



Article

# Pasta Consumption and Connected Dietary Habits: Associations with Glucose Control, Adiposity Measures, and Cardiovascular Risk Factors in People with Type 2 Diabetes—TOSCA.IT Study

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**Abstract:** Background: Pasta is a refined carbohydrate with a low glycemic index. Whether pasta shares the metabolic advantages of other low glycemic index foods has not really been investigated. The aim of this study is to document, in people with type-2 diabetes, the consumption of pasta, the connected dietary habits, and the association with glucose control, measures of adiposity, and

major cardiovascular risk factors. **Methods:** We studied 2562 participants. The dietary habits were assessed with the European Prospective Investigation into Cancer and Nutrition (EPIC) questionnaire. Sex-specific quartiles of pasta consumption were created in order to explore the study aims. **Results:** A higher pasta consumption was associated with a lower intake of proteins, total and saturated fat, cholesterol, added sugar, and fiber. Glucose control, body mass index, prevalence of obesity, and visceral obesity were not significantly different across the quartiles of pasta intake. No relation was found with LDL cholesterol and triglycerides, but there was an inverse relation with HDL-cholesterol. Systolic blood pressure increased with pasta consumption; but this relation was not confirmed after correction for confounders. **Conclusions:** In people with type-2 diabetes, the consumption of pasta, within the limits recommended for total carbohydrates intake, is not associated with worsening of glucose control, measures of adiposity, and major cardiovascular risk factors.

**Keywords:** type 2 diabetes; pasta consumption; dietary habits; glucose control; body mass index; cardiovascular risk factors

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## 1. Introduction

Type-2 diabetes mellitus is a condition primarily defined by the level of hyperglycemia, and it is associated with a reduced life expectancy, significant morbidity due to specific diabetes related microvascular complications, increased risk of macrovascular complications (ischemic heart disease, stroke, and peripheral vascular disease), and diminished quality of life. It is a chronic condition requiring continuous medical care with multifactorial risk-reduction strategies beyond glycemic control. Lifestyle management is a fundamental aspect of diabetes care and includes diabetes self-management education, nutrition therapy, physical activity, smoking cessation counseling, and psychosocial care.

Regarding nutrition therapy, the main goal is to improve glucose control and the cardiovascular risk factors profile, and to reduce weight, when needed [1]. Nutritional recommendations have been issued by several scientific societies in order to support clinicians in the management of diabetes [2–4]. Although there is no full agreement about the optimal composition of the diet, cereal foods—particularly those with a high glycemic index—are generally restricted because of their adverse effects on post-prandial plasma glucose and lipids [4,5]. Cereal foods are a major source of energy and are largely consumed worldwide under different forms; but there is remarkably little information on the health impact of the specific food items consumed in different cultures. Pasta, a traditional Italian dish now largely available around the world, is an important example of a food that is considered to be a refined carbohydrate, but has a low–moderate glycemic index (GI) and a low caloric density, [6–8], and therefore may be a more suitable source of carbohydrates than other cereal foods for people with type-2 diabetes. Recently, the consumption of pasta has considerably declined, as the concept of a low carbohydrate and high protein diet against obesity emerged. The scientific evidence linking pasta consumption with obesity/overweight is however scarce. A recent meta-analysis of randomized controlled trials has shown that in people with or without diabetes, pasta within the context of low-GI dietary patterns does not adversely affect adiposity; on the contrary, it slightly reduces body weight as compared with higher-GI dietary patterns [9]. These findings are in line with the results of observational studies showing benefits of pasta consumption on body weight and markers of adiposity [10,11]. In one of these studies, a greater pasta consumption was also associated with a better adherence to the Mediterranean diet [10]—a dietary pattern with proven beneficial effects on type-2 diabetes and cardiovascular risk [12].

As for the other relevant effects, a systematic review on the relation between pasta consumption and cardio-metabolic risk factors convincingly shows that pasta meals induce lower postprandial glucose responses compared with other starch-dense foods, like white bread and potatoes, commonly consumed in Western diets [13]. Most of the studies reviewed, however, are of a relatively short

duration, and seldom include measurements of major cardiovascular risk factors, so that no conclusions on the effect of pasta consumption on major cardiovascular risk factors can be drawn. It remains therefore unclear to date whether, and to what extent, pasta shares the cardiometabolic advantages of other low GI foods in people with diabetes.

Furthermore, foods are consumed based on an exchange list principle (i.e., a larger consumption of some foods is often counterbalanced by a lesser consumption of other foods), and a greater consumption of pasta may be associated with a lower intake of energy-dense foods, rich in animal fat, which may adversely affect body weight, plasma lipids, and in turn cardiovascular health. Finally, as adherence to the nutritional recommendations in people with diabetes is generally low [14,15], the inclusion in the dietary plan of local, traditional dishes to be shared with other members of the community may considerably improve adherence.

The aim of this study is to document, in clinical practice, the consumption of pasta and the connected dietary habits in a cohort of people with type-2 diabetes, and to evaluate the association of pasta consumption with glucose control, measures of adiposity, and major cardiovascular risk factors.

