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## Dietary carbohydrate intake and high sensitivity C reactive protein in at-risk women and men

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### Abstract

**Background—** The quality and quantity of dietary carbohydrate intake, measured as dietary glycemic load (GL), is associated with a number of cardiovascular disease (CVD) risk factors and, in healthy young women, is related to increased high sensitivity C-reactive protein (hsCRP) concentrations. Our objective was to determine if GL is related to hsCRP and other measures of CVD risk in a population of sedentary, overweight, dyslipidemic middle-aged women and men enrolled in an exercise intervention trial (STRRIDE).

**Methods—** This was a cross-sectional evaluation of the relationships between measures of dietary carbohydrate intake, calculated from food frequency questionnaire data, and CVD risk factors, including plasma hsCRP, measured in 171 subjects.

**Results—** After adjusting for energy intake, GL and other measures of carbohydrate intake were not independently related to hsCRP ( $P > 0.05$  for all). In analyses performed separately for each gender, only the quantity of carbohydrate intake was independently related to hsCRP ( $R^2 = 0.28$ ;  $P < 0.04$ ), and this relationship was present for women but not for men. The strongest relationship identified between GL and any CVD risk factor was for cardiorespiratory fitness ( $R^2 = 0.12$ ;  $P < 0.02$ ); an elevated GL was associated with a lower level of fitness in all subjects, and this relationship persisted even when the findings were adjusted for energy intake and gender ( $R^2 = 0.48$ ;  $P < 0.03$ ).

**Conclusions—** In middle-aged, sedentary, overweight to mildly obese, dyslipidemic individuals, consuming a diet with a low GL is associated with better cardiorespiratory fitness. Our findings suggest that the current literature relating carbohydrate intake and hsCRP should be viewed with skepticism, especially in the extension to at-risk populations that include men.

### INTRODUCTION

A number of recent reports have shown that elevated post-prandial blood glucose concentrations lead to increased metabolic and cardiovascular risk [reviewed in <sup>1</sup>]. Since diet is a major determinant of post-prandial glucose response, these findings have re-focused

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attention on a well-known method for characterizing the impact of dietary carbohydrate on blood glucose response—the dietary glycemic load (GL). GL describes the quality, characterized by the glycemic index (GI), as well as quantity of carbohydrates consumed in the diet. The terms GI and GL are often used interchangeably, but they are not synonymous<sup>2</sup>. The GI of a food reflects the post-prandial glucose response to a single serving of that food; the GI of a diet reflects the average post-prandial glucose effect of single servings of all foods in a diet. GL takes into account the both the type and amount of carbohydrate consumed in a diet to indicate the overall predicted impact of quantity and quality of the dietary carbohydrate intake on the post-prandial glucose response.

Dietary GL has been associated with coronary heart disease<sup>3</sup> and a number of cardiovascular risk factors, including risk of type 2 diabetes mellitus<sup>4, 5</sup>, reduced HDL-cholesterol<sup>6–8</sup>, and elevated triacylglycerol concentrations<sup>8</sup>. These associations suggest that diets producing a high post-prandial glucose load may lead to detrimental metabolic alterations in lipid profiles and insulin sensitivity. Moreover, a report linking GL with serum concentrations of high sensitivity C-reactive protein (hsCRP) in healthy women<sup>9</sup> adds strength to the accumulating evidence that determinants of insulin action and inflammatory pathways are connected<sup>10</sup>. As such, interventions comparing weight loss achieved by using a low GL diet versus a low fat diet<sup>11</sup> or high GL diet<sup>12</sup> suggest that a low GL diet imposes superior metabolic cardiovascular changes, including reductions in hsCRP concentrations. It is particularly notable that these interventions have been performed primarily in young, healthy, overweight, females (over 75%).

An acute phase reactant and independent predictor of cardiovascular disease risk, hsCRP shows promise as an indicator of metabolic responses to environmental influences such as dietary intake patterns. Since studies to date have included only healthy subjects and with a predominance of women, the primary objective of this work was to assess the relationship between dietary GL and serum hsCRP concentrations in a population of women and men at-risk for cardiovascular disease. A secondary objective was to assess the relationship between GL and other metabolic cardiovascular risk factors in this population.

