

Correlation Between Daily Energy Intake from Fat with Insulin Resistance in Patients with Polycystic Ovary Syndrome

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Objective: The aim of the present study was to investigate the possible correlation between the percentage of daily energy intake from fat (PEF) with insulin resistance (IR) in women with polycystic ovary syndrome (PCOS).

Methods: In this cross-sectional study, a total of 186 females with PCOS were screened. Daily dietary intake data were collected by a trained nutritionist using the 24-h dietary recall method over three consecutive days. A total of 111 subjects who had complete data were divided into two groups based on the percentage of daily energy intake from fat (PEF): the normal PEF (NPEF) group (PEF < 30%) and the high PEF (HPEF) group (PEF ≥ 30%). Pearson's correlation analysis and stepwise multivariate linear regression analysis were used to analyze the correlation of PEF with homeostasis model assessment of insulin resistance (HOMA-IR).

Results: The total prevalence rate of overweight/obesity was 80.2%. There were significant differences in waist circumference (WC), body mass index (BMI), fasting insulin, and HOMA-IR ($P < 0.001$) among the normal weight, the overweight, and the obese groups, but no significant differences were observed in total energy and dietary macronutrients intake in the three groups. The daily intake of fat and protein, fasting insulin, and HOMA-IR in the NPEF group were significantly higher than those in the HPEF group. Pearson's correlation analysis showed PEF in PCOS women was negatively correlated with BMI ($r = -0.189$, $p = 0.047$) and HOMA-IR (log-transformed) ($r = -0.217$, $p = 0.022$). Further, stepwise multivariate linear regression analysis showed PEF was negatively correlated with HOMA-IR ($p < 0.05$).

Conclusion: The percentage of daily energy intake from fat is negatively correlated with IR in women with PCOS.

Keywords: the percentage of daily energy intake from fat, polycystic ovary syndrome, insulin resistance, daily dietary intake, obesity

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Introduction

Polycystic ovary syndrome (PCOS) is the most common cause of anovulatory infertility in women of reproductive age, which is also one of the most common endocrine disorders.¹ Apart from impaired reproduction, PCOS is associated with obesity, glucolipid metabolism disorders and hypertension, which increases the risk of cardiovascular diseases (CVD) and the long-term CVD-related morbidity and mortality.² Obesity is one of the common features of PCOS, approximately 30–75% of PCOS patients are overweight or obese.³ It is reported that overweight or obese women with PCOS are at two- or three-fold higher risk compared with their non-PCOS counterparts, respectively.^{4,5} Moreover, obesity can aggravate IR and

hyperinsulinemia, even worsen the clinical, endocrine and metabolic characteristics of PCOS.⁶

Insulin resistance (IR) refers to the body's inability to utilize insulin for various reasons in the presence of elevated blood glucose concentrations, and then the body compensates to secrete excess insulin to maintain the stability of blood glucose. IR is prevalent in PCOS patients and affects approximately 75% of lean women and 95% of overweight women.⁷ The levels of IR and secondary hyperinsulinemia in overweight and obese PCOS women are significantly higher than those in nonobese PCOS women, and insulin sensitivity would worsen with increasing weight gain.^{8,9} Studies revealed that IR can increase the risk of developing diabetes and exhibit significantly higher androgens, eventually exacerbating metabolic and reproductive abnormalities in patients with PCOS.^{10,11}

Dietary factors are speculated to be the risk factors for PCOS. Unhealthy lifestyles, especially inappropriate diets, result in obesity and IR, and may even exacerbate the metabolic and reproductive disorders of PCOS.^{12,13} Increased mono/polyunsaturated fat diet yields statistically significant reductions in hyperinsulinemia in subjects with PCOS.¹⁴ Decreased trans fatty acid intake predicts reduced insulinogenic index in overweight/obese women with PCOS.¹⁵ A recent population-based case-control study found a lower energy percentage supplied by protein and carbohydrates and higher energy percentage supplied by fat in PCOS patients than in controls.¹⁶ However, there are limited data on the relationship between daily energy from dietary components with IR in subjects with PCOS. So, the aim of the present study was to explore the possible associations between the consumption of fat and IR in PCOS patients.