## 2. Materials and Methods

We studied 2562 men and women with type-2 diabetes, aged 5–75 years, enrolled in the TOSCA.IT study, a randomized clinical trial (clinicaltrials.gov NCT00700856) designed to compare the impact of glucose-lowering drugs on cardiovascular events [16]. The study protocol was approved by the Ethics Review Committee of the coordinating center and of each participating center; written informed consent was obtained from all participants. People with cardiovascular events in the previous six months, cancer in the previous five years, or any disease, other than diabetes, requiring special dietary treatment(s), were excluded from the study. The details of the study protocol and the main results of the trial have been previously published [17]. For the purposes of the present analysis, we used data collected at study entry, prior to randomization to the study treatments. The data were analyzed according to a cross-sectional observational design.

The body weight and height, waist, and hip circumference were measured according to a standard protocol. The body mass index (BMI) was calculated as weight (kg)/height (m<sup>2</sup>). The percentage of people with abdominal obesity was also calculated according to the ATP III criteria (waist circumference  $\geq 102$  cm for men and  $\geq 88$  cm for women). Biochemical analyses were performed in a central laboratory on fasting blood samples. HbA1c was measured with high-performance liquid chromatography (HPLC). The total cholesterol, HDL-cholesterol, and triglycerides were measured by standard methods. The LDL-cholesterol was calculated according to the Friedewald equation, only for people with triglycerides values  $< 400$  mg/dL. For people with triglyceride values  $> 400$  mg/dL (seven in total), the LDL cholesterol was considered as a missing value. By protocol, all patients were taking the same treatment for diabetes (i.e., metformin 2 g/day) [17]. Information on the concomitant treatments were collected by standard questionnaires.

### 2.1. Collection of Dietary Information

Dietary habits were investigated with the European Prospective Investigation into Cancer and Nutrition (EPIC) food frequency questionnaire, adapted for the Italian population [18,19]. The questionnaire contains 248 items, including the type of fat used as a condiment or added after cooking. The respondent indicates the absolute frequency of consumption of each item (per day, week, month, or year). The quantity of foods consumed was assessed based on the respondent's selection of an image of a food portion showing a small, medium, or large portion size, with additional quantifiers (i.e., "smaller than the small portion" or "between the small and medium portion"). Software developed ad hoc by the Epidemiology and Prevention Unit, Fondazione IRCCS, Istituto Nazionale dei Tumori, Milan, was used to convert the questionnaire dietary data into frequencies of consumption (times per day) and average daily amounts of foods (grams per day). The energy intake (kcal per day) and nutrient composition of the habitual diet were estimated by the software,

through the link of the consumed foods to the Italian Food Composition Tables [20,21] for energy and nutrients assessment. The glycemic index was calculated according to the tables by Brighenti [7], and the International Table by Foster and Powell [8]. Incomplete questionnaires and questionnaires with implausible data were excluded from the analyses. The intake of macronutrients is given as a percentage of the total energy or g/1000 Kcal/day, or an absolute value, as appropriate.

## 2.2. Statistical Analysis

The data are expressed as means and standard deviations, or frequencies and percentages, as appropriate. To correct for the different energy intake between subjects, the intake of pasta was expressed as g/1000 Kcal/day. The relation of pasta consumption with anthropometric variables, blood pressure, glucose control, and plasma lipids, was explored by stratifying the study population according to quartiles of pasta consumption; in order to account for gender differences in pasta consumption, allocation into quartiles was based on the sex-specific distribution of the amount consumed. To compare variables across the quartiles, an analysis of variance with a test for a linear trend was used for the continuous variables, and the  $\chi^2$  test was used to compare proportions. The multivariate linear regression model was used to evaluate the separate and combined association of pasta and other covariates with the variables of interest. Covariates were age, gender, BMI, smoking, fiber, and alcohol intake. The statistical analyses were performed with SPSS version 19.0. Statistical significance was assessed as  $p < 0.05$  (two-sided).

## 3. Results

The study population consisted of 1531 men and 1031 women with type-2 diabetes. The average age was  $62.1 \pm 6.5$  years and the average BMI was  $30.3 \pm 4.4$  Kg/m<sup>2</sup>. The average pasta consumption was  $53.0 \pm 40.4$  g/day, and was higher in men than in women ( $p < 0.05$ ). The clinical and socio-demographic characteristics of the study population are given in Table 1, according to quartiles of pasta consumption. Age, diabetes duration, smoking habits, educational status, and geographical area of residence were not significantly different across the categories of pasta consumption.

**Table 1.** Characteristics of the study population by quartiles of pasta consumption expressed as (g/1000 Kcal).

	Quartile 1 (n = 640)	Quartile 2 (n = 641)	Quartile 3 (n = 641)	Quartile 4 (n = 640)	p-Value
Pasta consumption (g/1000 Kcal)	7.5 ± 4.4 (0.0–15.8)	19.4 ± 4.2 (15.9–27.4)	30.8 ± 5.0 (27.5–41.1)	52.4 ± 14.0 (41.2–120.0)	
Age (years)	62.0 ± 6.5	62.3 ± 6.5	62.0 ± 6.5	62.1 ± 6.4	0.842
Diabetes Duration (years)	8.5 ± 5.8	8.3 ± 5.4	8.7 ± 6.0	8.5 ± 5.6	0.658
Smoking status (%) <sup>a</sup>					
Current Smokers	125 (19.5)	114 (17.8)	97 (15.1)	106 (16.6)	0.339
Educational status (%)					
None	194 (30.4)	197 (30.7)	180 (28.1)	218 (34.1)	0.204
Primary	226 (35.4)	248 (38.6)	250 (39.0)	231 (36.1)	
Secondary	185 (29.0)	160 (24.9)	174 (27.1)	169 (26.4)	
University	34 (5.3)	36 (5.8)	37 (5.8)	22 (3.4)	
Geographical area (%)					
North	224 (34.9)	216 (33.8)	218 (34.0)	233 (36.4)	0.681
Center	156 (24.6)	182 (28.3)	176 (27.5)	159 (24.8)	
South	260 (40.5)	243 (37.9)	247 (38.5)	248 (38.8)	

<sup>a</sup> Subjects were classified as “current smokers” if they smoked five or more cigarettes/day.

