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# Dietary fiber intake, dietary glycemic load, and the risk of gestational diabetes mellitus during the second trimester: A nested case-control study

doi: 10.6133/apjcn.202109/PP.0003 Published online: September 2021

Running title: Dietary fiber in various food groups and GDM risk

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#### ABSTRACT

Background and Objectives: The effect of fiber, especially the effect of specific fiber in different food groups, on gestational diabetes mellitus (GDM) has seldomly been investigated. This study aimed to examine the association between GDM risk and consumption of total fiber, fiber in specific food groups, and glycemic load (GL) in the second trimester in Chinese women. Methods and Study Design: A total 162 GDM cases were matched to 324 controls on women's age and pre-pregnancy BMI. Dietary survey was conducted twice to evaluate dietary factors between 13-16 gestational weeks (GW) and 21-24 GW respectively. Multivariable logistic regression analysis was used to compute the odds ratios (ORs) and 95% confidence intervals (CIs). Results: Intake of total fiber and fruit fiber in both 13-16 GW and in 21-24 GW were significantly correlated with decreased risk of GDM, with adjusted ORs (95% CIs): 0.06(0.03-0.13) and 0.03(0.01-0.08) for total fiber in the highest quartile, 0.003(0.0002-0.02) and 0.01(0.001-0.02) for fruit fiber in the highest quartile, respectively. In contrast, consumption of cereal fiber in 21-24 GW and daily average GL in 13-16 GW were positively associated with GDM risk, with adjusted ORs (95% CIs) of the highest quartile: 3.34(1.45-7.92) and 3.88(1.43-10.89) respectively. Conclusions: Our findings suggested consumption of dietary fiber in various food groups in the second trimester might be associated with GDM risk. Particularly, diet rich in total fiber and fruit fiber may play a protective role.

# Key Words: gestational diabetes mellitus, dietary fiber, glycemic load, total fiber, fruit fiber

# **INTRODUCTION**

Gestational diabetes mellitus (GDM) is a common complication during pregnancy that brings a wide range of significant short-term and long-term effects on mother and offspring. It comprises 5.8%-12.9% of all pregnancies worldwide.<sup>1</sup> Surprisingly, the prevalence of GDM in mainland China has sharply increased to 14.8% according to a meta-analysis of research published between 2010-2017.<sup>2</sup> GDM poses adverse effects on both mother and children. Women with GDM had an increased risk of preeclampsia, cesarean delivery, and type 2 diabetes (T2DM) in later years after pregnancy.<sup>3</sup> Children from GDM pregnancy are more likely to develop hypoglycemia after birth, become obese and develop diabetes in their adulthood.<sup>3</sup> Substantial evidence from observational studies and interventional studies supports that lifestyle and diet are modifiable factors for preventing and treating GDM. For instance, a systematic review of intervention and observational studies has shown that consumption of fiber-rich foods - fruits, vegetables, nuts and legumes -- was protective factor for populations at the risk of GDM in North America, Europe, Australia, Malaysia, Korea and Iran.<sup>4,5</sup> Although some large observational studies in the U.S. have identified that dietary fiber is adversely associated with T2DM risk in middle-aged and older men and women,<sup>6-10</sup> the evidence for population at the risk of GDM is limited. Particularly, the effect of specific fiber from different food groups on GDM risk in Chinese population has seldom been investigated.

In addition, studies have also emphasized on the beneficial effects of a low glycemic index (GI) diet.<sup>5</sup> Glycemic load (GL), which represents overall glycemic response of an individual food with specific amount of carbohydrate,<sup>11</sup> covers more comprehensive effects of glycemia. It measures not only the GI of each food but also the amount of carbohydrate in each food. Research have shown that high GL was a risk factor for diabetes mellitus (DM) in non-pregnant populations.<sup>12,13</sup> The association between GL and the risk of GDM has also been examined in some observational studies;<sup>14-16</sup> however, findings of these studies were controversial. A large cohort study investigated the habitual fruit intake prior GDM revealed that GL was positively related to GDM risk, while two studies indicated that GL was not associated with GDM. Meanwhile, researchers have suggested that dietary fiber and its source, such as whole grains, fruits, and vegetables, also play a role in the associations between GL and glucose homeostasis.<sup>17</sup>

For most women, the second trimester is more comfortable, compared with the first trimester when the gastrointestinal symptoms frequently happened and the third trimester when there was only limited space left for the digestive system after the fetus growing bigger. Therefore, dietary intake in the second trimester is relatively stable for most women. Accordingly, the present study aimed to investigate the relationship among fiber and its sources, GL and GDM in a sample of Chinese pregnant women in their second trimester.

#### **MATERIALS AND METHODS**

#### Study design and population

A 1:2 nested case-control analysis matched by age and pre-pregnancy BMI was conducted from a cohort of 1261 Chinese women. Participants were recruited at 11 hospitals from May 20th, 2012 to Dec 31st, 2013. The primary purpose of the cohort study was to investigate the relationship of dietary intake and maternal body composition with the risk of GDM, which was described in our previous paper.<sup>18</sup>

Pregnant women were recruited when they established their obstetric records for the regular prenatal examination. The inclusion criteria were: 1) age between 18 to 45 years old; 2) conceive a singleton fetus; 3) 12-18 gestational weeks (GW); 4) plan to deliver in the current maternity units. Exclusion criteria were a history of GDM or any types of diabetes mellitus, or any concomitant disease. Written informed consent was acquired by each subject before participation. The study was approved by the Human Subjects Committees of the PLA General Hospital and written informed consent was acquired by each enrolled subject before participation.

