Overall Glycemic Index and Glycemic Load of Vegan Diets in Relation to Plasma Lipoproteins and Triacylglycerols

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The data do not support the hypothesis that a carbohydrate-rich diet per se is associated with unfavorable effects on triacylglycerols that would be predicted to increase the risk of coronary heart disease.

Introduction

To date, coronary heart disease (CHD) remains the leading cause of mortality in high-income countries [1, 2] and the prevalence of type 2 diabetes continues to rise by epidemic proportions [3]. There is considerable epidemiological evidence that these diseases are influenced by the life course, and especially by dietary factors [2, 4, 5]. Recently, dietary carbohydrates have received particular attention. Depending on their chemical composition and physical structure, carbohydrates are digested and absorbed at different rates and, therefore, have different physiological effects, especially on blood glucose and insulin concentrations [6].

In order to characterize the rate of carbohydrate absorption after a meal, Jenkins et al. [7] proposed the concept of the glycemic index (GI) as a physiological basis for qualitative classification of carbohydrate-containing...
foods. The GI is defined as the area under the glucose response curve (AUC) after consumption of a standard amount (50 g) of carbohydrate from a test food, and values are expressed relative to that of a control food (white bread or glucose) [8, 9]. However, the postprandial blood glucose and insulin response depends both on the quantity and quality of carbohydrates, so there is a rationale to consider them together. Therefore, the concept of the glycemic load (GL) was established by Salmeron et al. [10]. The GL of a selected food is the arithmetic product of its GI and its carbohydrate amount [10]. Several observational studies have shown that the overall GL and/or GI of the diet is/are positively associated with the risk for CHD, unfavorable effects on plasma lipoproteins and triacylglycerols [11–15] and type 2 diabetes mellitus [16–18]. In the past, metabolic studies in humans have shown that low-GI diets improve blood glucose levels, urinary C-peptide output [19] and lipid profiles [20–23]. These findings suggest health benefits of plant-based, high-fiber diets with a low GI in the prevention of degenerative diseases like CHD and type 2 diabetes mellitus [4, 6, 24]. There is epidemiological evidence that vegetarians are at low risk for these life course-related diseases and it is believed that this fact is due to the composition of their diet [25–28].

In general, lacto-(ovo-)vegetarians and vegans consume more minimally processed plant foods like fresh fruits, vegetables, whole grain products, legumes and nuts [29, 30]. Therefore it can be hypothesized that such diets are characterized by a low GI and thus a low GL. Although a large number of studies on vegetarian [31, 32] and vegan nutrition [30, 33, 34] have been published during the last years, only one small study has examined the GI of vegan diets [35]. We therefore investigated the overall GI and GL of such diets assessed by a food-frequency questionnaire (FFQ). Furthermore, this cross-sectional study evaluated the relations of dietary GI and GL as well as total fat, carbohydrate and fiber intakes to plasma concentrations of lipoproteins and triacylglycerols (TG) in 154 apparently healthy adult vegans.

Methods

Ethical Considerations

This study was conducted in accordance with the Helsinki Declaration of 1964 as amended in 1996. Since there was no intervention, the Ethic Commission of the State of Lower Saxony decided that an ethical approval was not required. All subjects were volunteers who gave written consent prior to participation and who were free to withdraw from the study at any time.

Subjects: Recruitment and Screening

Participants were recruited by advertisements in eight German magazines. In total, 154 persons took part in all study segments (pre and main questionnaire, FFQ and blood sampling) and fulfilled the following study criteria: vegan diet for at least 1 year prior to the start of the study; a minimum age of 18 years; resident in Germany; no pregnancy or childbirth during the last 12 months. Exclusion criteria were defined as: severe illness during the last 12 months (such as malignant or cardiovascular diseases, renal failure, severe diseases of the gastrointestinal tract), diagnosed blood coagulation disorder, intake of inhibitors of blood coagulation, misuse of alcohol and/or drugs, participation in another study during the last month.