Table 2 describes the composition of the diet. The average daily energy intake was similar across the quartiles of pasta consumption, but the nutrient composition of the diet was significantly different.

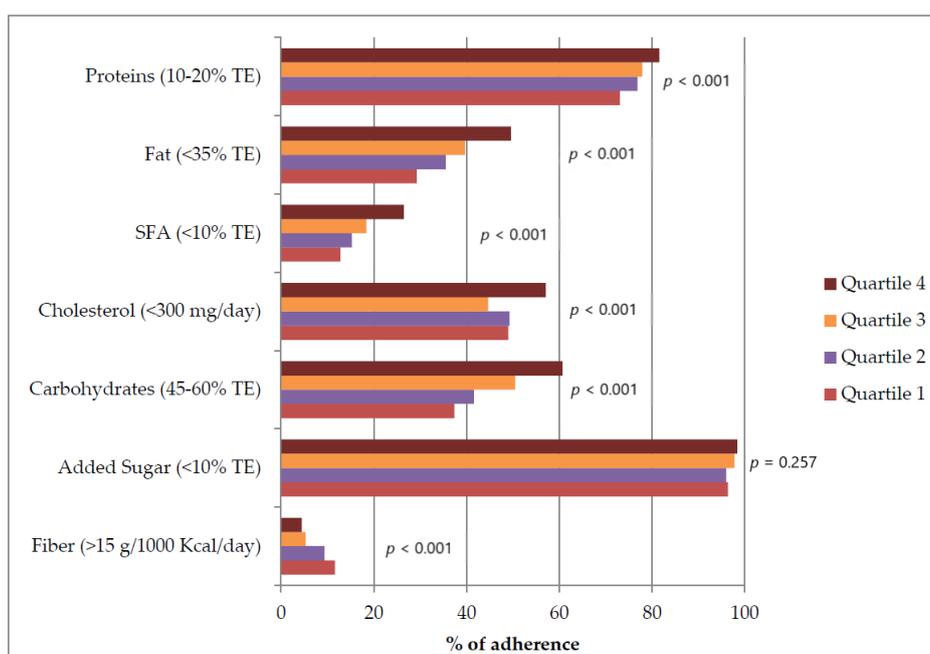
A higher pasta consumption was associated with a significantly higher intake of total carbohydrates, significantly lower intake of proteins, total fat, saturated fat, unsaturated fat, cholesterol, added sugar, alcohol, and fiber. The glycemic load of the diet was not significantly different across quartiles of pasta consumption, whereas the glycemic index was inversely associated with pasta consumption.

**Table 2.** Total energy and macronutrient intake in the study population by quartiles of pasta consumption (g/1000 Kcal).

	Quartile 1 (n = 640)	Quartile 2 (n = 641)	Quartile 3 (n = 641)	Quartile 4 (n = 640)	p for Anova	p for Trend
Pasta (g/1000 Kcal)	7.5 ± 4.4 (0.0–15.8)	19.4 ± 4.2 (15.9–27.4)	30.8 ± 5.0 (27.5–41.1)	52.4 ± 14.0 (41.2–120.0)		
Energy (Kcal)	1841 ± 820	1962 ± 881	1941 ± 683	1852 ± 673	0.067	0.519
Protein (% of TE)	18.6 ± 2.8	18.3 ± 2.5	18.3 ± 2.4	17.9 ± 2.5	<0.0001	<0.0001
Total Fat (% of TE)	38.3 ± 6.4	37.1 ± 6.1	36.3 ± 5.8	35.1 ± 5.7	<0.0001	<0.0001
SFA (% of TE)	12.8 ± 2.7	12.4 ± 2.5	12.1 ± 2.4	11.4 ± 2.3	<0.0001	<0.0001
MUFA (% of TE)	18.4 ± 4.1	18.0 ± 3.9	17.8 ± 3.6	17.5 ± 3.7	0.002	<0.0001
PUFA (% of TE)	4.7 ± 1.3	4.5 ± 1.0	4.3 ± 1.0	4.2 ± 1.0	<0.0001	<0.0001
Cholesterol (mg/1000 Kcal)	191 ± 54	183 ± 54	180 ± 50	172 ± 49	<0.0001	<0.0001
Carbohydrates (% of TE)	43.1 ± 7.7	44.6 ± 7.5	45.4 ± 7.0	46.9 ± 6.8	<0.0001	<0.0001
Added Sugar (% of TE)	2.7 ± 3.7	2.4 ± 3.3	2.3 ± 3.2	1.9 ± 2.5	<0.0001	<0.0001
Fiber (g/1000 Kcal)	11.5 ± 3.0	11.0 ± 2.9	10.5 ± 2.6	10.1 ± 2.4	<0.0001	<0.0001
Glycemic index (%)	53.0 ± 3.6	52.4 ± 3.2	51.5 ± 2.9	49.7 ± 2.9	<0.0001	<0.0001
Glycemic load	111 ± 64	118 ± 65	119 ± 50	111 ± 46	0.078	0.341
Alcohol (g/day)	10.7 ± 15.9	12.2 ± 18.4	11.2 ± 14.6	9.1 ± 13.8	0.046	0.023

TE—total energy; SFA—saturated fatty acids; MUFA—monounsaturated fatty acids; PUFA—polyunsaturated fatty acids.