#### Assessment of dietary intakes

The method of 24-hour dietary recall on 3 consecutive days was used to evaluate the intakes of energy, protein, fat, carbohydrate, and dietary fibers in all subjects twice during the second-trimester (13-16 GW and 21-24 GW) before the screening of GDM. The quantization of dietary fiber was based on the China Food Composition Table and Food Nutrition Composition Database from U.S. Department of Agriculture.

GL value of each food item by one serving size was searched in International Table of Glycemic Index and Glycemic Load Values (International Table)<sup>19</sup> and University of Sydney online databases (http://www.glycemicindex.com). The daily total GL was calculated by the following formula:

total GL (g)= $\Sigma$  [(daily consumption of food item i (g))/(serving size in Intenational Table for food i ) × GL of food i]

The final daily average GL was the average of total GL values of three surveyed days.

# Assessment of outcomes

The primary outcome was GDM which was diagnosed as one or more abnormal values from a 2h, 75g oral glucose tolerance test (OGTT) between 24 and 28 GW according to the criteria of The International Association of Diabetes and Pregnancy Study Groups (IADPSG) standard protocols. Threshold glucose values were as follows: fasting: 5.1 mmol/L, 1 h: 10.0 mmol/L, 2 h: 8.5 mmol/L.

#### Assessment of covariates

Sociodemographic characteristics, lifestyle, and pregnancy-relating medical information were collected by questionnaire during the first visit. Covariates were selected based on clinical

relevance or previously reported as risk factors for GDM in other studies,<sup>20-22</sup> including education level, cigarette smoking status, passive smoking status, alcohol intake, gravidity, and parity. Pre-pregnancy body mass index (BMI) was calculated with self-reported height and pre-pregnancy body weight. Baseline clinical characteristics, such as albumin, triglycerides, and cholesterol were tested routinely before they established the obstetric records and were retrieved as baseline information.

#### Statistical analysis

To calculate the sample size, there are six factors to consider:<sup>23</sup> 1) number of controls per case which is 2; 2) the estimated odds ratio being 0.41 when comparing the highest versus the lowest quartile of the primary independent variable (fruit consumption), according to a previous study; 24 3) a two-sided test with  $\alpha$ = 0.05; 4) a power=0.8; 5) probability that a control is exposed to the primary independent variable (adequate fruit fiber) which we chose 0.55, representing 55% of the control group who consumed more than 200g fruits (the lower level of recommended fruit consumption for pregnant women) based on our previous work;18 6) correlation between case and control which is 0.2. The calculated sample size with PASS11.0 (NCSS) for the case group is at least 73. Actually, we have included all the pregnant women who have developed GDM (n=162) in our multicenter cohort of 1123 women which we have described in the results part.

For the baseline data, continuous variables were presented as mean  $\pm$  standard deviation and Student t-test was performed to test differences between two groups; while discrete variables were presented as frequencies (%) and Pearson's  $\chi 2$  test/ Fisher's exact test was used to analyze group differences.

Multivariable logistic analysis was used to examine the association between GDM risk and dietary factors adjusting nondietary and dietary covariates. Each dietary factor was analyzed separately to estimate odds ratios (ORs) and 95% confidence intervals (CIs). Three models were established to adjust for different covariates. In Model 1, education level (junior high or less, middle school and vocational school, college, university or higher), cigarette smoking status (never, less than 1 cigarette per day, 1-5 cigarettes per day, or more than 6 cigarettes per day), passive smoking status (None, sometimes, or often), alcohol intake (none or sometimes), gravidity (0 or  $\geq$ 1), and parity (0 or  $\geq$ 1) were adjusted. In Model 2, we adjusted for variables in Model 1, energy(quartile), protein intake (quartile), fat intake (quartile), and gestational weight gain (GWG). In the stage of 13-16 GW, GWG was weight change before 13 GW; whereas in the stage of 21-24 GW, GWG was weight change between 13 to 24 GW.

In Model 3, we adjusted for variables in Model 2 and other fiber types (cereal, fruit, vegetable, and legume in quartile).

All statistical analysis was performed with SPSS 22.0 for windows (SPSS, Chicago, IL) and R version 3.6.3 (R Foundation for Statistical Computing). A two-sided p value <0.05 was considered statistically significant.

## RESULTS

# Sociological and clinical characteristics of study population

Within the original cohort of 1261 women, 138 subjects were excluded for the following reasons: not singleton pregnancy (n=69), had more than 15% blanks in their questionnaires (n=26), had apparent mistake in their dietary recall (n=23), underwent miscarriage (n=12), and lost to follow-up (n=8). Therefore, 1123 women finished the observation after they completed their OGTT, among which 162 subjects developed GDM. For the 162 GDM cases, each was matched two controls on maternal age and pre-pregnancy BMI.

Sociological and clinical characteristics are given in Table 1. All sociodemographic and medical characteristics were comparable between two groups except the use of alcohol. Women in GDM group had significantly higher frequency of alcohol use than the control group (p<0.05). No significances were seen in education, smoking habits/environment, gravidity, and parity. Among those who consumed alcohol in two groups, there were no difference in wine category, drinking frequency, or consumption amount.

# Baseline clinical characteristics and weight (BMI) change

Women with GDM had significant higher weight gain between 13 GW and OGTT (medians: 9.3 kg vs 8.5 kg, p<0.05; Table 2) and BMI gain between 13 GW and OGTT (medians: 3.5 kg/m<sup>2</sup> vs 3.2 kg/m<sup>2</sup>, p <0.05; Table 2) than controls. Other baseline clinical characteristics were comparable between the two groups.

#### Dietary intake in 13-16 GW and association with gestational diabetes

Dietary intakes between GDM group and control group in 13-16 GW were significantly different except seafood, eggs, soybean and legume fiber, with higher total energy, protein, fat, carbohydrate, cereal fiber, and daily average GL in GDM group and higher total dietary fiber, fruit fiber, and vegetable fiber in control group (Table 3).

