake is related to good glycemic control and weight loss [13-18], severe restriction of carbohydrate intake has been linked to poor glycemic control, because it results in an increase of other dietary components such as fat [19-21].

The aim of the present study was to evaluate the association between daily energy intake or each proportion of macronutrient, especially carbohydrate, and glycemic control in diabetic Koreans.

METHODS

Study population

This study was based on the data obtained from the third Korea National Health and Nutrition Examination Survey (KNHANES III) among non-institutionalized civilians in the Republic of Korea, which was conducted by the Korean Ministry of Health and Welfare in 2005. This survey was a nationwide representative study using a stratified and multistage probability sampling design for the selection of household units. The survey consisted of a health interview, health behavior, health examination, and nutrition components. A total of 34,145 individuals from these sampling frames were included in the health interview survey. Among them, 334 persons with diabetes (fasting glucose concentration ≥ 126 mg/dL, taking insulin, or oral anti-diabetic medication) were identified as participants, via a laboratory test and nutritional survey data. Glycemic control was categorized based on the concentration of glycated hemoglobin (HbA1c $\leq 6.5\%$; 6.6% to 8.0%; $\geq 8.1\%$).

Health examination survey and laboratory test

Trained interviewers visited each participant's dwelling and administered a standardized questionnaire on smoking, alcohol consumption, regular exercise, and diabetes treatments (e.g., taking insulin or oral hypoglycemic agents). A dietary recall method was used to collect data on food items consumed by participants during the past 24 hours. Height and weight were obtained using standardized techniques and equipment. Height was measured to the nearest 0.1 cm using a portable stadiometer (Seriter, Bismarck, ND, USA). Weight was measured to the nearest 0.1 kg using a calibrated balance-beam scale (Giant-150N; Hana, Seoul, Korea). Body mass index

(BMI) was calculated by dividing weight by height squared (kg/m^2). Waist circumference (WC) was measured on standing participants with a soft tape midway between the lowest rib and the iliac crest. Blood samples were collected on the morning after fasting for at least 8 hours. Fasting plasma glucose (FPG) was measured in a central and certified laboratory; an Advia1650 (Siemens, Tarrytown, NY, USA) was used. HbA1c was measured using a Bio-Rad Varian II (Bio-Rad, Hercules, CA, USA) in all participants.

Statistical analysis

Baseline characteristics were summarized for men and women and presented as means \pm SEM. All statistical techniques were performed using the SPSS for personal computer (SPSS Inc., Chicago, IL, USA). ANCOVA was done to examine age- and sex-adjusted characteristics according to glycemic control. To compare the difference of glycemic control according to daily energy intake or carbohydrate % of energy after adjusting for age, sex, diabetes duration, alcohol intake, exercise, and modality of anti-diabetes treatment, ANCOVA with post hoc analysis (least significant difference [LSD] multiple comparison test) was used. To determine which parameter was associated with glycemic control, logistic regression analysis was done with variables including age, sex, diabetes duration, daily alcohol intake, regular exercise, modality of anti-diabetes treatment, daily energy intake, carbohydrate % of energy, protein % of energy, and fat % of energy. Significance was determined by two-tailed analysis where $P < 0.05$.

Ethics statement

This study was approved by the Institutional Review Board of the Ilsan Paik Hospital (IB-3-1106-018).

RESULTS

Table 1 displays data on age and age-adjusted characteristics of the study population. There were 175 diabetic men (mean age, 57.4 ± 0.8 years) and 159 diabetic women (mean age, 60.9 ± 0.9 years) included in the analysis. Mean HbA1c was $7.39 \pm 0.13\%$ in men and $7.70 \pm 0.13\%$ in women. BMI was $24.6 \pm 0.2 \text{ kg}/\text{m}^2$ in men and $25.1 \pm 0.3 \text{ kg}/\text{m}^2$ in women. DM duration was about 5 years. Total energy intake in men was $2,229 \pm 54$ kcal, which was higher than that in women ($1,724 \pm 56$ kcal) ($P < 0.001$).