Patients and Methods

Participants

In the current cross-sectional study, a total of 186 female patients with PCOS from the Department of Endocrinology and Diabetes, the First Affiliated Hospital of Xiamen University (Xiamen, China) were screened. PCOS was diagnosed according to the Rotterdam definition, which has been described in our previous publication.¹⁷ Face-to-face interviews were conducted to collect subjects' health information and daily dietary intake assessment data. Of 186 patients, 111 subjects who had complete data on clinical and daily dietary intake assessment were left for the present analysis. This study was approved by the Human Research Ethics Committee

of the First Affiliated Hospital of Xiamen University (Xiamen, China). Written informed consent was obtained from each participant.

Data Collection

The anthropometric indices were recorded including height, weight, and waist circumference (WC) measured by using a calibrated scale. Blood pressure was measured with OMRON electronic sphygmomanometer after 15 minutes rest in sitting position. WC was measured at the midpoint between the lowest rib and the iliac crest. Body mass index (BMI) was calculated as the weight in kilograms divided by the square of the height in meters. We defined a BMI of 25–30 kg/m² as overweight and BMI greater than 30 kg/m² as obese according to the World Health Organization's criteria.¹⁸

Fasting blood samples were used to measure hormonal and biochemical parameters. Lipid profiles including triglyceride (TG), total cholesterol (TC), and high-density lipoprotein cholesterol (HDL-c) were determined on HITACHI 7450 analyzer (HITACHI, Tokyo, Japan). Low-density lipoprotein cholesterol (LDL-c) was calculated by Friedewald's formula.¹⁹ Fasting blood glucose (FBG) was measured by the hexokinase method. Serum fasting insulin (Roche Elecsys Insulin Test, Roche Diagnostics, Mannheim, Germany), luteinizing hormone (LH), and follicle-stimulating hormone (FSH), and testosterone (T) (Siemens Healthcare Diagnostics Inc, Massachusetts, USA) were measured by electrochemiluminescence immunoassay. Oral glucose tolerance test with 75 g of glucose (OGTT) was performed in all subjects. After a fasting blood sample was obtained, subjects ingested 250 mL solution containing 75 g of dextrose within five minutes, and then venous blood samples were obtained at 120 min for determination of plasma glucose and plasma insulin, respectively. Homeostasis model assessment of insulin resistance (HOMA-IR) was calculated using the following formula: (fasting insulin (μIU/mL) × fasting glucose (mmol/L))/22.5. Subjects were considered as having IR when the HOMA-IR index was $\geq 2.6 \times 10^{-6} \text{ mol} \times \text{U/L}^2$.²⁰

Dietary Intake Assessment

Daily dietary intake data were collected by a trained nutritionist using the 24-h dietary recall method over three consecutive days.²¹ The trained nutritionist used standard forms for the dietary recalls. To ensure that the subjects accurately reported their dietary intake, visual aids were used to assist with portion size descriptions such as food

models and photographs. The trained nutritionist did face-to-face interviews with the participants to collect all food consumption information including type, amounts, type of meal, and place of consumption during the preceding 24 hours. Subjects were divided into two groups based on the percentage of daily energy intake from fat: the normal percentage of daily energy intake from fat (less than 30% of total daily energy intake) (NPEF) group and the higher percentage of daily energy intake from fat (more than or equal to 30% of total daily energy intake) (HPEF) group.