The effects of dietary fiber intakes and GL in 13-16 GW on GDM risks are shown in Table 4. The likelihood of GDM decreased as the consumption of total fiber and fruit fiber increased

in quartiles. Compared with the lowest quartile, higher total fiber consumption was associated with lower likelihood of GDM (highest vs. lowest quartile in Model 2: OR, 0.06; 95% CI 0.03 to 0.13). Likewise, as the consumption of fruit fiber increased in quartiles, the likelihood of GDM decreased in gradient. The adjusted ORs in Model 3 across the lowest to highest quartiles of fruit fiber consumption were 1.00 (reference), 0.03(95% CI 0.01 to 0.08), 0.01(95% CI 0.001to 0.02), and 0.003(95% CI 0.0002 to 0.02), respectively. Vegetable fiber consumption was initially negatively associated with GDM after adjusting energy, protein, fat, weight change, and other non-dietary covariables, but this association was attenuated to null in Model 3 after adjusting other types of fiber (highest vs. lowest quartile in Model 3: OR, 0.45; 95% CI 0.18 to 1.11).

On the contrary, greater consumption of GL was associated with higher likelihood of GDM. Compared with the lowest quartile, the highest quartile for daily average GL consumption was associated with a higher likelihood of GDM (adjusted OR in Model 3, 3.88; 95% CI 1.43 to 10.89). At first, cereal fiber consumption shown positive association with GDM after adjusting energy, protein, fat, weight change, and other non-dietary covariables, but this association was attenuated to null in Model 3 (highest vs. lowest quartile in Model 3: OR,1.21; 95% CI 0.47 to 3.07). Additionally, no significant association was observed between GDM and legume fiber consumption.

#### Dietary intake in 21-24 GW and association with gestational diabetes

In 21-24 GW, GDM group still consumed higher total energy, protein, carbohydrate, cereal fiber, and high GL food with significance compared with control group, while control group ate significantly more vegetables and fruits which result in higher total dietary fiber and fruit fiber consumption. Meanwhile, GDM group consumed more legume fiber in a similar manner to 13-16 GW, but with statistical significance. There was no difference in intakes of fat and vegetable fiber between GDM group and control group.

As shown in Table 5, similar to the GW 13-16, the likelihood of GDM decreased as the intake of total fiber and fruit fiber increased. Compared with the corresponding lowest quartiles, the highest quartile for consumption of total fiber and fruit fiber intake were each related to a lower likelihood of GDM (adjusted OR in Model 2, 0.03; 95% CI 0.01 to 0.08; adjusted OR in Model 3, 0.01; 95% CI 0.001 to 0.03, respectively). In parallel with the findings in 13-16 GW, the association between vegetable fiber intake and GDM was inverse after adjusting energy, protein, fat, weight change, and other non-dietary covariables, but was

attenuated to null after adjusting other types of fiber (highest vs. lowest quartile in Model 3: OR, 0.49; 95% CI 0.19 to 1.28).

In contrast, compared with the lowest quartile, the highest quartile for consumption of cereal fiber was associated with a higher likelihood of GDM even after adjusted other types of dietary fiber and other confounding factors (adjusted OR in Model 3, 3.34; 95% CI 1.45 to 7.92). Compared with the lowest quartile, the highest quartile for GL consumption was positively associated with the likelihood of GDM in Model 2, but the association attenuated to null after adjusting all dietary fibers in each subgroup (adjusted OR in Model 3, 2.11; 95% CI 0.76 to 5.98). Moreover, no significant association was observed between legume fiber consumption and GDM.

#### DISCUSSION

In our previous study, we found that body weight, energy, carbohydrate, fat, and protein during the second trimester were associated with an increased risk of GDM,<sup>18</sup> which drew attention to the importance of dietary factors in the second trimester. In our nested casecontrol study, we collected data of dietary intake at the beginning of the second trimester and the time point right before the OGTT test. In this study, we observed that intake of total fiber and fruit fiber in 13-16 GW and in 21-24 GW were significantly correlated with decreased risk of GDM. In contrast, consumption of daily average GL in 13-16GW and cereal fiber in 21-24 GW was associated with increased risk of GDM. We are unaware of other studies that focused on the association between fiber consumption in the second trimester and GDM risk. Our findings of the protective role of total fiber and fruit fiber in GDM were conforming to the NHSII which involved 13,110 subjects and explored the association between habitual prepregnancy fiber intake and GDM risk.<sup>15</sup> There are several potential mechanisms that lead to the protective role of total fiber and fruit fiber. First, dietary fiber may increase satiety and lower the intake of energy.<sup>25</sup> Second, fiber, especially the soluble fiber, can also slow glucose absorption and subsequently promote insulin levels.<sup>26</sup> Third, adequate amount of fiber could promote growth of gut bacteria that could generate short-chain fatty acids and decrease the likelihood of insulin resistance in pregnant women.<sup>27</sup>

Studies focused on fruit fiber intake in the second trimester and GDM is limited. However, there are many studies examining relationship between fruit consumption and GDM risk. A Chinese prospective cohort study in Wuhan involving 3,300 participants found that higher overall fruit consumption in second trimester is protective in GDM.24 The likelihood of GDM decreased 59% in the highest quintile of fruit consumption (median, 671 g/d). There are

many studies that support the protective role of fruit intake in type 2 diabetes.<sup>28-30</sup> Yet, another Chinese cohort study in Guangdong indicated that excessive consumption of fruit (the median of the highest quartile was 710 g) during the second trimester was associated with risk of GDM.<sup>31</sup> In fact, they consumed much more citrus fruit and tropical fruit with moderate-to-high GI values than our population due to geographic difference, and fruits consumed the most in our study were low-GI, fiber-rich fruit, such as apple, pear, peach, and berries. Therefore, GI/GL value of fruits may play a contributive role in the association between fruit/fruit fiber intake and GDM risk. This is in accordance with the finding in the Wuhan cohort that low GL fruits, rather than high GL fruits, were associated with a lower GDM risk.<sup>24</sup>