Life course factors (except alcohol consumption; FFQ) were evaluated by means of the study’s main questionnaire, in which frequency, duration and kind of physical activity as well as amount of smoking (number of cigarettes per day) were assessed. During the (physical) examinations the participants were screened for health status by a general practitioner (standardized questionnaire with questions regarding (severe) diseases and (minor) complications) in order to screen out the existence of any of the exclusion criteria. Body height and weight were determined by trained personnel. The body weight of participants was measured without shoes and with light clothes using a calibrated SECA scale and recorded to the nearest 0.1 kg; their body height without shoes was determined with a tape measure and recorded to the nearest centimeter. BMI was calculated from weight (kg) divided by height squared (m²) of each subject.

Definition of ‘Vegan’

As previously described, participants were classified as vegan when adhering to either a strict vegan diet (SV, n = 98) or a moderate vegan diet (MV, n = 56). The latter was considered a vegan diet when a maximum of 5% of the ingested energy from eggs, milk and/or dairy products was included [30]. On the one hand, the ingested amounts of eggs (0.88 ± 3.31 g/day), milk and dairy products (7.95 ± 10.1 g/day) in the moderate vegan subgroup was so low that it seemed not to be appropriate to name them restricted lacto-ovo-vegetarians. On the other hand, although all persons reported to be vegans, some persons included food of animal origin in their diet. Moreover, the consumption of eggs, milk and dairy products makes it difficult – if not impossible at all – to name those persons (strict) vegans. Therefore, we decided to name them moderate vegans rather than restricted lacto-ovo-vegetarians or strict vegans.

Dietary Data Collection

A slight modification of the validated FFQ used in the Giessen Raw-Food Study [36] was used in the German Vegan Study (GVS) for data collection. The original FFQ was complemented for vegan foods; foods of animal origin – except eggs, butter, milk and dairy products – were excluded. The GVS-FFQ included 199 vegan foods and 7 non-vegan foods such as dairy products and eggs. In the GVS-FFQ, common household measures and their equivalents in grams or milliliters were given for each food item. In some cases, portion sizes were made clear with photos or specific comments (i.e. nine strawberries with a size of walnuts weigh 150 g; an orange of the size of a tennis ball equals 250 g). In addition, participants were asked for copies of recipes of homemade vegan dishes.
Each study participant was asked to complete two estimated 9-day FFQs. In order to minimize seasonal differences, one FFQ was completed in autumn, the other one in spring. As there was no adequate software or database for vegan nutrition to calculate the amounts of ingested nutrients, we developed a piece of software (Paradox database) on the basis of the German nutrient database (BLS II.2). Missing nutritional information of vegan foods and dishes were obtained from the food manufacturers. In the end, 245 foods (incl. 18 beverages) were listed in the database.

Computing the Dietary GI and GL

For the present analysis, most values of GI (GI = 100 corresponding to glucose) and GL of the foods were taken from the ‘International Table of Glycemic Index and Load’ by Foster-Powell et al. [37]. GIs for some foods, for which no published data were available, were estimated using the mean of published indexes from other, similar foods as suggested by Buyken et al. [38]. In these cases, the GI of each food was computed by multiplying the food’s GI and the carbohydrate content of one serving. The overall dietary GI for each participant was calculated by multiplying the GI per serving of food by the frequency of consumption and summed these products over all food items to produce the dietary GI. Therefore the formula for the calculation of overall GI is as follows [14]:

\[
\text{Overall dietary GI} = \sum_{i=1}^{n} \left( \text{GI}_i \times \text{CHO}_i \times \text{FPD}_i \right)
\]

where \( \text{GI}_i \) is the GI for food \( i \), \( \text{CHO}_i \) is the carbohydrate content in food; (in grams per serving), and \( \text{FPD}_i \) is the frequency of servings of food \( i \) per day.

