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The relationship between anthropometric and metabolic risk factors and testicular function in healthy young men

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Objective: This study investigated the relationship of anthropometric and metabolic risk factors with seminal and sex steroidal hormone parameters in a screened population of healthy males.
Methods: The participants were healthy young men without chronic or congenital diseases. The body composition parameters that we investigated were measured weight, height, and waist circumference (WC), as well as bioelectrical impedance analysis. Semen samples were analyzed for semen volume, sperm concentration, sperm motility and morphology, seminal pH, and liquefaction time. Biochemistry analysis, including glucose and lipid metabolism parameters, was conducted on fasting blood samples. Testicular volume was calculated separately for each testis using ultrasonography.
Results: Body mass index exhibited an inverse association with total sperm count. WC showed negative correlations with numerous seminal parameters, including sperm concentration, total sperm count, sperm morphology, and follicle-stimulating hormone levels. The basal metabolic rate was associated with seminal pH, liquefaction time, and sperm motility. WC, fat mass percentage, and triglyceride levels exhibited negative correlations with sex hormone binding globulin. The measures of glucose metabolism were associated with a greater number of seminal parameters than the measures of cholesterol metabolism. C-reactive protein levels were inversely associated with sperm concentration and total sperm count.
Conclusion: Anthropometric and metabolic risk factors were found to predict semen quality and alterations in sex steroidal hormone levels.

Keywords: Body mass index; C-reactive protein; Electric impedance; Glucose; Lipid metabolism; Semen analysis; Sex hormone-binding globulin; Waist circumference

Introduction

Sperm counts have been declining worldwide in recent decades, and infertility has become a serious health concern [1-4]. Male factor infertility has been identified in approximately 50% of infertile couples [5]. There has also been a parallel increase in chronic diseases and cancer. Studies have shown that men with poor semen quality are at risk for long-term morbidity and mortality [6-9]. Increased cancer and mortality rates related to lifestyle factors, obesity, and diabetes have been reported in men with impaired infertility [9,10], and there is growing evidence linking metabolic disturbances with male infertility. Obesity-related metabolic disorders, including metabolic syndrome, diabetes, hypogonadism, osteoporosis, and cardiovascular disease [11], may contribute to a potential decline in semen quality.

Metabolic syndrome, which is related to unhealthy dietary habits and a sedentary lifestyle, is also associated with poor general health and chronic systemic diseases. It is characterized by central obesity, dyslipidemia, insulin resistance, and hypertension [12]. In addition, both metabolic syndrome and male infertility have been associated with hypogonadism and high mortality risk [13,14]. Further studies

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have also shown that eugonadal men with a low sperm count had a higher risk for metabolic and cardiovascular diseases [13,14]. Body mass index (BMI), waist circumference (WC), and non-alcoholic fatty liver disease (the liver component of metabolic syndrome) have been associated with a decline in sperm output; however, the evidence regarding this point is still being debated and is far from conclusive [11].

Given the conflicting data and biological plausibility of this association, we conducted a screening study of healthy men. We investigated anthropometric and metabolic risk factors to identify predictive variables for seminal and sex steroidal hormone parameters.

Methods

1. Study population

This study was designed to measure the predictive associations between anthropometric, metabolic, hormonal, and seminal parameters in a population of healthy young men. Men being trained at a school for security services, arriving from different zones of the country, were included in the study. The participants underwent a thorough medical examination before acceptance to the school and had no chronic or congenital diseases, drug addiction, or anabolic steroid abuse. The study was approved by the Ethics Committee of Recep Tayyip Erdogan University (approval no. 224) and written informed consent was obtained from each participant.

All participants underwent a diagnostic workup with a thorough medical history that included comorbidities, a physical examination, anthropometric measurements, semen analysis, biochemistry assessment, and color Doppler ultrasound of the scrotum. For each participant, weight and height were measured, BMI was calculated, WC was measured, and bioelectrical impedance analysis (BIA) was performed using the Tanita BC-420MA (Tanita UK Ltd.). The physical examinations included a physical assessment for varicocele. Testis and epididymis size was measured by color Doppler ultrasound, and the volume of each testis was calculated using the following formula: testicular volume=[length (mm) ×width (mm)×height (mm)]× 0.71/1,000 (mL). Varicocele was identified by physical examination when the maximal venous diameter was >3 mm and increased with the Valsalva maneuver.