To the best of our knowledge, no previous studies have especially examined the association between legume fiber and GDM. Our study found no association between legume fiber and GDM likelihood. Large prospective studies involving 75,344 and 71,346 subjects respectively found that moderate to high intake of soy foods was associated with a slightly higher T2DM risk.<sup>32,33</sup> However, a series of other observational studies described a protective role of legume consumption on diabetes risk.<sup>34-36</sup> The reason behind this discrepancy could be different kinds of legume products consumed besides legume fiber. When divided the legume products into unsweetened and sweetened ones, researchers found that unsweetened one was associated with inversely associated with diabetes risk; whereas sweetened one was positively associated with diabetes risk.<sup>34</sup> Therefore, further studies analyzing the influence of legume fiber and added sugar in the legume products on GDM risks are needed in the future.

When analyzed in Model 2 adjusting energy, protein, fat, weight change, and other nondietary covariables, consumption of cereal fiber in both 13-16 GW and in 21-24 GW were positively associated with GDM. This association still existed in 21-24 GW after adjusting other types of dietary fibers. However, this finding was inconsistent with the NHSII in which the highest quintile of cereal fiber was related to a lower likelihood of GDM.<sup>15</sup> The cut point of the highest quartile/quintile levels of cereal fiber in the current study and the NHSII were 3.5 g/day and 7.2 g/day respectively, which revealed the consumption of cereal fiber was much more in the NHSII population. The probable reason for the discrepant role of cereal fiber in the two studies could be that the population in our study did not consume enough cereal fiber. Moreover, the percentage of whole grain fiber in cereal fiber is also important. In a meta-analysis,<sup>28</sup> researchers found that whole grains consumption, but not refined grains consumption, played a protective role in T2DM. In China, staple food is mainly refined rice and flour which are low in dietary fiber; therefore, it is hard to reach such a high consumption of cereal fiber in general population. The more cereal fiber one consumed in China; the more carbohydrate was consumed alongside. As a result, the association between cereal fiber and GDM may due to the excessive refined grains intake.

In this study, the protective association between vegetable fiber and GDM was approaching but not yet significant. Correspondingly, the NHSII found no association between prepregnancy vegetable fiber consumption and GDM risk even though there seemed to be a reduction in the risk in higher consumption subgroups.<sup>15</sup> Moreover, in a meta-analysis of 13 prospective studies of 63299 individuals about relationship between vegetable intake and T2DM risk found only a borderline inverse association for the high vs. low (RR: 0.95; CI 0.89-1.01),<sup>29</sup> with inverse association observed only in Asian and Australian studies and in studies with smaller samples. Therefore, studies investigating association between vegetable fiber/vegetable consumption with GDM risks in larger Chinese pregnant women populations is required.

The highest quartile of daily average GL (median 163 g/d) was positively associated with the risk of GDM in both 13-16 GW and 21-24 GW adjusting energy, protein, fat, weight change, and other non-dietary covariables. This association was still significant in 13-16 GW after adjusting other types of dietary fibers. This finding is in parallel with the NHSII in which the highest quintile of GL (median 212 g/d) was associated with increased likelihood of GDM.<sup>15</sup> A dose-response meta-analysis including prospective cohort studies from 1946 to December 2018 also indicated that higher GL substantially increased the likelihood of T2DM.<sup>37</sup> Another meta-analysis confirmed this robust and consistent positive relation between GL and T2DM risk, with 42% increase risk for a 100g increment in GL.<sup>38</sup> In a study of 306 young female, Arikawa et al found that high GL was marginally associated with inxidi and the stress which was considered to be involved in the etiology of diabetes process.<sup>39</sup> Additionally, a cross-sectional study found that high GL was associated with insulin resistance.<sup>40</sup>

The strength of this study including: a relatively large sample size from multi-centers prospective study, and analyzing association of GDM risk with dietary fiber in various food subgroups. A limitation of present study is the reliability of self-reported dietary recall. Also, the GL was calculated using an international database in which most foods were derived from western countries; therefore, GL of some foods in this study may not been calculated correctly, especially some traditional foods that were unique in Chinese culture. Moreover, we didn't include free sugar in our dietary survey since the major sources come not only from sugar-sweeten foods, but also as a condiment in the cooking process. Notably, the relatively

small OR values in fruit fiber consumption in all three models suggesting that sample size of this research may be limited. Besides, the other reason for these small OR values may be the very small variance of fruit fiber consumption as a continuous variable in the studied population. The interquartile of fruit fiber consumption lies between 3.35 g to 5.42 g. One gram of fruit fiber could mean a lot of real fruits, such as 250 g grapes/orange or 333 g cherries. Therefore, even though the number of fruit fiber didn't change much in each quartile, the actual fruit consumption could be quite different. Correspondingly, the small OR of highest quartile versus the lowest quartile may reveal the GDM risk of huge fruit consumption. Further research with larger sample size on fiber consumption of various food groups in the second trimester is needed to verify our findings.

#### Conclusion

In conclusion, we investigated relationship of GDM risk to the intake of different sorts of dietary fibers and total dietary GL by dietary surveys in the second trimester. Our findings suggested that total fiber and fruit fiber were associated with lower risk of GDM. On the contrary, cereal fiber and daily average GL were positively associated with risk of GDM. This implies that pregnant women should meet daily recommendations for each food category to gain proper fiber intake and limit high GL foods for preventing GDM.