## METHODS

### Study Design

This was a cross-sectional evaluation of subjects enrolled in the first Studies of Targeted Risk Reduction In Defined Exercise (STRRIDE I, ClinicalTrials.gov Identifier: NCT00200993), which examined individuals at-risk for cardiovascular disease living in Durham and Greenville, NC, from 1999–2002. Relevant institutional review boards approved the research protocol, and informed consent was obtained for each subject. The data presented here were all collected at baseline, i.e., prior to treatment randomization.

### Subjects

Inclusion criteria were inactivity (not participating in regular exercise), overweight to mild obesity (body mass index [BMI] 25–35 kg/m<sup>2</sup>), dyslipidemia (LDL-cholesterol of 130–190 mg/dl *or* HDL-cholesterol <40 mg/dl for men and <45 mg/dl for women), post-menopausal status for women, and age between 40 and 69. Exclusion criteria included the use of medications known to affect carbohydrate or lipid metabolism, participation in a dietary regimen designed to induce weight loss, presence of diabetes mellitus (fasting glucose >140 mg/dl), hypertension (BP >160/90 mmHg), known cardiovascular disease, tobacco use, and/or a musculoskeletal condition that would prohibit exercise training.

## Metabolic and Cardiovascular Risk Factor Assessments

We measured anthropomorphic characteristics as previously described<sup>13</sup>, and visceral adiposity and subcutaneous adiposity with a single slice abdominal CT scan at L4<sup>14, 15</sup>. Cardiorespiratory fitness was assessed with exercise treadmill testing and measured in maximal oxygen consumption (ml O<sub>2</sub>\*min/kg), as previously described<sup>15</sup>.

Fasting plasma was collected from each participant and stored at -80° Celsius. Lipoprotein analyses were performed by LipoScience (Cary, NC) using nuclear magnetic resonance spectroscopy<sup>16</sup>. Fasting insulin (Access Immunoassay System, Beckman Coulter, Fullerton, CA) and fasting glucose (YSI model 2300 Stat Plus, Yellow Springs Instruments, Yellow Springs, OH) were measured as previously described<sup>17</sup>.

## High sensitivity-CRP Measurement

We measured high sensitivity CRP concentrations with an ELISA (UBI) as previously described<sup>18</sup>. The manufacturer reported inter- and intra-assay variations of 6.6 to 10.8% and 3.9 to 9.2%, respectively. Each sample was assayed in duplicate with coefficients of variation less than 15%.

## Assessment of dietary carbohydrate

Reported carbohydrate intake and dietary GL and GI were assessed using a semi-quantitative food frequency questionnaire, the Block 1998 Revision of the Health Habits and History Questionnaire or HHHQ, (Block Dietary Data Systems, Berkeley, California). HHHQs were completed by STRRIDE participants who indicated their consumption frequency and portion size for over 100 food items. Completed forms were analyzed by Block Dietary Data Systems for nutrient content. This information along with GI values for foods assigned by the Nutritional Epidemiology Core at the University of North Carolina ([http://www.cnrc.unc.edu/core\\_b.htm](http://www.cnrc.unc.edu/core_b.htm)) was used to calculate GL and GI according to the following formulae:

$$\text{Daily GL} = \sum_{\text{all foods}} [\text{Frequency} \times \text{Portion size} \times (\text{g carbohydrate} - \text{g fiber}) \times \text{food GI}]$$

and Daily GI = [Daily GL/(Total g carbohydrate - Total g fiber)] × 100. GI was expressed relative to bread (Bread GI = 100).

The methodology for the development of the HHHQ<sup>19</sup>, and its validation for assessing the foods from which the GL and GI were derived<sup>20</sup> have been previously reported. Data collection via a food frequency questionnaire such as the HHHQ is the preferred approach for GL/GI analysis in large studies where food by food calculation is impractical<sup>9</sup>.

## Data analysis

We evaluated descriptive statistics including distributions for each variable. Variables with highly skewed distributions, fasting insulin, triacylglycerol, and hsCRP, were logarithmically transformed prior to statistical significance testing. Standard correlations were used to confirm assignment of GI values and to evaluate relationships between GL and cardiovascular risk factors. Analogous to the previously recommended residuals method,<sup>21</sup> we used multivariable linear regression to account for the impact of HDL-cholesterol and energy intake on hsCRP and GL. For all models with more than a single predictor, two-way and three-way interaction terms were initially included in the full model. If the interaction term was not significant, it was removed, and the main effects model was refit. For all analyses, statistical significance was established at  $P < 0.05$ , and statistics were performed using SAS Version 8.2 Enterprise Guide (SAS, Cary, NC). Power calculations were performed using Power and Precision 2.0 (Biostat, Englewood, NJ).