A higher pasta consumption was also associated with a better adherence to the nutritional recommendations for people with diabetes for all items except for fiber, which was inversely associated with pasta consumption (Figure 1).



**Figure 1.** Adherence to the nutritional recommendations [2,3] by sex-specific quartiles of pasta consumption, expressed as g/1000 Kcal. TE—total energy; SFA—saturated fatty acids.

Glucose control, evaluated by glycated hemoglobin, did not change significantly across the quartiles of pasta consumption (Table 3). In this regard, it is relevant to say that because of the eligibility criteria of the trial, the study participants were all taking the same hypoglycemic treatment (i.e., metformin 2 g/day). No relation was observed between pasta consumption and measures of adiposity. BMI, prevalence of obesity (BMI  $\geq 30$  Kg/m<sup>2</sup>), and surrogate measures of visceral adiposity—such as waist circumference, waist/hip ratio, and prevalence of abdominal obesity—were not significantly different across the categories of pasta consumption (Table 3). As for major cardiovascular risk factors, non-significant differences were observed for LDL-cholesterol and triglycerides, whereas HDL-cholesterol was significantly and inversely associated with pasta consumption. The LDL to HDL ratio and the non-HDL-cholesterol, two markers of the atherogenicity of the plasma lipids profile, remained stable across the quartile of pasta consumption. Systolic blood pressure was significantly and directly associated with pasta consumption, whereas no difference was detected for diastolic blood pressure. A high proportion of the cohort was taking lipid lowering medications (61.2%) and antihypertensive medications (92.3%), with no significant differences across categories of pasta consumption (Table 3).

**Table 3.** Clinical characteristics of the study population by quartiles of pasta consumption expressed as g/1000 Kcal.

	Quartile 1 (n = 640)	Quartile 2 (n = 641)	Quartile 3 (n = 641)	Quartile 4 (n = 640)	p for Anova or $\chi^2$	p for Trend
Pasta (g/1000 Kcal)	7.5 $\pm$ 4.4 (0.0–15.8)	19.4 $\pm$ 4.2 (15.9–27.4)	30.8 $\pm$ 5.0 (27.5–41.1)	52.4 $\pm$ 14.0 (41.2–120.0)		
BMI (Kg/m <sup>2</sup> )	30.1 $\pm$ 4.4	30.5 $\pm$ 4.5	30.2 $\pm$ 4.5	30.4 $\pm$ 4.4	0.362	0.569
BMI $\geq 30$ Kg/m <sup>2</sup> (%)	48.6	49.5	48.0	49.7	0.926	
Waist Circumference (cm)	103.9 $\pm$ 11.2	104.8 $\pm$ 11.1	104.7 $\pm$ 11.2	104.4 $\pm$ 11.3	0.497	0.524
Waist-to-hip ratio	0.98 $\pm$ 0.07	0.98 $\pm$ 0.06	0.98 $\pm$ 0.07	0.98 $\pm$ 0.07	0.545	0.154
With Abdominal obesity * (%)	71.9	74.6	75.0	74.1	0.589	
HbA1c (%)	7.67 $\pm$ 0.51	7.70 $\pm$ 0.50	7.67 $\pm$ 0.51	7.68 $\pm$ 0.51	0.707	0.915
Proportion with HbA1c <7.5%	26.0	23.7	25.8	24.4	0.445	
LDL-cholesterol (mg/dl)	104.7 $\pm$ 32.3	102.3 $\pm$ 31.5	102.2 $\pm$ 31.6	102.0 $\pm$ 30.3	0.364	0.142
HDL-cholesterol (mg/dl)	47.8 $\pm$ 12.5	45.5 $\pm$ 11.8	46.1 $\pm$ 11.9	44.9 $\pm$ 11.3	<0.0001	<0.0001
LDL/HDL Ratio	2.32 $\pm$ 0.90	2.36 $\pm$ 0.90	2.33 $\pm$ 0.90	2.39 $\pm$ 0.92	0.571	0.271
Triglycerides (mg/dl)	145.7 $\pm$ 71.5	153.4 $\pm$ 80.2	148.7 $\pm$ 72.6	155.2 $\pm$ 75.4	0.095	0.072
Non-HDL Cholesterol (mg/dl)	133.8 $\pm$ 36.0	133.3 $\pm$ 36.3	132.9 $\pm$ 37.9	133.2 $\pm$ 35.0	0.980	0.963
Systolic blood pressure (mm/Hg)	132.5 $\pm$ 14.3	133.7 $\pm$ 13.9	135.2 $\pm$ 15.2	133.8 $\pm$ 15.0	0.034	0.012
Diastolic blood pressure (mm/Hg)	78.9 $\pm$ 8.8	79.8 $\pm$ 8.6	80.1 $\pm$ 8.3	79.4 $\pm$ 8.4	0.103	0.300
On lipid lowering medications (%)	59.7	66.1	62.8	59.9	0.080	
On antihypertensive medications (%)	93.6	90.3	94.1	92.0	0.104	

\* Waist circumference: >102 cm for men and >88 cm for women. BMI—body mass index.