Statistical Analyses

Data were presented as the mean \pm standard deviation (SD) for continuous variables or as median (inter-quartile range, IQR) for number and percentage for categorical variables. Skewness and kurtosis tests for normality found that fasting glucose, fasting insulin, 2-h glucose, 2-h insulin, TG, LDL-c, testosterone, LH/FSH ratio, the percentage of dietary fiber intake and HOMA-IR did not follow normal distributions. Differences between two groups were analyzed on continuous variables using the Student's *t*-test for those with normal distribution and Kruskal–Wallis test for those with skewed distribution. Pearson's correlation analysis was used to analyze the correlation of the percentage of daily energy intake from fat (PEF) with HOMA-IR (log-transformed), BMI and total daily energy intake. Univariate and stepwise linear regression analyses were used to explore the association of PEF with HOMA-IR. For multivariable linear regression analyses, age, systolic blood pressure (SBP), diastolic blood pressure (DBP), TG, TC, LDL-c, and HDL-c were adjusted. All statistical analyses were performed using SPSS version 21.0 software (IBM Corporation, Armonk, NY). All *p*-values were two-sided and *p*-value < 0.05 was considered statistically significant.

Results

Among the 111 PCOS patients whose mean ages (\pm SD) were 28.3 ± 4.9 years, 30 (27.0%) were overweight and 59 (53.2%) were obese, respectively. The total prevalence rate of overweight/obesity was 80.2%.

Clinical Characteristics and Daily Dietary Intake Categorized by BMI

The clinical characteristics and energy intake of the studied groups were shown in Table 1. There were no significant differences in age, SBP, DBP, fasting glucose, 2-h glucose, 2-h insulin, lipid profiles (TG, TC, HDL-c, and LDL-c),

testosterone, LH/FSH ratio, total energy intake, and the amount of daily dietary macronutrients intake (protein, fat, and carbohydrates, respectively) among the normal weight, the overweight and the obese groups. But there were statistically significant differences in BMI (23.2 ± 1.7 kg/m², 27.7 ± 1.2 kg/m², 33.2 ± 2.6 kg/m², respectively, $p < 0.001$), WC (79.9 ± 5.5 cm, 90.3 ± 4.6 cm, 101.7 ± 9.0 cm, respectively, $p < 0.001$), fasting insulin (82.6 (55.7–105.2) pmol/L, 110.6 (91.4–141.4) pmol/L, 145.4 (107.0–190.4) pmol/L, respectively, $p < 0.001$) and HOMA-IR (2.4 (1.7–3.2), 3.4 (2.8–4.7), 4.8 (3.6–6.4), respectively, $p < 0.001$) among these three groups. However, PEF in the obese group was lowest ($34.6 \pm 6.6\%$, $35.3 \pm 8.3\%$, $30.6 \pm 10.2\%$, respectively, $p = 0.040$) among the three groups, and the percentages of daily energy from carbohydrates in the same group were highest ($46.2 \pm 7.1\%$, $46.2 \pm 9.9\%$, $52.3 \pm 11.5\%$, respectively, $p = 0.010$).

Data Categorized by the Percentage Energy Intake from Fat

Clinical practice guidelines recommend the percentage energy from dietary fat should not exceed 30% for the management of obesity.²² In order to further explore the effect of PEF on women with PCOS, the subjects were divided into the NPEF group and the HPEF group (Table 2). In the current cohort, 79 (71.2%) PCOS women had more than 30% energy intake from fat with the average fat intake of 78.9 ± 26.4 g/day in the HPEF group. However, the mean fasting insulin (142.7 (108.2–190.0) vs 108.6 (87.5–147.3), $p = 0.005$) and HOMA-IR (4.8 (3.5–6.4) vs 3.4 (2.7–5.0), $p = 0.008$) were significantly higher in the NPEF group than that in the HPEF group. Regarding the other dietary macronutrient intakes, patients in the NPEF group had significantly higher daily intake of protein (66.1 ± 28.7 vs 84.0 ± 26.7 g/day, $p = 0.002$), but there was no significant difference in daily carbohydrate intake (233.0 ± 96.2 vs 209.7 ± 61.5 g/day, $p = 0.131$) and dietary fiber intake (8.1 (5.8–10.6) vs 8.8 (6.2–12.1) g/day, $p = 0.264$). In terms of the energy ratio from macronutrients, there were significant differences between two groups in carbohydrates (61.1 ± 9.3 vs $44.7 \pm 6.9\%$, $p < 0.001$), and fat (21.1 ± 6.3 vs $37.4 \pm 5.3\%$, $p < 0.001$) but not in protein (17.8 ± 5.4 vs $17.9 \pm 4.0\%$, $p = 0.893$).