## RESULTS

### Relationships between carbohydrate intake and hsCRP

Descriptive data are presented in Table I. Our primary analysis of interest was to determine whether dietary carbohydrate quantity or quality is related to the surrogate of cardiovascular risk, hsCRP. Of the 171 STRRIDE participants who had completed HHHQs, 144 had plasma samples available for measurement of hsCRP concentrations. Since hsCRP screening recommendations for cardiovascular disease suggest excluding and repeating hsCRP values greater than 10 mg/L<sup>22</sup>, we evaluated the relationship between hsCRP and carbohydrate quality and quantity for only those subjects (n=136) with hsCRP concentrations less than 10 mg/L.

First, we wished to determine if a relationship existed for GL and hsCRP concentrations. Given the significant inverse relationship between HDL-cholesterol concentrations and GL, and the fact that we have previously noted a positive relationship between HDL-cholesterol and hsCRP<sup>18</sup>, we hypothesized that HDL-cholesterol may be a confounder in the relationship between GL and hsCRP. Indeed, after adjusting for HDL-cholesterol concentrations, GL ( $P<0.04$ ) was independently related to hsCRP and explained 14% of the variance in hsCRP (see Table II). After adjustment for energy intake and HDL-cholesterol, GL ( $P<0.18$ ) was no longer an independent predictor of hsCRP concentrations, but together, GL, energy intake, and HDL-cholesterol accounted for 14% of the variance on hsCRP (see Table II).

Second, we wished to determine if dietary carbohydrate quality, measured as GI, was related to hsCRP concentrations. In adjusted (HDL-cholesterol as well as energy intake + HDL-cholesterol) analyses, GI was not independently related to hsCRP concentrations, and the amount of variance in hsCRP was less than that found with GL (see Table II). When either GL or total daily carbohydrate were added to the energy intake and HDL-cholesterol-adjusted model containing GI, there was an improvement in the explanation of variance in hsCRP from 13% to 15%.

Next, we wished to determine whether the carbohydrate quantity alone, daily carbohydrate intake (DC), was related to hsCRP concentrations. Similar to the relationships seen for GL, after adjustment for HDL-cholesterol, total DC was significantly and independently related to hsCRP concentrations. Together, total DC ( $P<0.03$ ) and HDL-cholesterol accounted for 14% of the variance in hsCRP in this subject population (see Table II). However, after adjustment for energy intake plus HDL-cholesterol, DC ( $P<0.14$ ) was no longer an independent predictor of hsCRP concentrations ( $R^2=0.15$ , model  $P<0.001$ ). Furthermore, adding GL or GI to the model containing DC provided no improvement in its predictive capacity ( $R^2=0.15$  for all, model  $P<0.005$  for all).

### Relationships between carbohydrate intake and hsCRP for men and women

Taking note of published reports of gender differences in susceptibility to a number of cardiovascular risk factors, a key part of our primary objective for this analysis was to determine if the relationship between hsCRP and GL was different in men versus women. As part of this analysis, we tested the interaction term gender\*GL; it was not found to be a significant independent predictor for hsCRP (unadjusted GL\*gender:  $P<0.29$ ; HDLc adjusted GL\*gender:  $P<0.21$ ; HDLc and EI adjusted GL\*gender:  $P<0.21$ ). However, based on the aforementioned *a priori* evidence of relationships between hsCRP and metabolic cardiovascular risks in women<sup>18</sup> but no reports of these relationships in men, we did evaluate the associations between carbohydrate intake and hsCRP separately by gender group.