To disentangle the association of pasta per se with HDL-cholesterol and systolic blood pressure, we performed a multivariate regression analysis with age, gender, BMI, smoking, fiber, and alcohol intake as covariates. Pasta consumption remained significantly and inversely associated with HDL-cholesterol, but not with systolic blood pressure (Table 4).

**Table 4.** Multivariate regression analyses of the relation between pasta consumption and HDL-cholesterol and systolic blood pressure.

	Standardized $\beta$ -Coefficient	Standard Error	p-Value
Dependent variable HDL-cholesterol			
Intake of pasta (g/1000 Kcal)	−0.057	0.007	0.003
Gender (female/male)	0.330	0.535	0.000
BMI (Kg/m <sup>2</sup> )	−0.170	0.050	0.000
Age (years)	0.093	0.034	0.000
Fiber intake (g/1000 Kcal)	0.004	0.032	0.836
Alcohol (g/day)	0.171	0.015	0.000
Smoking (yes/no)	−0.018	0.266	0.370

Table 4. Cont.

	Standardized $\beta$ -Coefficient	Standard Error	<i>p</i> -Value
Dependent variable Systolic Blood Pressure			
Intake of pasta (g/1000 Kcal)	0.031	0.010	0.133
Gender (female/ male)	−0.026	0.833	0.261
BMI (Kg/m <sup>2</sup> )	0.108	0.078	0.000
Age (years)	0.126	0.054	0.000
Fiber intake (g/1000 Kcal)	−0.045	0.049	0.024
Alcohol (g/day)	0.001	0.023	0.970
Smoking (yes/no)	0.044	0.415	0.040

#### 4. Discussion

The study provides evidence that in people with type-2 diabetes, the consumption of cereals in the form of pasta, not exceeding the limits recommended for total carbohydrate intake, is associated with a lower intake of proteins, total and saturated fat, cholesterol, and added sugar; this increases the overall adherence to the nutritional recommendation for the management of diabetes, which emphasizes the replacement of foods high in fat and sugar, with carbohydrates low in energy density and with a low glycemic index [1–3,22]. Furthermore, pasta consumption is not associated with a worsening of glucose control, BMI, obesity, abdominal adiposity, and the overall atherogenicity of the plasma lipids profile. However, a higher pasta consumption is associated with a lower intake of fiber, which makes the public health message regarding the consumption of pasta more complex than previously considered. Pasta is, in fact, a rare example of a refined carbohydrate food with a low GI.

The beneficial effects of low GI foods in people with and without diabetes have been established [23], but low GI diets can be achieved through different ways. The low GI of pasta is mainly due to its dense structure, resulting in slower digestion and delayed gastric emptying, which makes pasta a unique example of a refined cereal food with a low GI. To what extent pasta shares the positive metabolic effects of other low GI foods with a high fiber content remains unclear [24,25]. A recent meta-analysis convincingly shows that, in relatively short-term controlled studies, pasta meals induce lower postprandial glucose and possibly lower insulin response, compared with other carbohydrate-dense foods such as white bread or potatoes—widely consumed in Western diets [13]. The results of our study expand the current knowledge by showing that in people with type 2 diabetes, habitual pasta consumption, not exceeding the limits recommended for total carbohydrate intake is not associated with a worsening of glycated hemoglobin, a marker of long-term glucose control and a significant predictor of the development of diabetes complications [26]. Furthermore, we show that pasta consumption is not associated with a higher BMI, prevalence of obesity, or abdominal adiposity. These findings do not lend support to the commonly held paradigm that pasta is “fattening”. In contrast with this misconception and in line with our results, a recent meta-analysis of randomized controlled studies has shown that in people with or without diabetes, pasta consumption, within the context of a low GI eating pattern, is associated with a lower BMI compared with high GI eating patterns [9]. Along the same line are the results of two observational studies, which demonstrate the benefits of pasta consumption on body weight and adiposity in large, free-living populations [10,11].

Unlike prior studies, we have explored the associations between pasta consumption and major cardiovascular risk factors. An unwanted effect of carbohydrate intake on HDL-cholesterol has been described previously [27]. In line with these findings, we observe a significant inverse association of pasta consumption with HDL-cholesterol; although statistically significant, the magnitude of the change was of moderate clinical relevance. Furthermore, because of a concomitant, non-significant reduction in LDL-cholesterol, in this study, the ratio LDL to HDL cholesterol, and the values of non-HDL cholesterol did not change across the pasta quartiles. This is relevant in that non-HDL cholesterol represents a good estimation of the total amount of potentially atherogenic apolipoprotein B-containing lipid fractions (i.e., LDL, lipoprotein-a, intermediate-density lipoprotein, very low-density lipoprotein,

and VLDL remnants), and has a well-established role as an independent predictor of cardiovascular disease, even in patients with normal LDL and triglyceride levels [28]. Non-HDL-cholesterol is, in fact, recommended as a secondary lipid target when the LDL-cholesterol goal is reached, particularly in people with insulin resistance—a common feature of type-2 diabetes [29].