2. Assays

Fasting blood samples were drawn from all participants in the morning, between 9:00 AM and 12:00 PM. Serum sex steroidal hormones including testosterone (T), follicle-stimulating hormone (FSH), luteinizing hormone (LH), prolactin, and 17-β-estradiol were measured using a chemiluminescent microparticle immunoassay method with an Abbott Architect i2000 autoanalyzer (Abbott). The remaining metabolic parameters, including glucose and lipid metabolism, were measured by automated analyzers. The homoeostasis model assessment was calculated as fasting insulin (μU/mL)×fasting glucose (mmol/L)/22.5.

3. Semen analysis

Semen analysis was performed by two experienced biologists in the laboratory of the Andrology Unit. Semen samples were analyzed for semen volume, sperm concentration, sperm motility and morphology, seminal pH, and liquefaction time and were assessed according to the 5th edition of the World Health Organization (WHO) manual for human semen analysis. After 3 days of sexual abstinence, semen samples were collected by masturbation in a sterile container and were incubated at 37 °C during liquefaction. Subsequently, semen volume was estimated from the weight of the semen, assuming a sperm density of 1 g/mL, and was calculated by the gravimetric method. A Neubauer Improved hemocytometer (Shanghai Qijing Biochemical Instrument Co., Ltd.) was used to count the number of sperm (10^6/mL), and the total sperm count was calculated. Progressive motility was measured (WHO grades A+B) and morphology was assessed according to the Kruger strict criteria.

4. Statistics

1) Preprocessing the dataset

The R programming language (R Foundation for Statistical Computing) and SPSS version 23.0 (IBM Corp.) were used for statistical analysis. All quantitative data were summarized as minimum/maximum or arithmetic mean and standard deviation. While seminal and hormonal parameters were dependent variables in the model, metabolic and anthropometric variables were independent variables. Metabolic and anthropometric variables were used to predict the hormone concentrations and testicular volume by separately measuring the right testis and the left testis. Variations in the dependent variables were explained by the independent variables as estimated by multivariate linear regression analysis. There were no outliers in the data set. However, multiple normal distributions were not met with the Anderson-Darling test (Figure 1). Although data for the variables disrupting the multiple normality were transformed, the multiple normality assumption still could not be achieved (Figure 2).

2) Multilayer perceptron model

The multilayer perceptron (MLP) model, which is the most widely used artificial neural network model and does not require normality assumptions, has been deemed appropriate for use in statistical predictions [15]. The MLP is a feedforward model that maps input datasets to an appropriate output set by adjusting the weights between internal data nodes. Since multiple normality assumptions were not
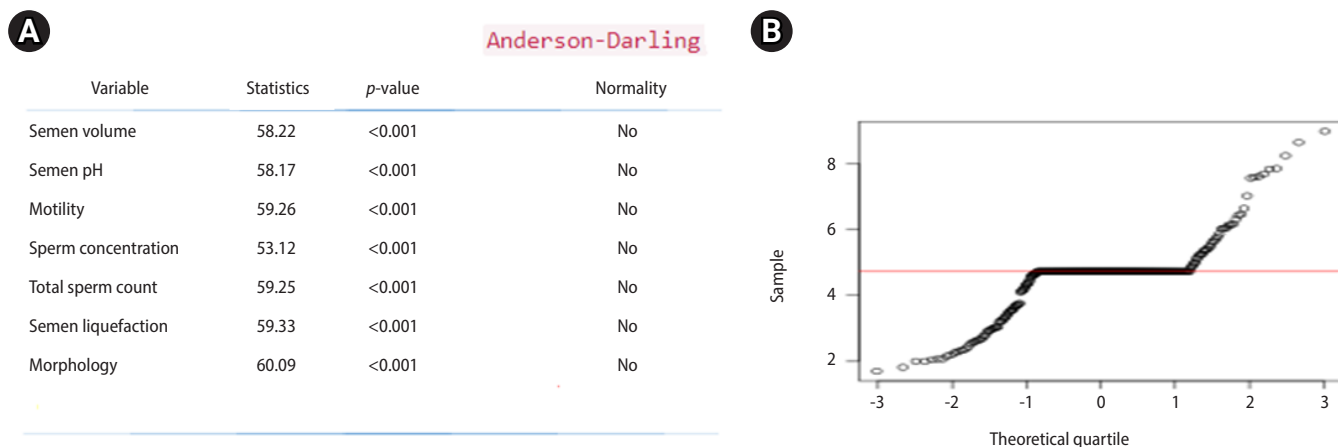
A