In women, both HDL-cholesterol and GL were independent predictors of hsCRP and accounted for 25% of the variance ( $P<0.002$ ) (see Figure 1). In contrast, even after adjusting for HDL-

cholesterol concentrations, we found no significant relationship between GL and hsCRP in men ( $P<0.78$ ). GI was not an independent predictor of hsCRP in any analysis (see Table II). Interestingly, we found that total DC intake ( $P<0.01$ ) and HDL-cholesterol ( $P<0.01$ ) were each independently related to and together explained 26% of the variance in hsCRP in women (model  $P<0.001$ ), but neither variable explained any portion of the variance in hsCRP in men (model  $P<0.81$ ). When we controlled for both energy intake and HDL-cholesterol, total DC was independently ( $P<0.04$ ) related to hsCRP in women and accounted for 28% of the variance in hsCRP (model  $P<0.003$ ). These relationships were not apparent for men (model  $P<0.84$ ).

### Relationships for carbohydrate intake and cardiovascular risk factors

In addition to evaluating the relationship between GL and hsCRP, a secondary objective of this study was to evaluate the relationship between GL and other cardiovascular risk factors. Prior to adjusting for energy intake, we found significant relationships between dietary GL and several known metabolic cardiovascular risk factors. GL was correlated with weight ( $r=0.24$ ,  $P<0.001$ ), minimal waist circumference ( $r=0.22$ ,  $P<0.007$ ), cardiorespiratory fitness ( $r=0.18$ ,  $P<0.02$ ), visceral adiposity ( $r=0.22$ ,  $P<0.005$ ), and HDL-cholesterol ( $r=-0.22$ ,  $P<0.01$ ). Additionally, we noted trends toward statistical significance in the relationships between GL and BMI ( $P<0.10$ ) and triacylglycerol ( $P<0.06$ ). We found no relationship between GL and age, subcutaneous adiposity, percent body fat, fasting plasma glucose, fasting insulin, mean blood pressure, or LDL-cholesterol.

After adjustment for energy intake, GL ( $P<0.02$ ) was independently and inversely related to cardiorespiratory fitness ( $VO_{2max}$ , ml\*min/kg) ( $R^2=0.12$ , model  $P<0.0001$ ) (see Figure 2). The association between energy intake adjusted-GL and fitness was also independent of gender. Together, GL, energy intake, and gender account for 48% of the variance in cardiorespiratory fitness (model  $P<0.0001$ ,  $P_{GL}<0.03$ ). There was no significant gender\*GL interaction for the relationship between GL and cardiorespiratory fitness ( $p<0.32$ ). All other relationships between metabolic cardiovascular risk factors and GL were no longer apparent after adjusting for energy intake.

Fasting insulin was correlated with GI ( $P<0.05$ ), but the relationship between GI and fasting insulin was no longer apparent after adjustment for energy intake. We found no significant relationship between GI and any other risk factor (age, weight, BMI, minimal waist circumference, percent body fat, subcutaneous adiposity, visceral adiposity, cardiorespiratory fitness, HDL-cholesterol, LDL-cholesterol, triacylglycerol, fasting plasma glucose, fasting insulin, or mean blood pressure,  $P>0.05$  for all) with or without adjustment for energy intake.

**Power of the study**—We believed that a meaningful effect of GL would be one that would explain 10% of the variance in hsCRP or contribute 5% to a known amount of variance. For the primary analysis, which included GL, HDL-cholesterol, and EI, we had 91% power ( $\alpha=0.05$ ) to detect a significant relationship explaining 10% of the variance in hsCRP. If HDL-cholesterol and EI together accounted for 10% of the variance in hsCRP, we had 80% power ( $\alpha=0.05$ ) to detect that GL might explain an additional 5% of the variance in hsCRP. Additionally, we had 79% power ( $\alpha=0.05$ ) to detect a significant interaction between gender and GL that would improve the explanation of variance from 10% to 15%.

## DISCUSSION

Our findings indicate that in a subject demographic of sedentary, overweight to mildly obese, dyslipidemic middle-aged men and women, GL was related to a number of cardiovascular risk factors, including hsCRP, weight, waist circumference, visceral adiposity, and low HDL-cholesterol concentrations. However, after adjustment for energy intake, GL was inversely related to cardiorespiratory fitness, but other no relationships between GL and metabolic risk

factors were sustained. The only measure of carbohydrate intake serving as an independent predictor of hsCRP concentrations after adjustment for energy intake was DC, and this relationship appeared stronger for women than for men.