Table 2 displays data on age, sex, and age- and sex-adjusted characteristics according to glycemic control. Poor glycemic

Table 1. Age and age-adjusted characteristics of the study population

Characteristic	Men (n=175)	Women (n=159)	P value
Age, yr	57.4±0.8	60.9±0.9	0.006
Fasting plasma glucose, mg/dL	152.8±3.9	149.7±4.1	0.578
HbA1c, %	7.39±0.13	7.70±0.13	0.091
Current smoking, %	43.7±3.0	7.3±3.2	<0.001
Alcohol intake, g/day	24.6±2.6	2.3±2.7	<0.001
Exercise x3/wk, %	45.6±3.7	37.2±3.9	0.125
Body mass index, kg/m ²	24.6±0.2	25.1±0.3	0.174
Waist circumference, cm	88.4±0.6	85.2±0.7	0.001
Diabetes duration, yr	5.05±0.50	5.40±0.53	0.628
OHA or insulin treatment, %	51.1±3.6	58.9±3.8	0.142
Daily energy intake, kcal	2,229±54	1,724±56	<0.001
Protein % of energy	14.6±0.3	14.5±0.3	0.930
Fat % of energy	15.3±0.6	14.7±0.6	0.491
Carbohydrate % of energy	62.3±1.0	69.5±1.0	<0.001
Daily fiber intake, g/1,000 kcal	4.14±0.16	4.22±0.16	0.743

Data are presented as mean or %±SEM.

OHA, oral hypoglycemic agent.

control was associated with a longer duration of diabetes. High total energy intake was associated with poor glycemic control (HbA1c ≤6.5%, 1,824±75 kcal; HbA1c 6.6% to 8.0%, 1,990±57 kcal; HbA1c ≥8.1%, 2,144±73 kcal, *P* for trend is 0.002). After adjusting for age, sex, alcohol intake, exercise, current smoking, diabetes duration, and oral anti-diabetic medication or insulin treatment, this relationship still persisted. Carbohydrate %, protein %, or fat % of daily energy intake was not associated with glycemic control. In comparison of HbA1c among three equally divided participant groups according to daily energy intake, HbA1c in the highest tertile group was significantly higher than that in the lowest tertile group after adjusting for age, sex, diabetes duration, alcohol intake, exercise, and modality of anti-diabetes treatment (Table 3). There was no difference of HbA1c among three equally divided participant groups according to carbohydrate % of energy intake.

Logistic regression analysis for HbA1c >6.5% showed that diabetes duration and daily energy intake were associated with glycemic control, respectively, after adjusting for the above mentioned variables. Other parameters, including each proportion of macronutrients, were not significant (Table 4). After dividing the participants into men and women, daily energy intake was associated with glycemic control independently

Table 2. Age, sex, and age- and sex-adjusted characteristics according to glycemic control

Characteristic	HbA1c, %			P value for trend
	≤6.5 (n=88)	6.6-8.0 (n=153)	≥8.1 (n=93)	
Age, yr	58.1±1.2	60.2±0.9	58.2±1.2	0.289
Men, %	60.2	52.9	44.1	0.030
Fasting plasma glucose, mg/dL	120.6±4.4	140.8±3.4	197.8±4.3	<0.001
HbA1c, %	5.97±0.01	7.18±0.08	9.60±0.10	<0.001
Current smoking, %	28.8±4.3	22.2±3.2	30.9±4.2	0.211
Alcohol intake, g/day	18.5±3.7	9.7±2.8	16.8±3.6	0.102
Exercise x3/wk, %	35.6±5.3	42.5±4.0	45.9±5.1	0.362
Body mass index, kg/m ²	25.1±0.3	24.9±0.3	24.7±0.3	0.676
Waist circumference, cm	86.9±0.9	87.0±0.7	86.7±0.9	0.967
Diabetes duration, yr	3.89±0.70	4.98±0.53	6.86±0.68	0.002
OHA or insulin treatment, %	50.6±5.1	54.3±3.9	59.6±5.0	0.451
Protein % of energy	14.3±0.4	14.8±0.3	14.4±0.4	0.668
Fat % of energy	14.7±0.8	15.1±0.6	15.2±0.8	0.885
Carbohydrate % of energy	64.3±1.4	66.9±1.0	65.2±1.3	0.271
Daily fiber intake, g/1,000 kcal	4.20±0.22	4.28±0.17	3.99±0.21	0.553
Daily energy intake, kcal	1,824±75	1,990±57	2,144±73	0.002
Daily energy intake, kcal ^a	1,817±73	2,003±55	2,129±71	0.01

Data are presented as mean or %±SEM.