The GL density was computed by dividing the dietary GI by energy intake. Additionally, the average GI of the individual’s diet, an indicator of the overall quality of carbohydrate intake, was determined. According to Liu et al. [14] the formula for computing the overall dietary GI is the following:

\[
\text{Overall dietary GI} = \frac{\sum_{i=1}^{n} \left( \text{GI}_i \times \text{CHO}_i \times \text{FPD}_i \right)}{\sum_{i=1}^{n} \text{CHO}_i \times \text{FPD}_i}
\]

Evaluation of Lipid Parameters

Fasting venous blood samples were taken for the measurement of lipid parameters. TG concentrations were measured using an enzymatic colorimetric test (GPO-PAP; Boehringer, Germany). Total cholesterol was assessed also by an enzymatic colorimetric test (CHOD-PAP; Boehringer), while LDL concentrations were measured by precipitation with polyvinylsulfate (Boehringer) and HDL cholesterol by precipitation with phosphotungstic acid (Boehringer).

Statistical Methods

A statistical analysis program (SPSS 12.0; Chicago, Ill., USA) was used to analyze the data. The results are shown as mean ± SD and median plus 5th/95th percentiles, respectively. The following two-tailed tests at the 5% level of significance were employed to evaluate the data: in case of skewness the Mann-Whitney U test was used to evaluate differences between the two subgroups (SV and MV). Given normal distribution, the independent sample t test was taken. Normal distribution of data was checked visually and by using the Kolmogorov-Smirnov test. Dealing with nominal data, the \( \chi^2 \) test was employed to evaluate statistically significant differences. For the subgroups characterized by the quartiles of average dietary GL, GI density, average dietary GI as well as intake of carbohydrate and dietary fiber, the Kruskal-Wallis test was carried out to reveal statistically significant differences regarding the demographic, life course, and biochemical factors. Using multiple linear regressions the plasma concentrations of TG, total cholesterol, HDL and LDL cholesterol were adjusted for potential confounding factors including age, BMI, cigarette smoking, alcohol intake, physical activity, intakes of energy, dietary fiber, protein, saturated, monounsaturated, and polyunsaturated fatty acids. The adjusted geometric means were calculated by first regressing the natural logarithm of plasma concentrations on dietary variables and by exponentiation of the resulting mean concentrations.

Because of the low variability in BMI (5th percentile: 17.5 kg/m², 95th percentile: 25.4 kg/m²), analyses were not repeated with subjects stratified by BMI classes.

Results

Of the recruited subjects, 56.5% were females (SV 51.0%, MV 66.1%, \( p = 0.07, \chi^2 \)). Furthermore, the two subgroups did not differ in their mean BMI and their mean age. Participants reported a total carbohydrate intake of 274 ± 90.3 g/day, which contributed 57.1 ± 7.48% to dietary energy. Computed dietary fiber intake was 56.7 ± 7.74 g/day. The average dietary GI was 144 ± 47.3, the mean GL density was 17.7 ± 3.06 and the average dietary GI was 51.4 ± 4.25. Table 1 shows the corresponding data for the dietary subgroups SV and MV. Strict vegans had significantly higher dietary intakes of polyunsaturated fatty acids (PUFA) and n–6 fatty acids respectively, whereas the intake of cholesterol was significantly lower.

Dietary GI and GL

Table 2 shows to which extent different food groups contributed to the overall dietary GI. In the total study population, whole grain products and fruits were the main contributors to the GL with a median contribution of 37.8 and 26.3%, respectively. There were no significant differences in age, exercise time, smoking status, and alcohol among the dietary quartiles of GI (neither when the crude GI was used nor when controlling for energy). But among the dietary GLD quartile groups, age distribution was significantly different. In the lowest GLD quartile, persons were older (47.8 ± 15.1 years) than in the highest quartile (40.6 ± 16.1, \( p = 0.018 \)).