Correlation of Percentage of Dairy Fat Intake with Insulin Resistance

Pearson's correlation analysis was performed to explore the correlation of PEF with HOMA-IR (log-transformed),

Table 1 Clinical Characteristics and Daily Dietary Intake of Patients with Polycystic Ovary Syndrome by BMI Category

	Normal Weight	Overweight	Obese	P value
N	22(19.8%)	30(27.0%)	59(53.2%)	
Age (years)	28.3 ± 4.9	27.8 ± 5.5	26.1 ± 4.9	0.184
BMI (kg/m ²)	23.2 ± 1.7	27.7 ± 1.2	33.2 ± 2.6	<0.001
WC (cm)	79.9 ± 5.5	90.3 ± 4.6	101.7 ± 9.0	<0.001
SBP (mmHg)	115.4 ± 10.6	121.8 ± 12.1	122.7 ± 13.1	0.085
DBP (mmHg)	78.3 ± 12.0	81.6 ± 7.4	82.9 ± 12.0	0.288
Fasting glucose (mmol/L)	4.7(4.6–5.3)	4.9(4.6–5.4)	5.1(4.8–5.4)	0.113
Fasting insulin (pmol/L)	82.6(55.7–105.2)	110.6(91.4–141.4)	145.4(107.0–190.4)	<0.001
2-h glucose (mmol/L)	7.7(6.1–8.9)	6.8(6.2–8.5)	6.9(5.9–8.5)	0.839
2-h insulin (pmol/L)	658.1(461.8–1143.5)	650.5(427.1–1025.7)	823.9(539.7–1191.9)	0.278
HOMA-IR	2.4(1.7–3.2)	3.4(2.8–4.7)	4.8(3.6–6.4)	<0.001
TG (mmol/L)	1.2(0.9–1.8)	1.4(1.1–2.1)	1.5(1.1–2.0)	0.378
TC (mmol/L)	4.9 ± 0.8	5.3 ± 1.0	5.2 ± 0.7	0.351
HDL-c (mmol/L)	1.3 ± 0.3	1.2 ± 0.2	1.2 ± 0.2	0.148
LDL-c (mmol/L)	2.6(2.1–2.8)	2.8(2.3–3.4)	2.9(2.6–3.4)	0.084
Testosterone (ng/dL)	43.7(30.9–81.3)	53.4(40.7–71.8)	52.1(43.6–77.6)	0.563
LH/FSH ratio	1.6(0.8–2.6)	1.3(0.7–2.1)	1.4(1.0–1.9)	0.899
Total energy (Kcal)	1710.9 ± 379.2	1900.4 ± 522.9	1744.3 ± 606.5	0.361
Carbohydrate intake (g/day)	197.0 ± 47.0	216.7 ± 65.2	223.6 ± 84.5	0.355
Daily energy intake from carbohydrates (%)	46.2 ± 7.1	46.2 ± 9.9	52.3 ± 11.5	0.010
Protein intake (g/day)	82.5 ± 26.3	85.9 ± 27.7	73.9 ± 28.9	0.135
Daily energy intake from protein (%)	19.1 ± 3.7	18.5 ± 5.2	17.1 ± 4.1	0.124
Daily fat intake (g/day)	65.8 ± 19.3	76.7 ± 32.4	61.6 ± 33.3	0.097
PEF (%)	34.6 ± 6.6	35.3 ± 8.3	30.6 ± 10.2	0.040
Daily dietary fiber intake (g/day)	9.0(5.9–12.2)	8.7(5.3–11.4)	8.4(5.9–11.0)	0.929

Note: Values are expressed as mean ± SD or median (inter-quartile range).