OHA, oral hypoglycemic agent.

^aAfter adjusting for age, sex, alcohol intake, exercise, current smoking, diabetes duration, and OHA or insulin treatment.

in men (odds ratio [OR], 1.001 [1.000 to 1.001]; *P*=0.013, per unit of daily energy intake). However, in women there was no significant association (OR, 1.000 [0.999 to 1.001]; *P*=0.641). In participants with no anti-diabetes medication, daily energy intake was associated with glycemic control independently (OR, 1.001 [1.000 to 1.001]; *P*=0.032, per unit of daily energy intake). However, there was no significant association in those receiving oral anti-diabetes or insulin treatment (OR, 1.000 [1.000 to 1.001]; *P*=0.176). In the case of carbohydrate % of

Table 3. HbA1c according to daily energy intake or carbohydrate % of energy after adjusting for age, sex, diabetes duration, alcohol intake, exercise, and modality of anti-diabetes treatment

	Daily energy intake						P value for trend
	Tertile 1,	Tertile 2,	Tertile 3,	P value between two groups with post hoc analysis			
	≤1,613 kcal	1,614-2,208 kcal	≥2,209 kcal	1 vs. 2	2 vs. 3	1 vs. 3	
HbA1c, %	7.33±0.16	7.42±0.16	7.85±0.16	0.687	0.062	0.032	0.072
	Carbohydrate % of energy						P value for trend
	Tertile 1,	Tertile 2,	Tertile 3,	P value between two groups with post hoc analysis			
	<63%	63-73%	>73%	1 vs. 2	2 vs. 3	1 vs. 3	
HbA1c, %	7.40±0.17	7.69±0.16	7.51±0.16	0.248	0.418	0.667	0.483

Data are presented as % ± SEM.

Table 4. Logistic regression analysis for HbA1c > 6.5%

	OR (95% CI)	P value
Age per unit	1.005 (0.982-1.029)	0.677
Women	1.622 (0.921-2.856)	0.094
Diabetes duration per unit	1.057 (1.001-1.117)	0.047
Daily alcohol intake per unit	1.005 (0.982-1.029)	0.647
Regular exercise (≥x3/wk)	1.377 (0.813-2.333)	0.234
OHA or insulin treatment	0.811 (0.426-1.542)	0.523
Daily energy intake per unit	1.000 (1.000-1.001)	0.029
Carbohydrate % of energy per unit	1.045 (0.970-1.126)	0.244
Protein % of energy per unit	1.045 (0.951-1.148)	0.357
Fat % of energy per unit	1.033 (0.954-1.118)	0.424
R ²	0.093	

OR, odds ratio; CI, confidence interval; OHA, oral hypoglycemic agent.

energy, there was no significant association with glycemic control in each subgroup (data not shown).

DISCUSSION

The present study examined the relationship between glycemic control and macronutrient intake, specifically total daily energy vs. proportion of carbohydrate in daily energy, in a diabetic Korean population. The data suggests that total energy intake is more closely related to glycemic control than the proportion of macronutrients in Korean individuals.

The staple food of most countries in Asia is carbohydrate. According to KNHANES III [24], the daily total energy intake was 2,016.3 kcal, and the percent of carbohydrate intake of total energy intake was 64.5%. Furthermore, the major component of energy intake was raw rice (38%). Koreans consume

more carbohydrate than Americans. In the United States, NHANES 1999-2000 data indicated that carbohydrate intake was 51.9% of total energy intake and fat intake was 32.7% of total energy intake [25].

Medical nutrition therapy (MNT) is the most important aspect of diabetes treatment including preventing diabetes, managing existing diabetes, and preventing or delaying the rate of development of diabetes complications [26,27]. Optimizing energy intake and macronutrient composition are important in MNT.