We found a direct association of pasta consumption with systolic blood pressure, which was, however, largely mediated by major confounders such as age and BMI. The influence of dietary carbohydrates on blood pressure is controversial—both the quality and quantity of carbohydrates may play a role [30]. To our knowledge, the relationship of pasta consumption with blood pressure has not been explored before. The present study is the first exploring, in real life clinical practice, the eating habits and quality of the diet associated with pasta consumption in people with type-2 diabetes, with a focus on the cardio-metabolic profile.

A major study strength is that the intake of food and nutrients reflects real life conditions. Furthermore, the use of a central laboratory for biochemical analyses ensures good quality standards for the measurements.

Among the limitations, we acknowledge the cross-sectional design, which may limit the assessment of a cause-effect relationship. Reverse causation is a concern when the relation of dietary habits with cardio-metabolic markers and adiposity measures is assessed cross-sectionally. The individual's perception of cardio-metabolic and weight status may influence both the dietary habits and their reporting, and may produce spurious associations. In order to limit the possibility of spurious associations, we have excluded from the analysis implausible dietary questionnaires and the people with acute cardiovascular events in the prior six months. Furthermore, we observed a significant and positive association between BMI and energy intake in the whole population, which supports the internal validity of the data. Additional problems are the lack of information on wholegrain cereal intake, their consumption, however, is rather limited; a previous study conducted in an Italian cohort has shown that the average consumption of wholegrain cereals is about 5 g per day [31]. In addition, the dietary data were collected only once, and could be prone to recall bias and seasonal variation. Finally, the data collected do not allow us to analyze the different types of pasta dishes. This may be relevant, as both healthy and unhealthy eating patterns have been described in pasta eaters [32], and this may depend on the way pasta dishes are prepared.

Regarding the study population, we cannot exclude that the use of hypoglycemic medications, as well as the extensive use of lipid lowering and anti-hypertensive drugs, may have partly masked a possible association of pasta with worsening of glucose control and cardiovascular risk factors. It is however relevant in this regard to underline that the use of medications was evenly distributed across the categories of pasta consumption.

Despite these, the study findings are relevant, in as much as the study population represents most of the patients with type-2 diabetes encountered in clinical practice.

## 5. Conclusions

In conclusion, in people with type-2 diabetes, the consumption of cereal foods in the form of pasta, not exceeding the limits recommended for total carbohydrate intake is not associated with the worsening of glucose control, BMI, adiposity measures, obesity, and major cardiovascular risk factors. If these results are confirmed, pasta may represent a suitable food choice to widen the dietary option in people with diabetes, particularly when consumed in dishes that combine pasta with vegetables and legumes, according to the Mediterranean traditional diet.

**Author Contributions:** O.V., G.R., A.A.R., M.M., and E.B. designed the study; A.C.B. (Anna Carla Babini), G.S., S.G., L.C., R.B., G.C., M.P.A.B., A.C.B. (Antonio Carlo Bossi), C.G., S.A., E.D., R.I., L.T., G.D.C., G.C., and G.G. conducted the research; S.G. and V.K. provided essential materials; M.V. and M.S. prepared the data base and performed the statistical analysis; M.V. and O.V. drafted the paper; M.V. and O.V. had primary responsibility for the final content. All authors have read and agreed to the published version of the manuscript.