Anderson-Darling

Variable	Statistics	*p*-value	Normality
Semen volume	58.22	<0.001	No
Semen pH	58.17	<0.001	No
Motility	59.26	<0.001	No
Sperm concentration	53.12	<0.001	No
Total sperm count	59.25	<0.001	No
Semen liquefaction	59.33	<0.001	No
Morphology	60.09	<0.001	No

B

Figure 1. (A, B) Multiple normal distributions.

A

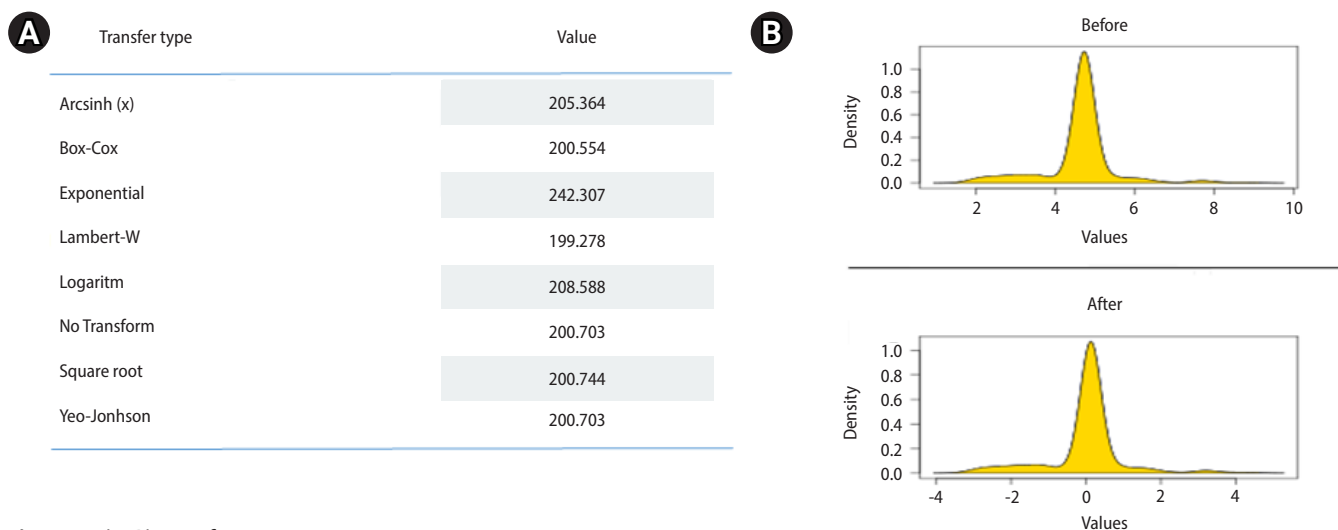| Transfer type	Value
Arcsinh (x)	205.364
Box-Cox	200.554
Exponential	242.307
Lambert-W	199.278
Logaritm	208.588
No Transform	200.703
Square root	200.744
Yeo-Jonhson	200.703

B

Figure 2. (A, B) Transformation operation.

met, we used a correlation matrix to select the variables to be used in the MLP model. For the correlation matrix, Spearman rank correlations were examined between the seminal and metabolic variables, and between the hormonal and metabolic parameters. Independent variables with the highest correlations to the dependent variables were included and analyzed in the MLP model.

The data set was divided into two divisions, training (80%) and testing (20%), and the sum of the squares error (SSE) and relative error (RE) values were checked to assess model performance. In addition, the variables that influenced the seminal or hormonal parameters were determined by their importance and relative importance values (relative importance >50%). The extent to which the independent variables could explain the change in dependent variables was evaluated by the lowest SSE and RE metrics.

3) Power analyses

A sample size calculation (power analysis) was conducted. If there are six or more independent variables in an estimation model with dependent and independent variables, the minimum sample size is obtained with the formula n=104+k (k=number of independent variables). In this study, because the sample size required for each group was 120 (5×24) or 5 times the number of variables, and was still 130 (104+24) when the number of variables was 6 or more, the sample size was considered sufficient.