# ACKNOWLEDGEMENTS

We are grateful to all subjects who participated in this study, and to Dr. Botao Guo for his help in data analysis. We also sincerely thank all the participants who worked on this study in the 11 clinical centers: the First Medical Center of PLA General Hospital General Hospital, the Fourth Medical Center of PLA General Hospital, Beijing Hospital, Tongren Hospital, Haidian District Health Center for Women and Children, Jishuitan Hospital, the Sixth People's Hospital of Shanghai, Chinese PLA General Air Force Center, People's Hospital of Sichuan Province and Bethune International Peace Hospital.

#### AUTHOR DISCLOSURE

The authors declare no conflict of interest.

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 Table 1. The characteristics of participating restaurants (n=90)

|  | GDM case subjects | Control subjects | p value |
|--|-------------------|------------------|---------|
| N  | 162               | 324              |         |
| Maternal age (years)                       | 30.65±3.23        | 30.23±2.90       |         |
| 18-25                                      | 6 (3.7)           | 12 (3.7)         |         |
| 26-30                                      | 69 (42.6)         | 138 (42.6)       |         |
| 31-35                                      | 77 (47.5)         | 154 (47.5)       |         |
| 36-40                                      | 9 (5.6)           | 18 (6.2)         |         |
| 41-45                                      | 1 (0.6)           | 1 (0.6)          |         |
| Pre-pregnancy BMI (kg/m <sup>2</sup> )     | 22.09±3.40        | 22.08±3.41       |         |
| Under weight (lower than 18.5)             | 17 (10.5)         | 34 (10.5)        |         |
| Normal weight (18.5-23.9)                  | 107 (66.0)        | 214 (66.0)       |         |
| Overweight (24-27.9)                       | 24 (14.8)         | 48 (14.8)        |         |
| Obese (28 or higher)                       | 14 (8.6)          | 28 (8.6)         |         |
| Education                                  |                   |                  | 0.362   |
| Lower than junior high school              | 1 (0.6)           | 9 (2.8)          |         |
| Senior middle school and vocational school | 10 (6.2)          | 15 (4.6)         |         |
| College                                    | 41 (25.3)         | 75 (23.1)        |         |
| Higher than university                     | 110 (67.9)        | 225 (69.4)       |         |
| Smoke                                      |                   |                  | 0.086   |
| None                                       | 160 (98.8)        | 256 (79.0)       |         |
| Less than 1 cigarette per day              | 1 (0.6)           | 68 (21.0)        |         |
| 1-5 cigarettes per day                     | 0                 | 0                |         |
| 6-9 cigarettes per day                     | 1 (0.6)           | 0                |         |
| More than 10 cigarettes per day            | 0                 | 0                |         |
| Passive smoking                            |                   |                  | 0.069   |
| None                                       | 65 (40.1)         | 122 (37.7)       |         |
| Sometimes                                  | 86 (53.1)         | 157 (48.5)       |         |
| Often                                      | 11 (6.8)          | 45 (13.9)        |         |
| Alcohol                                    |                   |                  | 0.042   |
| None                                       | 114 (70.4)        | 256 (79.0)       |         |
| Sometimes                                  | 48 (29.6)         | 68 (21.0)        |         |
| Wine category                              |                   |                  | 0.176   |
| Spirit                                     | 7 (4.3)           | 10 (3.1)         |         |
| Wine                                       | 20 (12.3)         | 36 (11.1)        |         |
| Beer                                       | 20 (12.3)         | 21 (6.5)         |         |
| Yellow rice or millet wine                 | 0                 | 0                |         |
| Others                                     | 1 (0.6)           | 1 (0.3)          |         |
| Drinking frequency                         |                   | · ·              | 0.064   |
| Less than 1-2 times per week               | 42 (25.9)         | 63 (19.4)        |         |
| 1-2 times per week                         | 3 (1.9)           | 1 (0.3)          |         |
| 3-4 times per week                         | 0                 | 2 (0.6)          |         |
| 5-7 times per week                         | 3 (1.9)           | 2 (0.6)          |         |
| Alcohol consumption                        | · · /             | . /              | 0.207   |
| Less than 1 cup per time                   | 44 (27.2)         | 62 (19.1)        |         |
| 1-2 cups per time                          | 3 (1.9)           | 5 (1.5)          |         |
| 3-4 cups per time                          | 1 (0.6)           | 1 (0.3)          |         |
| Gravidity                                  |                   | . /              | 0.918   |
| None                                       | 111 (68.5)        | 219 (67.6)       |         |
| More than 1 time                           | 51 (31.5)         | 105 (32.4)       |         |
| Parity                                     | ()                | ()               | 0.555   |
| None                                       | 162 (100.0)       | 322 (99.4)       |         |
| More than 1 time                           | 0                 | 2 (0.6)          |         |

BMI; body mass index, (kg/m<sup>2</sup>).  $^{\dagger}Data$  are presented as n (%) or mean  $\pm$  standard deviation.

Table 2. Clinical characteristics of the study population

|                            | GDM (n=162)     | Control (n=324)  | p value |
|----------------------------|-----------------|------------------|---------|
| Albumin (g/L)              | 42.8±5.04       | 42.9±3.71        | 0.711   |
| BUN (mmol/L)               | 2.9±0.89        | 2.9±0.86         | 0.501   |
| sCr (mmol/L)               | 47.8±8.3        | 47.6±12.3        | 0.855   |
| TG (mmol/L)                | $1.4\pm0.64$    | $1.4\pm0.70$     | 0.877   |
| TC (mmol/L)                | 4.7±0.84        | 4.6±0.78         | 0.222   |
| HDL-c (mmol/L)             | 1.7±0.34        | 1.7±0.31         | 0.808   |
| LDL-c (mmol/L)             | 2.5±0.63        | 2.4±0.60         | 0.137   |
| SBP (mmHg)                 | 109.2±8.19      | $109.9 \pm 8.01$ | 0.380   |
| DBP (mmHg)                 | 70.2±6.43       | 70.9±6.78        | 0.286   |
| WG before 13GW             | 1.5±1.79        | $1.3 \pm 1.50$   | 0.135   |
| WG between 13GW and OGTT   | 9.3±3.47        | 8.5±3.38         | 0.027   |
| BMIG before 13 GW          | $0.55 \pm 0.68$ | $0.47 \pm 0.70$  | 0.264   |
| BMIG between 13GW and OGTT | 3.5±1.30        | 3.2±1.26         | 0.025   |