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Lipid Profile

The unadjusted mean fasting TG concentration of the participants was 0.96 ± 0.72 mmol/l (SV 0.96 ± 0.80 mmol/l, MV 0.95 ± 0.55 mmol/l; p = 0.724). The unadjusted mean plasma total cholesterol concentration was 4.50 ± 1.08 mmol/l (SV 4.36 ± 0.97 mmol/l, MV 4.75 ± 1.22 mmol/l; p = 0.044). The unadjusted mean plasma HDL cholesterol concentration was 1.34 ± 0.34 mmol/l (SV 1.30 ± 0.33 mmol/l, MV 1.41 ± 0.34 mmol/l; p = 0.070). The unadjusted mean plasma LDL cholesterol concentration was 2.63 ± 0.98 mmol/l (SV 2.53 ± 0.90 mmol/l, MV 2.81 ± 1.09 mmol/l; p = 0.108). The unadjusted ratio of total cholesterol to HDL was 3.49 ± 0.97 (SV 3.49 ± 1.02, MV 3.48 ± 0.89; p = 0.956). An association between the intake of saturated fatty acids and the unadjusted concentrations of total, HDL, and LDL cholesterol was found (Spearman correlation coefficient each r < 0.3, p < 0.5). Therefore, the multivariate models were – besides other factors – adjusted for the intake of saturated, monounsaturated, and polyunsaturated fatty acid intake.
The calculated geometric mean concentrations of TG, total cholesterol, HDL, and LDL cholesterol by quartiles of dietary GL, GL density, average dietary GI, carbohydrate, and dietary fiber intake are shown in Table 3.

The adjusted geometric mean plasma concentrations of total cholesterol, HDL, and LDL cholesterol decreased across the quartiles of dietary GL, carbohydrate and dietary fiber intake. While the associations with GL density and average GI, respectively, were inconsistent. For the lowest and the highest quartiles of GL, the multivariate-adjusted geometric mean total cholesterol concentrations were 4.92 and 3.97 mmol/l, whereas the geometric mean HDL cholesterol concentrations were 1.44 and 1.15 mmol/l and the geometric mean LDL cholesterol concentrations were 2.87 and 2.18 mmol/l, respectively.
lowest and the highest quartiles of carbohydrate intake the multivariate-adjusted geometric mean total cholesterol concentrations were 4.93 and 3.97 mmol/l, whereas the geometric mean HDL cholesterol concentrations were 1.44 and 1.15 mmol/l and the geometric mean LDL cholesterol concentrations were 2.90 and 2.18 mmol/l, respectively. For the lowest and the highest quartiles of dietary fiber intake the multivariate-adjusted geometric mean total cholesterol concentrations were 4.88 and 3.99 mmol/l, whereas the geometric mean HDL cholesterol concentrations were 1.41 and 1.17 mmol/l and the geometric mean LDL cholesterol concentrations were 2.85 and 2.19 mmol/l, respectively.

However, statistically significant associations or trends were not observed for GI, GL, the intake of carbohydrates, and plasma TG concentrations.

**Discussion**

In the present study conducted with adult vegans from Germany it was assumed that a vegan diet is characterized by a low GI and GL. The average dietary GI of the GVS study population was 51. Compared to the results of several observational studies [10–18], the overall GI of the vegan diet seems to be low and, therefore, may be one factor to explain the lower risk for CHD and type 2 diabetes of vegan subjects in comparison to omnivores [25]. However, it is difficult to accurately compare the mean GIs in the different studies, because different GI values (glucose- or white bread-based) are used. While we have taken glucose-based values, eight of the observational studies cited above [10–18] either did not indicate which one they used or stated they used white bread-based values. Only the study of Amano et al. [15] used glucose-based values as we did in our study. When taking the conversion factor of approximately ~0.7 into account to convert white bread- into glucose-based GI values [37], the average GI of the GVS study falls within the lowest GI quintile of the most observational studies. For example, in the Nurses’ Health Study (NHS) with 65,173 participants, the lowest GI quintile was 64 and this was associated with the lowest risk for diabetes mellitus type 2 [10]. Similar results were observed in a cohort study with 42,759 men, who were 40–75 years old. In this study, the dietary GI of the lowest quintile was 65 [16]. After adjustment for age, BMI, smoking, physical activity, family history of diabetes, alcohol consumption, cereal fiber, and total energy intake, the relative risk for diabetes mellitus type 2 was 37% lower in the lowest GI quintile compared with the highest quintile of GI [16]. The cohort study of Schulze et al. [18] supports our assumption that the GI of a vegan diet is rather low. In this large study with 91,249 women the lowest GI quintile was 70. This quintile was associated with a 60% lower multivariate relative risk for diabetes mellitus type 2. In addition in the cohort study by Liu et al. [12], using data from 10 years of follow-up on 75,000 women, the GI of the lowest quintile was 72.