Total energy is an important contributor to glycemic control. Several studies focusing on the effects of reducing energy intake have shown improved glycemic control, both in energy restriction and energy balance [11,12]. Among individuals with type 2 DM, significant positive correlations were found between glycemic control and intakes of energy. Modest weight reductions (as little as 5%) confer long-term benefits in patients with type 2 DM due to reductions in FPG [28]. Whereas, it is well-known that caloric restriction is essential for the achievement of good glycemic control, mainly through weight loss, the optimal dietary macronutrient composition for patients with type 2 DM remains controversial.

A recent meta-analysis compared the effects of a low-fat, high-carbohydrate diet and a high-fat, low-carbohydrate diet on glycemic control [21]. There was no significant difference in the reduction in HbA1c. However, replacing fat with carbohydrate significantly elevated postprandial glucose levels when total energy intake was consistent.

In a meta-analysis of 13 trials on the influence of restricted carbohydrate diets in individuals with type 2 DM [13], a greater reduction in hyperglycemia was found in the low carbohydrate diet compared to high carbohydrate diets. A 10% increase

in caloric intake from carbohydrates was associated with a $3.2 \pm 1.2\%$ increase in glucose levels ($P=0.047$). However, the meta-analysis included a small number of studies and did not evaluate long-term risks or benefits.

In Korea, according to KNHANES III [29], when percent energy intake from carbohydrates exceeded 70%, the risk of impaired glucose tolerance and diabetes increased. According to another study conducted in elderly diabetic Korean individuals [30], a lower intake of protein or fat was related to higher HbA1c levels, while there was no association between carbohydrate intake and HbA1c levels. In our study, lower total energy intake was related to good glycemic control, whereas the proportion of carbohydrate, protein, and fat were not.

We hypothesized that the proportion of carbohydrate, which is major energy intake in Korea, could be associated with glycemic control. However, this association was not presently evident. Although our study was done using nationally representative data, the clinical characteristics of the participants should be defined before interpretation of the data. Many participants in the present study exhibited relatively good glycemic control (mean HbA1c, $7.39 \pm 0.13\%$ in men and $7.70 \pm 0.13\%$ in women) and a shorter duration of diabetes (5.05 ± 0.50 years in men and 5.40 ± 0.53 years in women), which might attenuate the association between the proportion of carbohydrate and glycemic control. Therefore, we cannot rule out the association of carbohydrate percentage with glycemic control in more advanced and more insulin-deficient diabetes patients with prominent postprandial hyperglycemia.

There were several limitations in the current study. A single day diet is a poor descriptor of a person's usual intake, because of intra-individual variability. Secondly, because the results were obtained from cross-sectional data, we are not able to draw conclusions about the temporal relationship between macronutrient intake and glycemic control. Another limitation might be the lack of information about diabetes treatment, including compliance to treatment and the type of medication. Although the strength of our study was its nationally representative data, there could be some limitation for generalizing these results to the general population in Korea, because we did not account for primary sampling units, stratification, and sampling weights for KNHANES. Moreover, the result could not extend to all diabetics because of the different effects of macronutrients based on age, sex, the duration of diabetes, status of glycemic control, and compliance to treatment. In this study, the association of daily energy intake with glycemic

control was observed only in men or participants with no anti-diabetes medication in subgroup analyses. There was no significant association in women or participants with current oral anti-diabetes or insulin treatment. Different health behavior in men compared to that in women or many other related factors for glycemic control, except for diet in the treatment group compared to the no treatment group, potentially explain these finding. However, we could not present a clear explanation due to limitations of this study, including sample size. Further clinical trials are needed to more conclusively establish the relationship between glycemic control and macronutrients in a subgroup of diabetics for optimizing MNT.

In conclusion, the data suggested that high total energy intake may be more closely related to poor glycemic control in diabetic Koreans, but the proportion of carbohydrate for total energy intake may not. Clinical trials are needed to test the association between carbohydrate and glycemic control in populations with longer diabetic duration and poor glycemic control. Such investigations may be useful in nutritional therapy for diabetic Koreans.

CONFLICTS OF INTEREST

No potential conflict of interest relevant to this article was reported.

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