Results

Of the 304 cohort members invited to participate in the study, 121 (39.8%) men provided a semen sample and underwent all analyses including a physical examination, biochemistry assessment, and testicular ultrasound. All participants had college degrees and similar

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socio-economic status (data not shown). The mean age of the participants was 27.9 years (range, 22 to 32), none were married, and 69 (57%) and 19 (15.7%) were smokers and consumers of mild alcohol, respectively. No participants had cardiovascular, metabolic, congenital, or chronic diseases. Varicocele (grade 2+3) was detected by physical and ultrasound examinations in the left testicle of 23 men (19%), and subclinical varicocele was observed in 10 (8.2%) and six (4.9%) men in the left and right testicles, respectively. Semen analysis revealed normospermia in 101 (83.5%) participants. Azoospermia was not detected. Metabolic syndrome was observed in three participants (2.5%) based on the National Cholesterol Education Program-Adult Treatment Panel III definition (2001). The characteristics of the participants are presented in Table 1.

Changes in the seminal and hormonal parameters (dependent variables) explained by the metabolic or anthropometric parameters (independent variables) are shown in Table 2 and 3, respectively. The highest percentage of explanation performance for seminal parameters in the training data set was found for liquefaction time (SSE=35.04, RE=0.694), and in the testing data set for sperm count (SSE=3.831) and liquefaction time (RE=0.692) (Table 2). However, the highest explanatory performance for hormonal and testicular parameters was found for LH (SSE=30.12, RE=0.76), and right testis volume (SSE=5.61) and testosterone (RE=0.76) in the training and testing data sets, respectively (Table 3).

Discussion

The associations observed in this study showed that WC may be a more useful measure for male infertility than BMI. Although BMI was associated with total sperm count, WC was additionally predictive of sperm concentration and morphology. However, basal metabolic rate (BMR) was observed as the only predictive anthropometric parameter for seminal pH, liquefaction time, and sperm motility. In terms of metabolic parameters, the measures of glucose metabolism were associated with a greater number of seminal parameters than cholesterol metabolism.

BMI has been identified as a risk factor for male infertility in several studies. However, conflicting results have been reported for numerous methodological reasons [16]. In addition, WC has been associated with a decline in semen volume [17-19], sperm concentration [19,20], total sperm count [17,18,21], sperm morphology [20], and sperm motility [14,18,19]. The studies were largely performed in infertile men whose testicular function was already impaired [17], and differences in the studies can be explained by variance in the study group proportions [17]. In addition to its associations with seminal parameters, WC was also a negative predictor of FSH, the most prominent hormone in spermatogenesis. In this study, BIA measure-

Table 1. The variables used to evaluate the relationship of anthropometric and metabolic risk factors with testicular function