The average dietary GL of the GVS cohort was 144. In comparison to the findings from observational studies with omnivores [12, 14, 15, 39] the reported GL with the vegan diet was low to moderate. For example, in the total cohort of the NHS the mean value for the lowest and highest quintile of GL was 117 and 206, respectively. In this study the GL and the risk of CHD were positively associated – even after multiple adjustments were made [12].

Since there are associations between higher dietary GI or GL and risk for CHD disease [11–15, 39, 40] the relations of dietary GI and GL to plasma concentrations of lipoproteins such as total cholesterol, HDL, LDL cholesterol, the ratio of TC:HDL, and TG were examined. Contrary to the results of several observational studies [11, 14, 15, 40], there were no significant associations between TG, total cholesterol, HDL, LDL, TC:HDL, and the average GI in the GVS study population. This observation may be explained by the fact that there is only a small difference between the lowest (Q1: 46) and highest (Q4: 57) quartile of dietary GI in our vegan study population. Therefore, all quartiles are characterized by a low GI when compared to the studies of Frost et al. [11], Liu et al. [14], and Amano et al. [15]. For this reason, the GVS results are not surprising, since all vegan subjects consumed a low-GI diet. In this context a recent meta-analysis is of interest. In this analysis of 14 randomized controlled trials of low-GI versus high-GI diets, low-GI diets caused a statistically significant improvement in total cholesterol concentrations, while there were no significant effects on LDL, HDL cholesterol and TG [41]. This indicates that the influence of the overall dietary GI on markers of lipid metabolism still remains controversial.

In our study the adjusted geometric mean plasma concentrations of cholesterol, HDL and LDL cholesterol decreased across the quartiles of dietary GI while there was no significant correlation between GI and TG. Except for HDL cholesterol, this result is in contrast to the findings of other investigators [13–15, 40]. For example, in the NHS cohort of Liu et al. [14] a significant rise of TG was observed with increasing GI. Furthermore, GL remained a significant predictor of fasting TG concentrations. Be-
cause GL is defined as the product of GI and carbohydrate content [10, 16], carbohydrate intake of the GVS cohort is of interest. With a mean intake of 274 g/day (57.1 energy %) the diet of the subjects in our study is comparatively high in carbohydrates. In observational studies like the postmenopausal female subcohort of the NHS, ascending quintiles of carbohydrate intake were associated with increasing TG concentrations [14]. In addition, total carbohydrate intake was inversely associated with HDL cholesterol concentrations [40].

Data from metabolic studies support the observation that high-carbohydrate diets increase TG or reduce HDL cholesterol and therefore could translate into a higher risk of CHD [42, 43]. However, our results and those of Goff et al. [35] show that vegan diets rich in carbohydrates are not or only partially associated with such negative effects on the lipid profile. Other studies have shown that the HDL cholesterol reduction and rise of TG seen with some high-carbohydrate diets may be short-term effects [44, 45] and depends mainly on the carbohydrate quality of such diets. For example, Harbis et al. [46] studied two mixed meals that were reasonably matched for macronutrient composition but differed in their content of slowly available glucose (flaké-based versus biscuit-based meal). The plasma TG and the apolipoprotein B-100 and B-48 concentration increased significantly after the flakes meal (high GI) but not after the biscuit meal (low GI) in obese, insulin-resistant subjects. Similar results were observed in healthy individuals with different GI meals [47]. A recently published randomized trial of overweight or obese adults with dyslipidemia or fasting hyperglycemia shows no significant undesirable effects of a high-carbohydrate, low-GI diet (Ornish) on TG and HDL cholesterol when compared with a low-carbohydrate diet (Atkins) [48]. Furthermore, a reduction in the overall dietary GI by at least 12 units reduces the TG concentration by approximately 9% in 10 of 11 long-term studies lasting from 1 week to 3 months [49]. Other investigators have shown that a high-carbohydrate, but low-GI diet significantly increased HDL cholesterol concentrations compared to an isocaloric high-carbohydrate diet with a high GI [50].