Previously, the relationship between GL and hsCRP was demonstrated in a cross-sectional investigation in a population with only women<sup>9</sup>. Similarly, prospective investigations demonstrating reductions in hsCRP with low GL diets have been performed in samples with predominantly women<sup>11, 12</sup>. Here, in a well-characterized mixed-gender population at-risk for cardiovascular disease, the relationship between GL and hsCRP concentrations did not persist when adjusted for energy intake. Most interestingly, the relationships presented for hsCRP and all measures of carbohydrate intake appeared much more robust for women than men. These results led us to speculate that the relationship between carbohydrate intake and hsCRP may be stronger or, perhaps, specific for women, which might explain the current lack of such relationships reported for mixed-gender or male only populations.

Spurred by this relative lack of published relationships between hsCRP and GL in men, which persists even at the conclusion of this analysis, we believe that our *a priori* hypothesis of a gender difference justified the stratified analyses presented here. However, it is worth noting that we did not detect a significant interaction between gender and GL for the relationship with hsCRP. This finding suggests that we were underpowered to evaluate a subtle gender difference for this relationship. As such, given that we do not have power to definitively argue that the relationship between GL and hsCRP occurs only for women, we believe the remarkable differences in the relationships for women and men merit attention and highlight the need for future observational and prospective studies to carefully evaluate gender differences in relationships between carbohydrate intake and cardiovascular risk.

Such a gender disparity in the relationship between GL and hsCRP may reflect a propensity for hsCRP to better represent metabolic risk in women than men. Consistent with this hypothesis, we have previously reported that metabolic cardiovascular risk factors could account for close to 50% of the variance in hsCRP in women but only 17% in men<sup>18</sup>. Similarly, Frost et al<sup>6</sup> reported that the inverse association between GI and HDL-cholesterol was stronger in women than in men.

Somewhat surprisingly, the relationship between GL and hsCRP was driven primarily by the total amount of carbohydrate consumed by the subjects. One might ask why an elevated consumption of carbohydrates would promote higher concentrations of an inflammatory molecule, such as hsCRP, when a high dietary GI would not. One possible explanation follows: CRP is produced in the liver, primarily in response to interleukin-6 (IL-6), but also to tumor necrosis factor- $\alpha$  (TNF- $\alpha$ ) and IL-1, each of which has been invoked in insulin resistance pathways<sup>10</sup>. A higher GL diet may induce higher rates of IL-6 secretion from adipocytes (and thus elevated hsCRP) primarily as a result the higher caloric (and carbohydrate) load.

In addition to potential mechanistic links between diet and hsCRP, as suggested above, relationships between GL and cardiovascular risk factors may simply indicate correlations between disease risk and health-related lifestyle choices. More importantly, it has not been determined whether lifestyle modifications, such as improving diet and exercise, can actually modulate hsCRP. A recent report demonstrated that a diet high in sucrose, a simple carbohydrate with a high GI, did not impose significant changes in hsCRP concentrations<sup>23</sup>. Such a finding would be consistent with observations by our group in this same population during the intervention portion of STRRIDE. While STRRIDE did not include a nutrition intervention, the exercise interventions produced marked improvements in a number of metabolic cardiovascular risk factors without producing any clinically relevant change in

hsCRP<sup>22</sup>. Further work addressing the true role of hsCRP as a surrogate for *changing* cardiovascular risk is needed to address this issue.

To our knowledge, we are the first to describe a relationship between GL and cardiorespiratory fitness. Other reports have examined the relationships of GL and GI with self reported physical activity<sup>4, 5, 7, 24</sup>, but cardiorespiratory fitness has not been previously assessed in this context. The inverse relationship between GL and cardiorespiratory fitness prompts the conclusion that individuals who are more physically fit consciously choose a lower GL diet. However, this supposition seems unlikely given that this is a sedentary, middle-aged population that is relatively unfit and untrained. Based on other studies of GI/GL effects on metabolism, we hypothesize that lower GL diets may promote a higher  $VO_{2max}$ . When administered 3 hours prior to a submaximal exercise bout, low GI diets promote increased fat oxidation and muscle glycogen sparing during exercise<sup>25</sup>. In the context of a low GL diet, performance on a maximal exercise test, represented here as  $VO_{2max}$ , may be enhanced by a longer duration of fat oxidation and subsequent delay in shift to anaerobic glycolysis. Further studies are necessary to support or refute this hypothesis.