Variable	Mean ± SD	Range (minimum-maximum)
Semen parameters		
Semen volume (mL)	4.45 ± 0.92	1.97–8.98
Sperm concentration (million/mL)	62.9 ± 23.6	0.1–200
Total sperm count (million/mL)	275 ± 1,132	0.824–98.7
Motility	49.3 ± 8.4	0–86
Morphology	9.8 ± 3.3	0–20
Semen pH	7.96 ± 0.25	7.2–9.0
Liquefaction (min)	25.4 ± 9.13	5–60
Anthropometric parameters		
Height (cm)	176.3 ± 2.65	167–188
Weight (kg)	73.9 ± 4.14	56.8–92.7
Waist circumference (cm)	87.7 ± 3.54	69–103
BMI (kg/m^2)	23.4 ± 1.3	18.2–27.4
Tanita measurements		
Fat mass percentage (%)	9.07 ± 1.85	3–16.9
Fat mass (kg)	6.68 ± 1.66	1.7–13.8
Total body water (%)	46.2 ± 2.2	37.4–60.1
Bone mass (kg)	3.31 ± 0.13	2.8–4
Basal metabolic rate (kJ)	8,162.8 ± 536.5	711–10,134
Metabolic age (yr)	12.23 ± 1.03	12–22
Metabolic parameters		
Glucose (mg/dL)	88.7 ± 4.12	66–109
TSH (μIU/mL)	1.39 ± 0.54	0.16–7.01
Creatinine (mg/dL)	0.83 ± 0.037	0.66–1.11
Total cholesterol (mg/dL)	159.1 ± 18.4	74–328
Triglycerides (mg/dL)	84.2 ± 26.03	24-301
HDL-C (mg/dL)	45.2 ± 5.29	22.8–76.1
LDL-C (mg/dL)	93.36 ± 16.48	21–260
Insulin (μU/mL)	5.39 ± 1.67	1.2–22.1
HOMA-IR	1.19 ± 0.32	0.42–4.25
GGT (U/L)	17.62 ± 4.27	6–60
CRP (mg/L)	0.13 ± 0.16	0.1–1.8
Hemoglobin (g/dL)	14.94 ± 1.01	12.3–17.2
Folate (ng/mL)	5.59 ± 0.85	1.5–11.8
RDW-CV (%)	13.25	11.9–18.3
Hormonal parameters		
FSH (mU/mL)	2.69 ± 0.87	0.87–8.68
LH (mU/mL)	3.06 ± 0.67	1.35–7.43
Testosterone (ng/dL)	548.92 ± 104.01	196–1,089
Estradiol (pg/mL)	26.6 ± 5	10–49
SHBG (μg/dL)	2641.2 ± 736.8	1,084–8,693
DHEA-S (nmol/L)	320 ± 51.2	109–560
Prolactin (μg/L)	6.6 ± 2.8	3.3–45
Testicular volume parameters		
Right testis volume (mL)	20.4 ± 3.3	13.6–39.3
Left testis volume (mL)	19.8 ± 3.2	7.7–45.1

SD, standard deviation; BMI, body mass index; kJ, kilojoule; TSH, thyroid-stimulating hormone; HDL-C, high density lipoprotein cholesterol; LDL-C, low density lipoprotein cholesterol; HOMA-IR, homoeostasis model assessment of insulin resistance; GGT, gamma glutamine transaminase; CRP, C-reactive protein; RDW-CV, red blood cell distribution width-coefficient of variation; FSH, follicle-stimulating hormone; LH, luteinizing hormone; SHBG, sex hormone binding globulin; DHEA-S, dehydroepiandrosterone sulfate.

Table 2. The perceptron model of the relationships between semen parameters and metabolic variables