TG: triglyceride; TC: total cholesterol; HDL-c: high density lipoprotein cholesterol; LDL-c: low density lipoprotein cholesterol; SBP: systolic blood pressure; DBP: diastolic blood pressure; WG: weight gain; GW: gestation week; OGTT: oral glucose tolerance test; BMIG: body mass index gain.

Data are n (%) or mean  $\pm$  standard deviation.

**Table 3.** Dietary intake in the second trimester pregnancy

|                           | 13-1              | 6 GW            |               | 21-2              |                   |           |  |
|---------------------------|-------------------|-----------------|---------------|-------------------|-------------------|-----------|--|
|                           | GDM (n=162)       | Control (n=324) | - p value $-$ | GDM (n=162)       | Control (n=324)   | - p value |  |
| Energy (kcal/d)           | 2018.3±202.7      | 1862.3±220.7    | < 0.001       | 2074.0±216.7      | 1966.3±192.2      | < 0.001   |  |
| Protein (g/d)             | 81.1±15.3         | 73.9±16.2       | < 0.001       | 84.4±16.0         | 79.8±15.1         | 0.002     |  |
| Fat (g/d)                 | 75.6±14.5         | 70.3±14.0       | < 0.001       | 77.4±17.4         | 75.6±14.0         | 0.217     |  |
| Carbohydrate (g/d)        | 245.9±32.8        | 231.0±32.0      | < 0.001       | 253.4±33.5        | 240.5±30.1        | < 0.001   |  |
| Total dietary fiber (g/d) | 11.0±3.5          | $14.4\pm4.5$    | < 0.001       | 11.7±4.7          | 15.2 <u>+</u> 4.9 | < 0.001   |  |
| Cereal fiber (g/d)        | 3.5±1.4           | 2.9±1.3         | < 0.001       | 3.5±1.4           | 3.1±1.4           | < 0.001   |  |
| Legume fiber (g/d)        | 2.9±4.0           | 2.3±3.2         | 0.108         | 3.0±3.4           | 2.3±3.5           | 0.046     |  |
| Fruit fiber (g/d)         | $1.1 \pm 1.2$     | $4.6\pm2.0$     | < 0.001       | 1.2±1.2           | 4.6±1.8           | < 0.001   |  |
| Vegetable fiber (g/d)     | $1.1 \pm 1.2$     | 4.2±3.2         | 0.032         | 4.2±3.7           | 4.7±3.2           | 0.147     |  |
| Daily average GL          | 167.2±25.6        | $148.1\pm22.4$  | < 0.001       | 170.5±26.2        | 153.0±4.9         | < 0.001   |  |
| Grains (g/d)              | 219.9±74.7        | 193.7±80.8      | < 0.001       | 233.7.5±86.7      | 209.3±78.1        | 0.002     |  |
| Vegetables (g/d)          | 231.4±42.6        | 404.7±39.4      | < 0.001       | 416.3±45.4        | 464.4±55.4        | < 0.001   |  |
| Fruits (g/d)              | $142.5 \pm 154.2$ | 462.0±154.4     | < 0.001       | $151.4 \pm 100.2$ | 415.6±125.3       | < 0.001   |  |
| Meat (g/d)                | 87.3±29.6         | 73.5±28.3       | < 0.001       | 89.2±29.3         | 81.1±23.3         | 0.001     |  |
| Seafood (g/d)             | 15.3±9.9          | 16.7±9.2        | 0.124         | 32.8±24.2         | 17.7±17.1         | < 0.001   |  |
| Eggs (g/d)                | 56.7±15.4         | 59.0±14.8       | 0.112         | 58.3±19.2         | 49.2±17.3         | < 0.001   |  |
| Dairy products (g/d)      | 210.0±80.9        | 227.0±57.3      | 0.008         | 260.4±100.0       | 258.0±92.6        | 0.793     |  |
| Soybean (g/d)             | 9.3±7.2           | 8.2±6.8         | 0.100         | 9.6±5.3           | 8.6±4.5           | 0.030     |  |
| Nuts (g/d)                | 6.8±4.3           | 7.5±3.2         | 0.044         | 13.7±8.9          | 12.5±9.4          | 0.178     |  |
| Oil (g/d)                 | 33.4±5.7          | 28.0±6.5        | < 0.001       | 38.6±18.4         | 32.3±9.6          | 0.166     |  |