The exact mechanisms related to lipid modulation by low-GI diets still remain poorly understood. It is known that the glycemic and insulimic response after consumption of a low-GI meal is relatively slow compared to a high-GI meal [51–53]. The resulting comparatively low insulin concentration after a low-GI meal may reduce the insulin-stimulated HMG-CoA reductase activity and thus may reduce cholesterol synthesis [24]. Another possible mechanism is that the low insulin response could modulate VLDL synthesis in a favorable way [46].

Because the low-GI vegan diet of our subjects was based on minimally processed plant foods like fresh fruits, vegetables, whole grain products, legumes, and nuts [30], the low GI observed in this study is only one aspect of such a carbohydrate-rich diet. Concerning the risk factors of CHD such as plasma lipids (LDL, HDL cholesterol), another important aspect is the high fiber intake (56.7 g/day) observed in our study. In the GVS study population the adjusted geometric mean plasma concentrations of total cholesterol, HDL and LDL cholesterol decreased across the quartiles of dietary fiber intake. Except for HDL cholesterol, these findings are consistent with the results of several observational as well as clinical studies which have shown favorable effects of fiber-rich diets on the lipid profile. Especially soluble fiber abundant in fruits, legumes, and oat cereals exhibit LDL cholesterol-lowering properties and reduce TG levels [54]. A meta-analysis from 10 prospective cohort studies supported the CHD protective effect of a fiber-rich vegan diet. In this analysis dietary fiber from cereals and fruits was inversely associated with the risk of CHD [55]. Of course, there are a lot of other bioactive components in a vegan diet that alter plasma lipoproteins and TG. For example, phytosterols, monounsaturated fatty acids and vegetable protein in nuts have LDL cholesterol-lowering effects [56, 57].

Furthermore, the vegan diet in our study collective was characterized by a low intake of saturated fatty acids (6.0 ± 1.7 energy %) and dietary cholesterol (21.5 ± 27.0 mg/day), which may have a favorable impact on lipid profile. The multivariate analysis of the data allowed for considering some of these confounders by including them in the analysis.

Taking our data together with a series of clinical studies conducted from the Jenkins group [58–60] and the Ornish group [61–64] with high-fiber, carbohydrate-rich and low-GI diets, our results support the assumption that carbohydrate quality, not quantity is primarily important for lipid markers and CHD risk reduction. From the background of the current low-carbohydrate debate, our results, therefore, may be of interest for public health.

The most important limitation that should be mentioned when examining the results of this study is the use of a FFQ, which was not specifically designed to assess GI and GL. But this limitation occurs in some other studies as well [18]. Furthermore, the overall accuracy of the measurement of food intake via FFQ in general [65] and particularly the validity in view of measures the GI and

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GL of the diet [66] have been questioned. Because FFQs have substantial limitations [67], the above-mentioned questions arise also when interpreting the results of this study. However, up to now, the FFQ is the standard tool for dietary assessment in all large epidemiological studies and the results of those studies are still relevant [68, 69]. Another limitation is the measurement of surrogate markers (i.e. lipid parameters) instead of CHD endpoints such as CHD-related mortality or cardiac events. Also other markers (i.e. lipid parameters) instead of CHD endpoints such as CHD-related mortality or cardiac events. Another limitation is the measurement of surrogate markers (i.e. lipid parameters) instead of CHD endpoints such as CHD-related mortality or cardiac events. Also the relatively small number of vegan persons in this study might weaken the power of our results. To our knowledge, publications from the British EPIC cohort (e.g. Dav-ey et al. [70]) are the first data source with a vegan sample size of more than 250 persons, while other studies on vegetarian and vegan cohorts have smaller sample sizes too – not at least due to the small number of vegans found in a population.

Conclusion

A vegan diet including large amounts of fruit, vegetables, whole grains, legumes, and nuts is characterized by a low GI and GL. Especially the latter is associated with decreased total and LDL cholesterol plasma concentrations which are expected to be cardioprotective in the long run. Overall, these data do not support the hypothesis that a carbohydrate-rich diet per se is associated with unfavorable effects on TG that would be predicted to increase the risk of CHD.

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