Abbreviations: BMI, body mass index; WC, waist circumference; SBP, systolic blood pressure; DBP, diastolic blood pressure; HOMA-IR, homeostasis model assessment of insulin resistance; TG, triglycerides; TC, total cholesterol; HDL-c, high-density lipoprotein cholesterol; LDL-c, low-density lipoprotein cholesterol; LH, luteinizing hormone, FSH, follicle stimulating hormone; PEF, the percentage of daily energy intake from fat.

BMI and total energy intake in Figure 1. PEF in PCOS patients was negatively correlated with BMI ($r = -0.189$, $p = 0.047$) and HOMA-IR (log-transformed) ($r = -0.217$, $p = 0.022$). We performed univariate and stepwise multivariate linear regression analysis to determine if PEF was still correlated with HOMA-IR (log-transformed) (Table 3). In the univariate linear regression analysis, the coefficients (95% CI) of per 1 mmHg increase of SBP and 1% of PEF for HOMA-IR (log-transformed) were 0.003 (0.0002–0.007, $p = 0.039$) and -0.005 (-0.009 – -0.001 , $p = 0.022$) respectively, and the coefficients (95% CI) of per 1 mmol/L increase of TC, TG and LDL-c for HOMA-IR (log-transformed) were 0.074 (0.026–0.123, $p = 0.003$), 0.090 (0.051–0.129, $p < 0.001$) and 0.088 (0.033–0.142, $p = 0.002$), respectively. Further stepwise multivariate linear regression analysis showed the standardized OR of per 1 SD increase of PEF for HOMA-IR (log-transformed) was -0.248 ($P = 0.008$).

Discussion

In the current study, we found the prevalence rate of overweight and obesity of 80.2% in women with PCOS, and 71.2% of PCOS women had more than 30% daily energy intake from fat. Stratification by BMI revealed there were significant differences in WC, fasting insulin, and HOMA-IR, the percentage of daily energy intake from fat and carbohydrates, while other clinical characteristics were similar among the three groups. The levels of mean fasting insulin and HOMA-IR were significantly higher in the NPEF group than that in the HPEF group. Pearson's correlation analysis demonstrated that PEF in women with PCOS was negatively correlated with BMI and HOMA-IR. Furthermore, stepwise multivariate linear regression analysis showed an independently negative correlation between PEF and HOMA-IR.

The reported prevalence of overweight and obesity in PCOS women has been described previously, and briefly,

Table 2 Clinical Characteristics and Daily Dietary Intake of Patients with Polycystic Ovary Syndrome by the Percentage of Daily Energy Intake from Fat