Parameter	Semen volume		Sperm concentration		Total sperm count		Motility		Morphology		Semen pH		Liquefaction time	
	SSE	RE	SSE	RE	SSE	RE	SSE	RE	SSE	RE	SSE	RE	SSE	RE
Training (80%)	48.88 [a]	1.07	53.52 [a]	1.08	47.520 [a]	0.941	47.030 [a]	0.990	41.11 [a]	0.894	41.46 [a]	0.99	35.04 [a]	0.69
Testing (20%)	15.09	1.00 [b]	3.83	0.93 [b]	5.903	0.895 [b]	6.290	0.866 [b]	15.940	0.791 [b]	15.09	0.97 [b]	6.21	0.69 [b]
Metabolic variable														
Anthropometric														
Height	0.05	44.90	0.03	31.90	0.007	7.2	0.041	38.7	0.024	32.5	0.04	30.60	0.03	27.10
Weight	0.03	29.60	0.03	31.10	0.042	44.5	0.011	10.8	0.051	69	0.03	23.50	0.03	26.00
WC	0.03	24.80	0.08	79.90	0.052	54.2	0.045	41.7	0.049	67.3	0.04	35.00	0.01	5.70
BMI	0.00	2.50	0.03	27.70	0.064	67	0.012	11.4	0.021	29.2	0.03	29.70	0.02	17.10
Tanita														
Fat mass percentage	0.02	13.10	0.00	4.10	0.017	18.2	0.042	39.6	0.022	29.9	0.02	14.00	0.05	38.10
Fat mass	0.01	11.00	0.05	48.20	0.032	33.1	0.013	11.8	0.025	34.5	0.03	29.60	0.05	38.00
TBW	0.12	100.0	0.10	100.00	0.050	52.3	0.018	16.7	0.022	29.4	0.03	22.60	0.02	14.90
BM	0.02	15.20	0.07	68.70	0.029	29.9	0.015	14.5	0.040	54	0.03	26.20	0.03	23.90
BMR	0.08	67.40	0.03	29.90	0.030	31.9	0.076	71.2	0.033	45.5	0.07	60.10	0.12	100.00
Metabolic age	0.04	30.70	0.04	46.50	0.040	42.3	0.026	24.5	0.071	96.3	0.02	18.80	0.02	18.20
Biochemistry														
Glucose	0.08	67.10	0.01	8.70	0.025	26	0.087	81.2	0.033	44.6	0.02	20.60	0.05	39.60
TSH	0.00	1.60	0.04	39.50	0.047	49.1	0.064	60.4	0.030	40.5	0.05	46.90	0.01	6.90
Creatinine	0.10	84.20	0.01	5.90	0.081	84.7	0.024	22.2	0.062	84.7	0.01	4.60	0.05	41.20
TC	0.06	52.70	0.03	34.00	0.053	55.9	0.044	41	0.045	61.7	0.07	57.10	0.04	36.20
TG	0.05	40.90	0.05	47.70	0.043	44.9	0.018	17.3	0.025	34.7	0.06	49.10	0.04	34.30
HDL-C	0.02	15.80	0.03	27.30	0.025	26.5	0.014	12.7	0.026	35.8	0.04	37.90	0.01	10.40
LDL-C	0.00	3.60	0.04	39.20	0.004	3.8	0.016	15.1	0.039	53.3	0.09	74.50	0.04	35.10
Insulin	0.04	31.40	0.06	65.50	0.043	45.1	0.082	76.5	0.068	92.2	0.06	52.10	0.08	62.90
HOMA-IR	0.00	2.40	0.04	44.50	0.095	100	0.083	78.2	0.073	100	0.04	34.70	0.04	33.30
GGT	0.09	77.20	0.05	48.40	0.055	57.5	0.032	30	0.045	60.9	0.04	32.80	0.03	26.40
CRP	0.07	56.70	0.06	63.90	0.053	55.7	0.028	26.1	0.054	73.5	0.03	23.10	0.02	12.50
Hemoglobin	0.04	37.60	0.06	59.50	0.042	44.1	0.072	67.8	0.060	81.4	0.12	100.00	0.05	40.10
Folate	0.03	29.40	0.03	35.40	0.031	32.6	0.107	100	0.070	94.9	0.03	22.90	0.05	40.80
RDW-CV	0.01	11.40	0.06	63.80	0.039	41.4	0.030	28.3	0.012	16.7	0.01	10.80	0.11	91.60

SSE, sum of squares error; RE, relative error; WC, waist circumference; BMI, body mass index; TBW, total body water; BM, bone mass; BMR, basal metabolic rate; TSH, thyroid-stimulating hormone; TC, total cholesterol; TG, triglyceride; HDL-C, high density lipoprotein cholesterol; LDL-C, low density lipoprotein cholesterol; HOMA-IR, homoeostasis model assessment of insulin resistance; GGT, gamma glutamine transaminase; CRP, C-reactive protein; RDW-CV, red blood cell distribution width.

[a]Importance; [b]Relative importance (%).

Table 3. The perceptron model of the relationships between hormonal parameters and metabolic variables