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## References

1. American Diabetes Association. Lifestyle Management: Standards of Medical Care in Diabetes—2019. *Diabetes Care* **2019**, *42*, S46–S60. [[CrossRef](#)] [[PubMed](#)]
2. Standard Italiani per la Cura del Diabete Mellito 2018. Available online: <https://aemmedi.it/wp-content/uploads/2009/06/AMD-Standard-unico1.pdf> (accessed on 16 October 2019).
3. Mann, J.I.; De Leeuw, I.; Hermansen, K.; Karamanos, B.; Karlström, B.; Katsilambros, N.; Riccardi, G.; Rivellese, A.A.; Rizkalla, S.; Slama, G.; et al. Evidence-based nutritional approaches to the treatment and prevention of diabetes mellitus. *Nutr. Metab. Cardiovasc. Dis.* **2004**, *14*, 373–394. [[CrossRef](#)]
4. Ajala, O.; English, P.; Pinkney, J. Systematic review and meta-analysis of different dietary approaches to the management of type 2 diabetes. *Am. J. Clin. Nutr.* **2013**, *97*, 505–516. [[CrossRef](#)] [[PubMed](#)]
5. Kirk, J.K.; Graves, D.E.; Craven, T.E.; Lipkin, E.W.; Austin, M.; Margolis, K.L. Restricted-carbohydrate diets in patients with type 2 diabetes: A meta-analysis. *J. Am. Diet Assoc.* **2008**, *108*, 91–100. [[CrossRef](#)] [[PubMed](#)]
6. The World Pasta Industry in 2011. Available online: <http://www.internationalpasta.org/resources/extra/file/IPO%20AGM%202012/IPOreport2012fin2stat.pdf> (accessed on 16 October 2019).
7. Scazzina, F.; Dall’Asta, M.; Casiraghi, M.C.; Sieri, S.; Del Rio, D.; Pellegrini, N.; Brighenti, F. Glycemic index and glycemic load of commercial Italian foods. *Nutr. Metab. Cardiovasc. Dis.* **2016**, *26*, 419–429. [[CrossRef](#)] [[PubMed](#)]
8. Atkinson, F.S.; Foster-Powell, K.; Brand-Miller, J.C. International tables of glycemic index and glycemic load values: 2008. *Diabetes Care* **2008**, *31*, 2281–2283. [[CrossRef](#)] [[PubMed](#)]
9. Chiavaroli, L.; Kendall, C.W.C.; Braunstein, C.R.; Blanco Mejia, S.; Leiter, L.A.; Jenkins, D.J.A.; Sievenpiper, J.L. Effect of pasta in the context of low-glycaemic index dietary patterns on body weight and markers of adiposity: A systematic review and meta-analysis of randomised controlled trials in adults. *BMJ Open* **2018**, *8*, e019438. [[CrossRef](#)] [[PubMed](#)]
10. Pounis, G.; Castelnuovo, A.D.; Costanzo, S.; Persichillo, M.; Bonaccio, M.; Bonanni, A.; Cerletti, C.; Donati, M.B.; de Gaetano, G.; Iacoviello, L. Association of pasta consumption with body mass index and waist-to-hip ratio: Results from Moli-sani and INHES studies. *Nutr. Diabetes* **2016**, *6*, e218. [[CrossRef](#)] [[PubMed](#)]
11. Shay, C.M.; Van Horn, L.; Stamler, J.; Dyer, A.R.; Brown, I.J.; Chan, Q.; Miura, K.; Zhao, L.; Okuda, N.; Daviglus, M.L.; et al. Food and nutrient intakes and their associations with lower BMI in middle-aged US adults: The International Study of Macro-/Micronutrients and Blood Pressure (INTERMAP). *Am. J. Clin Nutr.* **2012**, *96*, 483–491. [[CrossRef](#)] [[PubMed](#)]
12. Estruch, R.; Ros, E.; Salas-Salvadó, J.; Covas, M.I.; Corella, D.; Arós, F.; Gómez-Gracia, E.; Ruiz-Gutiérrez, V.; Fiol, M.; Lapetra, J.; et al. Primary prevention of cardiovascular disease with a Mediterranean diet. *N. Engl. J. Med.* **2013**, *368*, 1279–1290. [[CrossRef](#)]
13. Huang, M.; Li, J.; Ha, M.A.; Riccardi, G.; Liu, S. A systematic review on the relations between pasta consumption and cardio-metabolic risk factors. *Nutr. Metab. Cardiovasc. Dis.* **2017**, *27*, 939–948. [[CrossRef](#)] [[PubMed](#)]
14. Vitale, M.; Masulli, M.; Cocozza, S.; Anichini, R.; Babini, A.C.; Boemi, M.; Bonora, E.; Buzzetti, R.; Carpinteri, R.; Caselli, C.; et al. Sex differences in food choices, adherence to dietary recommendations and plasma lipid profile in type 2 diabetes—The TOSCA.IT study. *Nutr. Metab. Cardiovasc. Dis.* **2016**, *26*, 879–885. [[CrossRef](#)] [[PubMed](#)]