|              | Quartiles     | GDM        | Control<br>(n=324) | Model 1             |         | Model 2             | Model 2 |                     | Model 3 |  |
|--------------|---------------|------------|--------------------|---------------------|---------|---------------------|---------|---------------------|---------|--|
|              |               | (n=162)    |                    | Odds ratio          | p value | Odds ratio          | p value | Odds ratio          | p value |  |
|              |               | (II=102)   |                    | (95% CI)            |         | (95% CI)            |         | (95% CI)            |         |  |
| Total fiber  | <11.11        | 93 (57.4)  | 81 (25)            | Reference           |         | Reference           |         | N/A                 | N/A     |  |
| (g)          | 11.12-13.69   | 40 (24.7)  | 81 (25)            | 0.38 (0.22-0.62)    | < 0.001 | 0.23 (0.13-0.42)    | < 0.001 |                     |         |  |
|              | 13.70-16.68   | 15 (9.3)   | 81 (25)            | 0.15 (0.08-0.29)    | < 0.001 | 0.08 (0.04-0.16)    | < 0.001 |                     |         |  |
|              | >16.68        | 14 (8.6)   | 81 (25)            | 0.14 (0.07-0.26)    | < 0.001 | 0.06 (0.03-0.13)    | < 0.001 |                     |         |  |
| Cereal fiber | <2.02         | 23 (14.2)  | 81 (25)            | Reference           |         | Reference           |         | Reference           |         |  |
| (g)          | 2.23-2.63     | 22 (13.6)  | 81 (25)            | 0.97 (0.49-1.90)    | 0.924   | 0.89 (0.43-1.83)    | 0.746   | 0.58 (0.20-1.67)    | 0.315   |  |
| -            | 2.64-3.50     | 46 (28.4)  | 81 (25)            | 1.96 (1.09-3.61)    | 0.027   | 1.47 (0.78-2.81)    | 0.231   | 0.53 (0.20-1.38)    | 0.202   |  |
|              | >3.50         | 71 (43.8)  | 81 (25)            | 3.09 (1.76-5.57)    | < 0.001 | 2.36 (1.28-4.42)    | 0.006   | 1.21 (0.47-3.07)    | 0.686   |  |
| Legume       | < 0.49        | 41 (25.3)  | 81 (25)            | Reference           |         | Reference           |         | Reference           |         |  |
| fiber (g)    | 0.50-1.32     | 30 (18.5)  | 81 (25)            | 0.77 (0.43-1.37)    | 0.380   | 0.69 (0.36-1.30)    | 0.248   | 0.22 (0.08-0.57)    | 0.002   |  |
|              | 1.33-3.01     | 38 (23.5)  | 81 (25)            | 0.94 (0.54-1.65)    | 0.841   | 0.75 (0.41-1.36)    | 0.342   | 0.61 (0.23-1.57)    | 0.305   |  |
|              | >3.01         | 53 (32.7)  | 81 (25)            | 1.48 (0.87-2.53)    | 0.151   | 1.12 (0.62-2.03)    | 0.694   | 0.68 (0.27-1.68)    | 0.407   |  |
| Fruit fiber  | <3.34         | 151 (93.2) | 81 (25)            | Reference           |         | Reference           |         | Reference           |         |  |
| (g)          | 3.35-4.28     | 8 (4.9)    | 81 (25)            | 0.06 (0.02-0.12)    | < 0.001 | 0.04 (0.02-0.10)    | < 0.001 | 0.03 (0.01-0.08)    | < 0.001 |  |
|              | 4.29-5.42     | 2 (1.2)    | 81 (25)            | 0.01 (0.002-0.04)   | < 0.001 | 0.01 (0.001-0.03)   | < 0.001 | 0.01 (0.001-0.02)   | < 0.001 |  |
|              | >5.42         | 1 (0.6)    | 81 (25)            | 0.005 (0.0003-0.02) | < 0.001 | 0.003 (0.0002-0.02) | < 0.001 | 0.003 (0.0002-0.02) | < 0.001 |  |
| Vegetable    | <2.29         | 44 (27.2)  | 81 (25)            | Reference           |         | Reference           |         | Reference           |         |  |
| fiber        | 2.30-3.55     | 52 (32.1)  | 81 (25)            | 1.21 (0.71-2.05)    | 0.482   | 1.02 (0.57-1.83)    | 0.932   | 0.91 (0.39-2.10)    | 0.818   |  |
| (g)          | 3.56-5.04     | 35 (21.6)  | 81 (25)            | 0.81 (0.46-1.41)    | 0.458   | 0.76 (0.41-1.39)    | 0.370   | 0.73 (0.30-1.73)    | 0.476   |  |
|              | >5.04         | 31 (19.1)  | 81 (25)            | 0.69 (0.39-1.22)    | 0.202   | 0.44 (0.23-0.81)    | 0.010   | 0.45 (0.18-1.11)    | 0.085   |  |
| Daily        | <136.53       | 19 (11.7)  | 81 (25)            | Reference           |         | Reference           |         | Reference           |         |  |
| average GL   | 136.54-150.31 | 17 (10.5)  | 81 (25)            | 0.96 (0.45-2.00)    | 0.905   | 0.86 (0.39-1.89)    | 0.707   | 1.38 (0.49-3.96)    | 0.540   |  |
| -            | 150.32-162.69 | 40 (24.7)  | 81 (25)            | 2.18 (1.16-4.22)    | 0.018   | 2.04 (1.00-4.22)    | 0.052   | 3.50 (1.33-9.56)    | 0.012   |  |
|              | >162.69       | 86 (53.1)  | 81 (25)            | 4.61 (2.57-8.60)    | < 0.001 | 3.40 (1.67-7.10)    | 0.001   | 3.88 (1.43-10.89)   | 0.009   |  |

**Table 4.** Relative risks of GDM according to quartiles of dietary fiber intakes and glycemic load (GL) in 13-16GW

Model 1: adjusted for education level, cigarette smoking status, passive smoking status, alcohol intake, gravidity, and parity. Model 2: adjusted for variables in Model 1, energy(quartile), protein intake (quartile), fat intake (quartile), and weight change before 13GW. Model 3: adjusted for variables in Model 2 and other fiber types (cereal, fruit, vegetable, and legume in quartile).