Parameter	FSH SSE	FSH RE	LH SSE	LH RE	Testosterone SSE	Testosterone RE	Estradiol SSE	Estradiol RE	SHBG SSE	SHBG RE	DHEA-S SSE	DHEA-S RE	PRL SSE	PRL RE	Right testis volume SSE	Right testis volume RE	Left testis volume SSE	Left testis volume RE
Training (80%)	37.20	0.83	30.12	0.76	36.15	0.80	40.47	0.92	30.32	0.81	38.07	1.00	38.11	0.87	43.99	0.99	44.42	1.01
	a)		a)		a)		a)		a)		a)		a)		a)		a)	
Testing (20%)	12.81	0.97	20.64	0.85	7.35	0.76	14.99	0.81	25.07	0.87	22.88	0.99	8.82	0.87	5.61	1.02	22.68	1.04
	b)		b)		b)		b)		b)		b)		b)		b)		b)	
Metabolic variable																		
Anthropometric																		
Height	0.024	23.900	0.028	25.400	0.048	27.100	0.013	11.200	0.042	33.100	0.032	29.300	0.029	21.900	0.033	33.900	0.010	12.400
Weight	0.031	30.900	0.042	38.700	0.011	26.000	0.011	9.700	0.002	1.400	0.04	36.500	0.030	22.800	0.069	71.800	0.061	78.200
WC	0.057	57.000	0.016	15.100	0.030	5.700	0.086	74.600	0.066	52.300	0.037	34.200	0.076	57.100	0.061	63.700	0.049	63.600
BMI	0.024	23.500	0.021	19.300	0.011	17.100	0.018	15.800	0.017	13.300	0.023	20.800	0.017	12.800	0.029	30.000	0.001	1.500
Tanita																		
Fat mass percentage	0.027	26.700	0.074	68.00	0.038	38.100	0.022	19.600	0.085	67.100	0.031	28.800	0.080	60.100	0.026	27.400	0.029	37.700
Fat mass	0.034	33.500	0.105	97.200	0.071	38.000	0.075	65.300	0.006	5.000	0.006	5.900	0.023	17.200	0.042	44.000	0.030	38.500
TBW	0.016	16.200	0.004	3.500	0.041	14.900	0.041	36.000	0.058	45.400	0.042	38.300	0.026	19.800	0.036	37.300	0.057	73.700
BM	0.036	35.600	0.007	6.100	0.014	23.900	0.019	16.600	0.109	85.900	0.109	100.000	0.041	31.300	0.004	4.100	0.005	6.400
BMR	0.070	69.300	0.051	46.800	0.121	100.000	0.032	28.300	0.006	4.800	0.039	35.600	0.043	32.300	0.062	64.500	0.069	88.400
Metabolic age	0.038	38.000	0.033	30.700	0.029	18.200	0.049	42.800	0.033	26.300	0.053	48.200	0.036	26.900	0.049	51.300	0.019	24.100
Biochemistry																		
Glucose	0.013	13.000	0.050	46.5000	0.040	39.600	0.028	24.200	0.036	28.400	0.038	34.800	0.080	60.500	0.006	6.700	0.066	84.600
TSH	0.033	32.500	0.005	4.500	0.074	6.900	0.003	2.700	0.011	8.600	0.035	32.300	0.085	63.800	0.045	47.000	0.054	69.100
Creatinine	0.038	38.200	0.108	100.000	0.044	41.200	0.026	22.900	0.011	8.800	0.055	50.700	0.004	3.000	0.002	2.400	0.036	45.800
TC	0.038	38.100	0.048	44.200	0.053	36.200	0.109	95.200	0.050	39.500	0.054	49.800	0.078	58.500	0.046	48.200	0.038	48.700
TG	0.049	49.200	0.056	51.900	0.050	34.300	0.011	10.000	0.080	63.200	0.006	5.400	0.027	20.100	0.020	20.500	0.005	5.800
HDL-C	0.030	29.700	0.035	32.100	0.024	10.400	0.081	70.800	0.009	7.200	0.009	8.700	0.044	33.300	0.045	46.600	0.062	79.700
LDL-C	0.035	34.500	0.063	58.500	0.026	35.100	0.023	20.300	0.036	28.600	0.056	51.000	0.009	6.600	0.074	77.000	0.071	91.000
Insulin	0.040	39.700	0.050	46.200	0.087	62.900	0.019	16.400	0.042	32.700	0.030	27.100	0.026	19.400	0.033	34.500	0.028	35.600
HOMA-IR	0.088	87.400	0.062	57.300	0.009	33.300	0.016	13.800	0.033	25.800	0.069	62.900	0.011	8.500	0.061	62.900	0.050	64.600
GGT	0.100	100.000	0.050	46.400	0.033	26.400	0.090	78.700	0.042	32.800	0.097	89.000	0.046	35.000	0.074	77.100	0.059	75.800
CRP	0.015	14.500	0.029	26.500	0.007	12.500	0.057	49.400	0.076	60.200	0.003	2.800	0.005	3.600	0.052	53.800	0.033	42.400
Hemoglobin	0.087	86.500	0.044	40.700	0.160	40.100	0.115	100.000	0.127	100.000	0.006	5.300	0.042	31.500	0.096	100.000	0.078	100.000
Folate	0.043	43.300	0.015	13.700	0.063	40.800	0.019	17.000	0.016	12.300	0.098	90.000	0.011	8.500	0.004	4.600	0.060	76.900
RDW-CV	0.035	35.200	0.004	3.500	0.103	91.600	0.035	30.500	0.006	4.500	0.031	28.100	0.133	100.000	0.026	27.400	0.033	42.300