15. Rivellese, A.A.; Boemi, M.; Cavalot, F.; Costagliola, L.; De Feo, P.; Miccoli, R.; Patti, L.; Trovati, M.; Vaccaro, O.; Zavaroni, I.; et al. Dietary habits in type II diabetes mellitus: How is adherence to dietary recommendations? *Eur. J. Clin Nutr.* **2008**, *62*, 660–664. [[CrossRef](#)] [[PubMed](#)]
16. Vaccaro, O.; Masulli, M.; Nicolucci, A.; Bonora, E.; Del Prato, S.; Maggioni, A.P.; Rivellese, A.A.; Squatrito, S.; Giorda, C.B.; Sesti, G.; et al. Effects on the incidence of cardiovascular events of the addition of pioglitazone versus sulfonylureas in patients with type 2 diabetes inadequately controlled with metformin (TOSCA.IT): A randomised, multicentre trial. *Lancet Diabetes Endocrinol.* **2017**, *5*, 887–897. [[CrossRef](#)]
17. Vaccaro, O.; Masulli, M.; Bonora, E.; Del Prato, S.; Giorda, C.B.; Maggioni, A.P.; Mocarelli, P.; Nicolucci, A.; Rivellese, A.A.; Squatrito, S.; et al. Addition of either pioglitazone or a sulfonylurea in type 2 diabetic patients inadequately controlled with metformin alone: Impact on cardiovascular events. A randomized controlled trial. *Nutr. Metab. Cardiovasc. Dis.* **2012**, *22*, 997–1006. [[CrossRef](#)] [[PubMed](#)]
18. Pala, V.; Sieri, S.; Palli, D.; Salvini, S.; Berrino, F.; Bellegotti, M.; Frasca, G.; Tumino, R.; Sacerdote, C.; Fiorini, L.; et al. Diet in the Italian EPIC cohorts: Presentation of data and methodological issues. *Tumori* **2003**, *89*, 594–607. [[CrossRef](#)] [[PubMed](#)]
19. Pisani, P.; Faggiano, F.; Krogh, V.; Palli, D.; Vineis, P.; Berrino, F. Relative validity and reproducibility of a food frequency dietary questionnaire for use in the Italian EPIC centres. *Int. J. Epidemiol.* **1997**, *26* (Suppl. 1), S152–S160. [[CrossRef](#)]
20. Salvini, S.; Parpinel, M.; Gnagnarella, P.; Maisonneuve, P.; Turrini, A. *Banca Dati di Composizione Degli Alimenti per Studi Epidemiologici in Italia*; Istituto Europeo di Oncologia: Milan, Italy, 1998.
21. Carnovale, E.; Marletta, L. *Tabella di Composizione degli Alimenti*; Crea-Nut (ex INRAN): Rome, Italy, 2000.
22. Vitale, M.; Masulli, M.; Rivellese, A.A.; Babini, A.C.; Boemi, M.; Bonora, E.; Buzzetti, R.; Ciano, O.; Cignarelli, M.; Cigolini, M.; et al. Influence of dietary fat and carbohydrates proportions on plasma lipids, glucose control and low-grade inflammation in patients with type 2 diabetes-The TOSCA.IT Study. *Eur. J. Nutr.* **2016**, *55*, 1645–1651. [[CrossRef](#)]
23. Augustin, L.S.; Kendall, C.W.; Jenkins, D.J.; Willett, W.C.; Astrup, A.; Barclay, A.W.; Björck, I.; Brand-Miller, J.C.; Brighenti, F.; Buyken, A.E.; et al. Glycemic index, glycemic load and glycemic response: An International Scientific Consensus Summit from the International Carbohydrate Quality Consortium (ICQC). *Nutr. Metab. Cardiovasc. Dis.* **2015**, *25*, 795–815. [[CrossRef](#)]
24. Giacco, R.; Brighenti, F.; Parillo, M.; Capuano, M.; Ciardullo, A.V.; Riviaccio, A.; Rivellese, A.A.; Riccardi, G. Characteristics of some wheat-based foods of the Italian diet in relation to their influence on postprandial glucose metabolism in patients with type 2 diabetes. *Br. J. Nutr.* **2001**, *85*, 33–40. [[CrossRef](#)]
25. Kristensen, M.; Jensen, M.G.; Riboldi, G.; Petronio, M.; Bügel, S.; Toubro, S.; Tetens, I.; Astrup, A. Wholegrain vs. refined wheat bread and pasta. Effect on postprandial glycemia, appetite, and subsequent ad libitum energy intake in young healthy adults. *Appetite* **2010**, *54*, 163–169. [[CrossRef](#)] [[PubMed](#)]
26. Gæde, P.; Oellgaard, J.; Carstensen, B.; Rossing, P.; Lund-Andersen, H.; Parving, H.H.; Pedersen, O. Years of life gained by multifactorial intervention in patients with type 2 diabetes mellitus and microalbuminuria: 21 years follow-up on the Steno-2 randomised trial. *Diabetologia* **2016**, *59*, 2298–2307. [[CrossRef](#)] [[PubMed](#)]
27. Mansoor, N.; Vinknes, K.J.; Veierød, M.B.; Retterstøl, K. Effects of low-carbohydrate diets v. low-fat diets on body weight and cardiovascular risk factors: A meta-analysis of randomised controlled trials. *Br. J. Nutr.* **2016**, *115*, 466–479. [[CrossRef](#)] [[PubMed](#)]
28. Catapano, A.L.; Graham, I.; De Backer, G.; Wiklund, O.; Chapman, M.J.; Drexel, H.; Hoes, A.W.; Jennings, C.S.; Landmesser, U.; Pedersen, T.R.; et al. 2016 ESC/EAS Guidelines for the Management of Dyslipidaemias. *Eur. Heart J.* **2016**, *37*, 2999–3058. [[CrossRef](#)]
29. Jellinger, P.S.; Smith, D.A.; Mehta, A.E.; Ganda, O.; Handelsman, Y.; Rodbard, H.W.; Shepherd, M.D.; Seibel, J.A. American Association of Clinical Endocrinologists' Guidelines for Management of Dyslipidemia and Prevention of Atherosclerosis. *Endocr. Pract.* **2012**, *18* (Suppl. 1), 1–78. [[CrossRef](#)]
30. Byun, S.S.; Mayat, Z.K.; Aggarwal, B.; Parekh, N.; Makarem, N. Quantity, Quality, and Timing of Carbohydrate Intake and Blood Pressure. *Curr. Nutr. Rep.* **2019**, *8*, 270–280. [[CrossRef](#)]
31. Piccinelli, R.; Arcella, D.; Buonocore, P.; Capriotti, M.; D'Addezio, L.; Le Donne, C.; Mistura, L.; Pettinelli, A.; Sette, S.; Turrini, A.; et al. L'Indagine Nazionale sui Consumi Alimentari in Italia INRAN-SCAI 2005–2006. I Consumi Alimentari in Termini di Alimenti. Osservatorio Consumi Alimentari. INRAN: Rome, Italy, 2011.

32. Fulgoni, V.L.; Bailey, R. Association of Pasta Consumption with Diet Quality and Nutrients of Public Health Concern in Adults: National Health and Nutrition Examination Survey 2009–2012. *Curr. Dev. Nutr.* **2017**, *1*, e001271. [[CrossRef](#)]



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