	NPEF Group	HPEF Group	P value
n	32(28.8%)	79 (71.2%)	
Age (years)	26.9 ± 4.7	27.08 ± 5.294	0.660
BMI (kg/m ²)	30.8 ± 4.3	29.4 ± 4.6	0.154
WC (cm)	95.4 ± 10.4	94.0 ± 11.7	0.556
SBP (mmHg)	122.6 ± 14.8	120.7 ± 11.6	0.509
DBP (mmHg)	83.0 ± 13.7	81.2 ± 9.9	0.5475
Fasting glucose (mmol/L)	5.0(4.6–5.2)	4.9(4.7–5.4)	0.997
Fasting insulin (pmol/L)	142.7(108.2–190.0)	108.6(87.5–147.3)	0.005
2-h glucose (mmol/L)	7.1(6.2–8.6)	6.9(6.0–8.6)	0.735
2-h insulin (pmol/L)	834.0(549.3–1198.6)	707.6(454.1–1048.2)	0.303
HOMA-IR	4.8(3.5–6.4)	3.4(2.7–5.0)	0.008
TG (mmol/L)	1.3(1.1–2.0)	1.4(1.1–2.0)	0.524
TC (mmol/L)	5.15 ± 0.77	5.15 ± 0.80	0.805
HDL-c (mmol/L)	1.14 ± 0.18	1.23 ± 0.22	0.054
LDL-c (mmol/L)	2.8(2.6–3.4)	2.8(2.3–3.4)	0.614
Testosterone (ng/dL)	56.2(40.1–82.4)	51.9(41.3–75.0)	0.615
LH/FSH ratio	1.3(0.9–1.4)	1.4(0.9–2.2)	0.123
Total energy (Kcal)	1519.2 ± 587.1	1885.4 ± 495.6	0.001
Daily carbohydrate intake (g/day)	233.0 ± 96.2	209.7 ± 61.5	0.131
Daily energy intake from carbohydrates (%)	61.1 ± 9.3	44.7 ± 6.9	<0.001
Daily protein intake (g/day)	66.1 ± 28.7	84.0 ± 26.7	0.002
Daily energy intake from protein (%)	17.8 ± 5.4	17.9 ± 4.0	0.893
Daily fat intake (g/day)	35.9 ± 18.3	78.9 ± 26.4	<0.001
PEF (%)	21.1 ± 6.3	37.4 ± 5.3	<0.001
Daily dietary fiber intake (g/day)	8.1(5.8–10.6)	8.8(6.2–12.1)	0.264

Note: Values are expressed as mean ± SD or median (inter-quartile range).

Abbreviations: BMI, body mass index; WC, waist circumference; SBP, systolic blood pressure; DBP, diastolic blood pressure; HOMA-IR, homeostasis model assessment of insulin resistance; TG, triglycerides; TC, total cholesterol; HDL-c, high-density lipoprotein cholesterol; LDL-c, low-density lipoprotein cholesterol; LH, luteinizing hormone; FSH, follicle stimulating hormone; PEF, the percentage of daily energy intake from fat; NPEF, within normal percentage of daily energy intake from fat (<30%); HPEF, the group with higher percentage of daily energy intake from fat (≥30%).

women with PCOS are generally more obese and more likely to have IR than age-matched control women.^{5,23} In our cohort, the prevalence rates of overweight and obesity were also higher and the severities of IR increased along with BMI (Table 1). Nutritional and dietary factors including dietary habits and patterns play an important role in prevention and treatment of PCOS.²⁴ As the first-line treatment of PCOS management,²⁵ adjusting macronutrient composition is an important part of dietary intervention. Although the optimal dietary composition in PCOS is unknown now, many previous studies focused on adjusting the proportion of protein and carbohydrates such as high-protein diet and low-carbohydrate diet²⁶ and low glycemic index (GI) diet,²⁷ which are effective in improving IR and blood glucose metabolism in PCOS women. Moreover, Nybacka et al¹⁵ demonstrated that an increase of dietary fiber intake in PCOS women was negatively correlated with increase in BMI, and contributed to the improvement

of insulin sensitivity and glucose metabolism. However, there are limited studies exploring the relationship between fat intake and endocrine abnormalities in women with PCOS.

There are few studies that identify percentage of fat intake as a significant factor of IR in women with PCOS. Some epidemiological studies revealed there was an association between higher fat intake and reduced insulin sensitivity.²⁸ Ahmadi et al²⁹ observed that PCOS women had a diet higher in fat intake compared to the healthy controls. In the present study, approximately 71.2% of subjects' daily energy percentage from fat intake exceeded the 30% recommended by the American Heart Association, which was similar to previous studies.²³ However, the percentage of daily fat intake was negatively correlated to IR and BMI, which meant the more obese PCOS women consumed less energy intake from fat per day even their daily percentages energy from dietary