FSH, follicle-stimulating hormone; LH, luteinizing hormone; SSE, sum of squares error; RE, relative error; SHBG, sex hormone binding globulin; DHEA-S, dehydroepiandrosterone sulfate; PRL, prolactin; SSE, sum of squares error; RE, relative error; WC, waist circumference; BMI, body mass index; TBW, total body water; BM, bone mass; BMR, basal metabolic rate; TSH, thyroid-stimulating hormone; TC, total cholesterol; TG, triglyceride; HDL-C, high density lipoprotein cholesterol; LDL-C, low density lipoprotein cholesterol; HOMA-IR, homoeostasis model assessment of insulin resistance; GGT, gamma glutamine transaminase; CRP, C-reactive protein; RDW-CV, red blood cell distribution width.

[a] Importance; [b] Relative importance (%).

ments were extensively studied for seminal and hormonal parameters. BIA is widely used for estimating body composition [22] and is a more accurate measure of adiposity than BMI. Adiposity is strongly associated with obesity-related health problems [23]. The BMR or daily minimum level of energy utilized by a body in a physical and psychological resting state [24], was found to be the only BIA measurement predictive of seminal pH, liquefaction time, and sperm motility. Although BMR may be an indirect indicator of energy expended during sperm motility, in this study a positive prediction was observed between testicular volume and testosterone. Androgen deficiency was associated with changes in whole-body metabolism including decreased rates of body protein turnover and decreased resting energy expenditure [25]. BMR decreases by 1% to 2% every 10 years, and testosterone levels fall at an average of 1.6% per year [26,27]. Like testosterone, BMR is related to the metabolic rate of tissues [25-27] and could be used as an indicator in the clinical evaluation of patients with hypogonadism. Further studies of this relationship are needed.

Cholesterol is the precursor of sex steroidal hormones; however, little is known about the relationship of cholesterol and lipid metabolism with seminal parameters. In this study, both cholesterol and glucose metabolism were found to explain the variance in seminal parameters. Preclinical models confirm our findings that a high-fat diet may result in decreased sperm motility, sperm count, and morphology, and in increased sperm cholesterol content [28,29]. In this study, WC, fat mass percentage and triglyceride level were negatively predictive of sex hormone binding globulin (SHBG). Lower levels of SHBG have been linked to obesity [30]. Obesity has been shown to affect the gonadotropin-releasing hormone-LH/FSH pulse, which may deteriorate testicular steroidogenesis and spermatogenesis [31]. Metformin, an oral insulin-sensitizing agent that improves insulin sensitivity, has been used in human and animal models and resulted in improvements in seminal parameters [32] and in obesity-induced poor spermatogenesis [33]. Insulin resistance and dyslipidemia are associated with oxidative stress [34], and increased levels of reactive oxygen species impair germ cell production and sperm motility and give rise to sperm DNA damage [35]. Apart from our findings on lipid and glucose metabolism, we found that serum levels of C-reactive protein negatively predicted sperm concentration and total sperm count. Chronic inflammation disrupts the hypothalamic–pituitary-testicular axis and may result in poor semen quality [36].

This study had limitations. Because the cohort included unmarried healthy young men without chronic and genetic diseases, selection bias was possible. Therefore, it is difficult to generalize the results to other male populations (e.g., men with infertility or advanced age). Although the cohort was recruited from a single academic outpatient clinic, the subjects participating in this study came from multi-

ple parts of the country, and thus can be considered to represent the whole country. Only one semen sample was obtained per subject; nevertheless, it has been reported that one sample is adequate to evaluate semen quality in epidemiological studies [37]. Differences in the predictive variables between right and left testis volume may be due to volume differences, the fact that varicocele was more common on the left side, or by chance. Finally, the cross-sectional design does not permit inference of a causal relationship between metabolic and seminal parameters.

In conclusion, this study demonstrated that anthropometric and metabolic parameters have predictive relationships with seminal parameters and sex steroidal hormones. These findings add further insight into the underlying mechanisms of these relationships, which can be used to develop preventive and therapeutic interventions that improve semen quality in the management of infertile men.

Conflict of interest

No potential conflict of interest relevant to this article was reported.

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Author contributions

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