To summarize, we have shown that, in sedentary, overweight to mildly obese, dyslipidemic individuals, GL is inversely related to cardiovascular fitness. Also, our work suggests a potential gender-specific relationship between elevated hsCRP concentrations and a diet high in simple carbohydrates and in carbohydrate quantity. In addition to our limited ability to detect interactions in a moderately sized sample, another potential limitation of this study is the use of a relatively homogenous population of sedentary, dyslipidemic, overweight to mildly obese individuals. This could explain the disappearance of the GL/hsCRP relationship after adjustment for energy intake. Finally, our cross-sectional design precludes our ability to infer causal relationships. However, while recognizing these limitations, we urge others to view current literature with skepticism and recommend that future investigations relating cardiovascular risk, specifically hsCRP, and dietary carbohydrate intake will carefully consider potential gender disparities.

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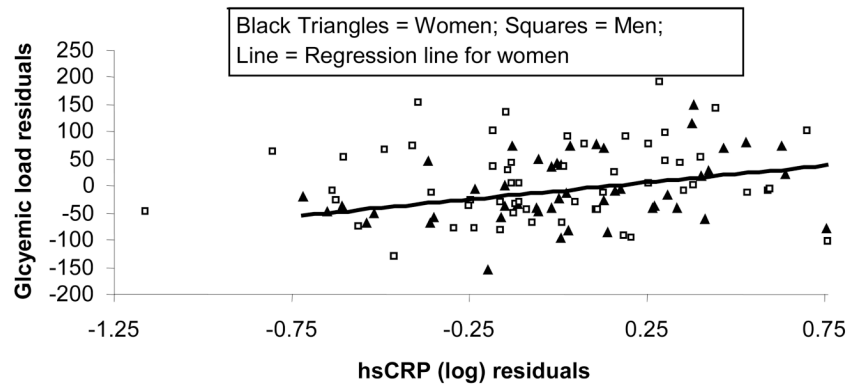
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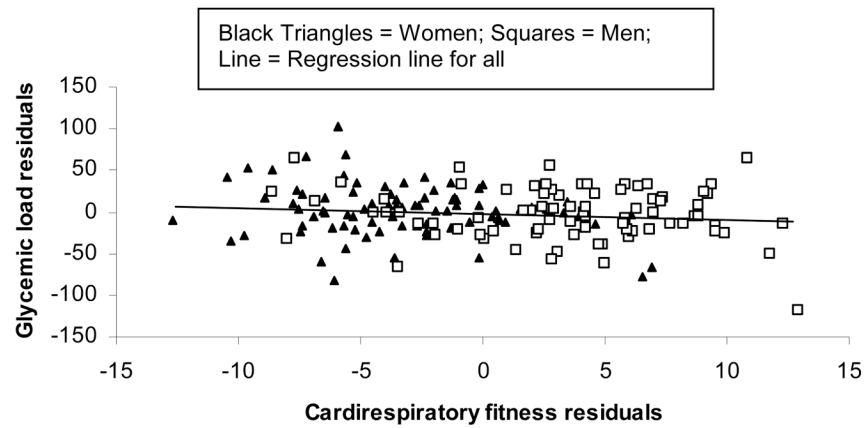
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**Figure 1. Relationship between glycemic load and hsCRP after adjusting each for their relationship with HDL-cholesterol concentrations**

$hsCRP (log) = 0.403 * HDL\text{-cholesterol} + 0.001 * Glycemic\ load - 0.361$ ;  $R^2=0.14$ ;  $P<0.0005$ ;  
 $hsCRP_{women} (log) = 0.357 * HDL\text{-cholesterol} + 0.002 * Glycemic\ load - 0.397$ ;  $R^2=0.25$ ;  
 $P<0.002$ . Relationship for men alone was not significant. hsCRP= high sensitivity C-reactive protein.



**Figure 2. Energy intake adjusted relationship between glycemic load and cardiorespiratory fitness (VO<sub>2</sub> max)**

Multivariable linear regression was used to predict cardiorespiratory fitness after adjustment for energy intake:  $n = 171$ , Cardiorespiratory fitness (ml\*min/kg) =  $-0.03*GL + 0.005*kcal/day + 23.2$ ;  $R^2 = 0.12$ , model  $P < 0.0001$ ,  $P_{GL} < 0.02$ . GL remains an independent predictor ( $P_{GL} < 0.03$ ) of cardiorespiratory fitness after accounting for gender. GL = Glycemic load.