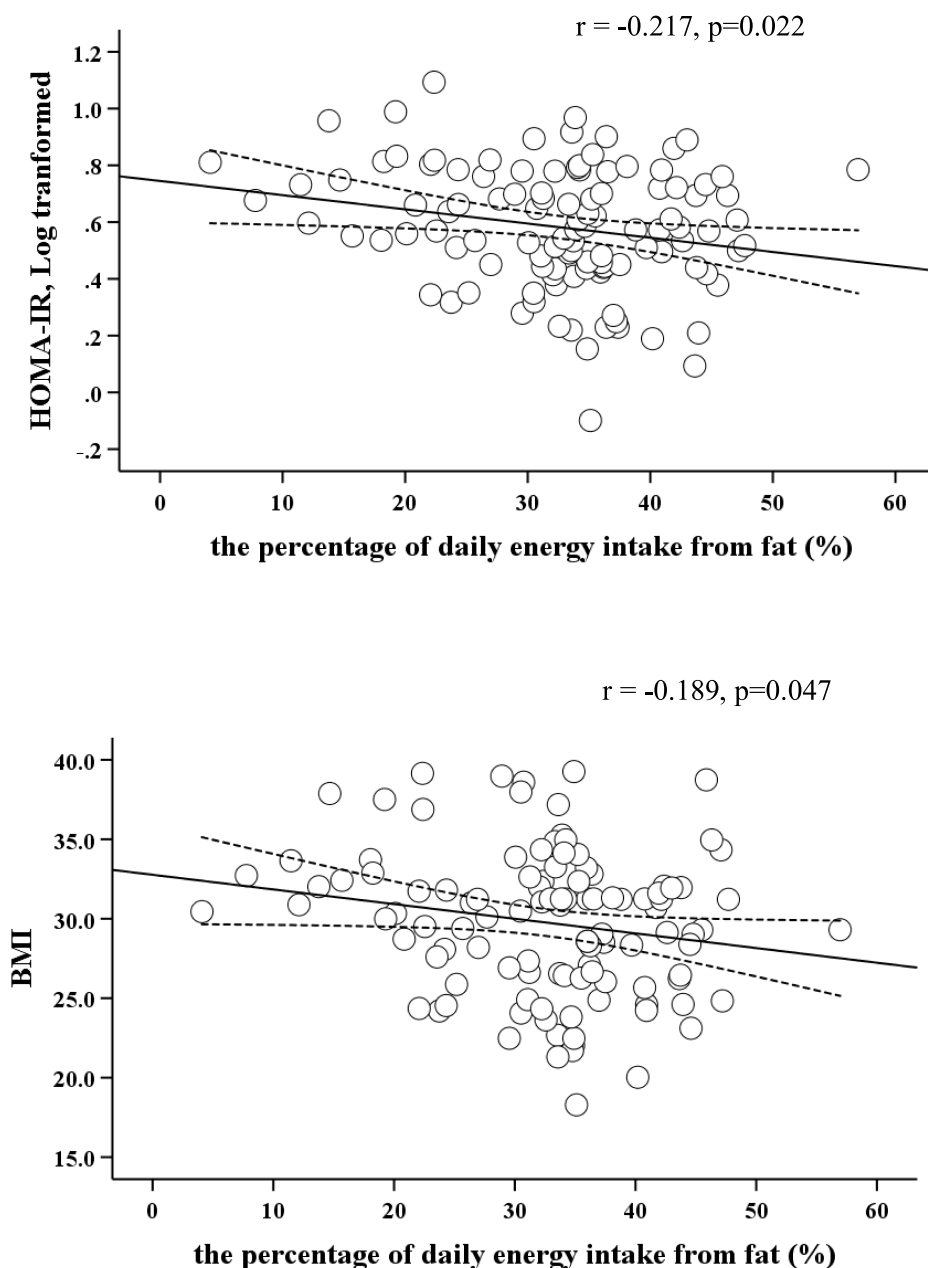


Figure 1 The correlation of the percentage of daily energy intake from fat with HOMA-IR (log-transformed) and BMI.