|              | Quartiles     | GDM<br>(n=162) | Control (n=324) | Model 1 Model 2   |         |                   |         | Model 3           |         |
|--------------|---------------|----------------|-----------------|-------------------|---------|-------------------|---------|-------------------|---------|
|              |               |                |                 | Odds ratio        | p value | Odds ratio        | p value | Odds ratio        | p value |
|              |               | (II=102)       | (11-324)        | (95% CI)          |         | (95% CI)          |         | (95% CI)          |         |
| Total fiber  | <11.11        | 93 (57.4)      | 81 (25)         | Reference         |         | Reference         |         | N/A               | N/A     |
| (g)          | 11.12-13.69   | 40 (24.7)      | 81 (25)         | 0.42 (0.25-0.70)  | < 0.001 | 0.32 (0.18-0.55)  | < 0.001 |                   |         |
|              | 13.70-16.68   | 15 (9.3)       | 81 (25)         | 0.24 (0.13-0.42)  | < 0.001 | 0.15 (0.08-0.29)  | < 0.001 |                   |         |
|              | >16.68        | 14 (8.6)       | 81 (25)         | 0.07 (0.03-0.15)  | < 0.001 | 0.03 (0.01-0.08)  | < 0.001 |                   |         |
| Cereal fiber | < 2.02        | 23 (14.2)      | 81 (25)         | Reference         |         | Reference         |         | Reference         |         |
| (g)          | 2.23-2.63     | 22 (13.6)      | 81 (25)         | 1.38 (0.71-2.70)  | 0.343   | 1.41 (0.71-2.86)  | 0.328   | 1.63 (0.64-4.25)  | 0.310   |
|              | 2.64-3.50     | 46 (28.4)      | 81 (25)         | 3.03 (1.67-5.66)  | < 0.001 | 3.59 (1.90-7.00)  | < 0.001 | 3.44 (1.49-8.25)  | 0.004   |
|              | >3.50         | 71 (43.8)      | 81 (25)         | 3.09 (1.71-5.75)  | < 0.001 | 3.48 (1.86-6.76)  | < 0.001 | 3.34 (1.45-7.92)  | 0.005   |
| Legume       | < 0.49        | 41 (25.3)      | 81 (25)         | Reference         |         | Reference         |         | Reference         |         |
| fiber (g)    | 0.50-1.32     | 30 (18.5)      | 81 (25)         | 1.20 (0.65-2.22)  | 0.557   | 1.08 (0.57-2.06)  | 0.808   | 0.62 (0.25-1.53)  | 0.304   |
|              | 1.33-3.01     | 38 (23.5)      | 81 (25)         | 1.25 (0.69-2.27)  | 0.465   | 0.93 (0.49-1.76)  | 0.829   | 0.71 (0.28-1.77)  | 0.464   |
|              | >3.01         | 53 (32.7)      | 81 (25)         | 2.17 (1.25-3.83)  | 0.006   | 1.68 (0.92-3.10)  | 0.091   | 1.60 (0.67-3.84)  | 0.291   |
| Fruit fiber  | < 3.34        | 151 (93.2)     | 81 (25)         | Reference         |         | Reference         |         | Reference         |         |
| (g)          | 3.35-4.28     | 8 (4.9)        | 81 (25)         | 0.04 (0.02-0.09)  | < 0.001 | 0.03 (0.01-0.08)  | < 0.001 | 0.03 (0.01-0.08)  | < 0.001 |
|              | 4.29-5.42     | 2 (1.2)        | 81 (25)         | 0.01 (0.002-0.04) | < 0.001 | 0.01 (0.001-0.03) | < 0.001 | 0.01 (0.001-0.03) | < 0.001 |
|              | >5.42         | 1 (0.6)        | 81 (25)         | 0.01 (0.002-0.04) | < 0.001 | 0.01 (0.001-0.02) | < 0.001 | 0.01 (0.001-0.03) | < 0.001 |
| Vegetable    | < 2.29        | 44 (27.2)      | 81 (25)         | Reference         |         | Reference         |         | Reference         |         |
| fiber (g)    | 2.30-3.55     | 52 (32.1)      | 81 (25)         | 1.04 (0.61-1.77)  | 0895    | 0.88 (0.50-1.57)  | 0.678   | 1.15 (0.51-2.57)  | 0.737   |
|              | 3.56-5.04     | 35 (21.6)      | 81 (25)         | 1.00 (0.59-1.69)  | 0.997   | 0.68 (0.38-1.21)  | 0.192   | 0.74 (0.31-1.72)  | 0.479   |
|              | >5.04         | 31 (19.1)      | 81 (25)         | 0.57 (0.31-1.01)  | 0.057   | 0.35 (0.18-0.67)  | 0.002   | 0.49 (0.19-1.28)  | 0.148   |
| Daily        | < 136.53      | 19 (11.7)      | 81 (25)         | Reference         |         | Reference         |         | Reference         |         |
| average GL   | 136.54-150.31 | 17 (10.5)      | 81 (25)         | 1.20 (0.59-2.43)  | 0.616   | 1.18 (0.56-2.48)  | 0.663   | 0.45 (0.16-1.26)  | 0.130   |
| -            | 150.32-162.69 | 40 (24.7)      | 81 (25)         | 1.42 (0.72-2.83)  | 0.317   | 1.50 (0.72-3.17)  | 0.278   | 0.55 (0.18-1.66)  | 0.296   |
|              | >162.69       | 86 (53.1)      | 81 (25)         | 5.17 (2.87-9.69)  | < 0.001 | 4.58 (2.26-9.61)  | < 0.001 | 2.11 (0.76-5.98)  | 0.156   |

Table 5. Relative risks of GDM according to quartiles of dietary fiber intakes and glycemic load (GL) in 21-24GW

Model 1: adjusted for education level, cigarette smoking status, passive smoking status, alcohol intake, gravidity, and parity. Model 2: adjusted for variables in Model 1, energy intake(quartile), protein intake (quartile), fat intake (quartile), and weight change between 13GW to OGTT test. Model 3: adjusted for variables in Model 2 and other fiber types (cereal, fruit, vegetable, and legume in quartile).