Table 1

## Baseline Characteristics\*

	All subjects	Women	Men
<b>n</b>	171	88	83
<b>Age<sup>†</sup> (y)</b>	53.0±6.6 (40–69)	54.6±5.3 (43–66)	51.3±7.4 (40–69)
<b>Race</b>			
Asian	4	3	1
Black	25	13	12
Hispanic	1	0	1
White	141	72	69
<b>BMI (kg/m<sup>2</sup>)</b>	29.6±3.0 (23.3–38.1)	29.2±3.1 (23.3–38.1)	29.9±2.8 (24.7–36.2)
<b>hsCRP<sup>‡</sup> (log) (mg/L)</b>	2.2±2.8 (0.1–58.5)	2.9±2.9 (0.2–58.5)	1.7±2.4 (0.1–22.8)
<b>Total energy<sup>†</sup> (kcal)</b>	1789±706 (595–4045)	1582±578 (595–3280)	2008±765 (703–4045)
<b>Glycemic load<sup>†</sup></b>	155.3±65.9 (14.2–381.4)	141.0±59.1 (14.2–381.4)	170.5±69.5 (16.5–367.3)
<b>Glycemic index (Bread = 100)</b>	75.9±5.9 (31.3–86.8)	76.0±5.3 (57.0–86.8)	75.8±6.6 (31.3–84.7)

hsCRP = high sensitivity C-reactive protein

\* Data presented as means ± SD. Ranges are depicted in parentheses.

<sup>†</sup> Significant gender differences present:  $P < 0.05$  (Unpaired Student's *t*-test).

<sup>‡</sup> Geometric means presented.

**Table II**  
Relationships for hsCRP (log) (mg/L) and glycemic load, glycemic index, and daily carbohydrate.\*

	All subjects n=136			Women n=66			Men n=76		
	$\beta$ coefficient	P value	Model $R^2$	$\beta$ coefficient	P value	Model $R^2$	$\beta$ coefficient	P value	Model $R^2$
<b>GL –</b>									
Unadjusted	0.000322	0.53	<0.01	0.001354	0.12	0.04	0.000253	0.68	<0.01
Adjusted- HDLc	0.001134	0.04	0.14 <sup>†</sup>	0.002126	0.01	0.25 <sup>†</sup>	0.000524	0.49	0.01
Adjusted- HDLc + EI	0.001610	0.18	0.14 <sup>†</sup>	0.002712	0.10	0.26 <sup>†</sup>	-0.000352	0.85	0.01
<b>GI (Bread = 100) –</b>									
Unadjusted	0.000067	0.99	<0.01	-0.009233	0.34	0.01	0.006140	0.36	0.01
Adjusted- HDLc	0.005164	0.36	0.11 <sup>†</sup>	-0.0012543	0.89	0.14 <sup>†</sup>	0.006512	0.38	0.01
Adjusted- HDLc + EI	0.003686	0.52	0.13 <sup>†</sup>	-0.0030889	0.74	0.21 <sup>†</sup>	0.005440	0.47	0.02
<b>DC (g) –</b>									
Unadjusted	0.000273	0.47	<0.01	0.001130	0.08	0.05	0.000152	0.75	<0.01
Adjusted- HDLc	0.000862	0.03	0.14 <sup>†</sup>	0.001636	0.01	0.26 <sup>†</sup>	0.000375	0.52	0.01
Adjusted- HDLc + EI	0.001433	0.14	0.15 <sup>†</sup>	0.002995	0.04	0.28 <sup>†</sup>	-0.0004429	0.75	0.02

hsCRP=high sensitivity C-reactive protein GI=glycemic load EI=energy intake GL=glycemic index DC=daily carbohydrate intake

\* Data analyzed with multivariable linear regression.

<sup>†</sup>  $P < 0.05$  for the model, indicating a significant relationship between all “predictors” and hsCRP; P value listed within the table reflects whether the individual carbohydrate intake measure was significantly and independently related to hsCRP.