fat had exceeded the recommended value. We speculate that these results may be related to the following factors. First of all, it is necessary to take into account that overweight/obese patients tend to restrict their energy intake and avoid high-fat products in order to control weight, which would result in a reduction in the percentage of fat intake and daily energy intake.³⁰ Another possibility is that the improvement of IR in obese or overweight PCOS women may be related to polyunsaturated fatty acids. Although fat is considered to be a dietary component

associated with obesity,³¹ in fact, fat is a blend of different fatty acids and different fatty acids have different effects on IR. Riserus³² et al found that polyunsaturated fatty acids and monounsaturated fatty acids had benefits in improving insulin sensitivity and preventing type 2 diabetes, which was consistent with the experimental animal studies which showed that polyunsaturated fatty acids had a positive effect on insulin sensitivity.³³ In contrast, saturated and trans fatty acids will worsen IR. In addition, daily dietary intake data using standard forms for dietary

Table 3 Correlation of the Percentage of Daily Energy Intake from Fat with HOMA-IR (Log Transformed) Using Univariate and Stepwise Linear Regression Analysis

	Univariate		Stepwise	
	Unadjusted Coefficient (95% CI)	P value	Standardized Coefficients	P value
Age	−0.004 (−0.012 to 0.004)	0.312	–	–
SBP	0.003 (0.0002 to 0.007)	0.039	0.191	0.041
DBP	0.003 (−0.001 to 0.007)	0.153	–	–
TC	0.074 (0.026 to 0.123)	0.003	–	–
TG	0.090 (0.051 to 0.129)	<0.001	0.346	<0.001
LDL-c	0.088 (0.033 to 0.142)	0.002	0.242	0.011
HDL-c	−0.117 (−0.300 to 0.065)	0.206	–	–
PEF	−0.005 (−0.009 to −0.001)	0.022	−0.248	0.008

Abbreviations: SBP, systolic blood pressure; DBP, diastolic blood pressure; HOMA-IR, homeostasis model assessment of insulin resistance; TG, triglycerides; TC, total cholesterol; HDL-c, high-density lipoprotein cholesterol; LDL-c, low-density lipoprotein cholesterol; PEF, the percentage of daily energy intake from fat.

recall for obese/overweight patients may underestimate the dietary intake information.

Our study has some limitations. First, the sample size in this study was relatively small, and further prospective studies with large sample size are needed to evaluate the relationship between PEF with IR. Second, we used dietary recall to reflect dietary intake evaluation in the present study. This assessment cannot avoid the risk of underestimation, and subjects with higher BMI are more likely to underreport their food intake.³⁰ Also, there may be great personal differences in the methods of diet recall. In a future study, diet photos or mobile application (APP) for 7-day weighed food diaries could be used to record and analyze diet more accurately. Third, the diagnosis of IR was based on HOMA-IR rather than hyperinsulinemic euglycemic clamp. Fourth, in the multivariate linear regression analysis, we did not use variables such as BMI, WC, and the percentage of daily energy intake from other macronutrients as confounding factors, which may have reduced the study's power. Fifth, in the present study there were also no healthy women with regular diet as a control, which could help guide the significance of PCOS patients whether they are lean or obese. In addition, this was a cross-sectional study and we did not analyze the proportion of various fatty acids in fat. Therefore, the causality between PEF and IR in PCOS cannot be drawn. Further research is needed to explore the true relationship of daily energy intake from fat with IR in patients with PCOS in the future.

Conclusion

In summary, the current results indicate that higher percentage of daily energy intake from fat was negatively

correlated with IR in women with PCOS. Further studies are needed to clarify the mechanisms underlying the association between the percentage of fat intake and metabolic abnormalities in PCOS in order to provide more information on dietary intake and dietary composition in women with PCOS to improve fertility and weight loss.

Statement of Ethics

All procedures performed in this study were in accordance with the Helsinki declaration and its later amendments or comparable ethical standards. This study was approved by the Human Research Ethics Committee of the First Affiliated Hospital of Xiamen University (Xiamen, China) (Reference number: KYH2018-017).

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Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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Disclosure

The authors declare that they have no conflicts of interest for this work.

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