Muscular resistance training using applied pressure and its effects on the promotion of growth hormone secretion

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Abstract

Objective: We compared the effects of resistance training with and without pressure applied to the muscles on the secretion of growth hormone, insulin-like growth factor - I (IGF - I), and serum levels of lactic acid and free fatty acids.

Methods: Ten volunteers were divided into 2 groups who underwent 15 - minute resistance exercise: the AP group used a device to apply pressure to the proximal part of both upper arms and the proximal part of both thighs (AP group) and the control group exercised without the device. The subjects' background factors and physical and mental status were determined by questionnaire, and laboratory blood tests were conducted 15 minutes before exercise, immediately after exercise (0 minutes), then at 15 minutes, 60 minutes, and 120 minutes.

Results: The AP group had significantly greater secretion of growth hormone at 15 minutes after exercise compared to the control group, which actually had a decrease in growth hormone secretion. The serum lactic acid level was significantly higher in the AP group 15 minutes after exercise compared to values in the control group. The AP group had a significant rise in serum free fatty acid levels at 60 and 120 minutes after exercise, while no significant change was observed in the control group. No significant differences in mental symptoms were seen between the 2 groups; the control group reported more dyspnea.

Conclusions: Resistance training using applied pressure induces the production of lactic acid and the secretion of growth hormones, and causes serum free fatty acid levels to rise.

Key words: anti-aging medicine, exercise therapy, lactic acid, free fatty acids

Introduction

The purpose of anti-aging medicine is not simply to prolong life but to prevent the physical and mental deterioration caused by aging, and as a result to maintain a high quality of life (QOL).¹ From an anti-aging perspective, exercise is significant in that it maintains muscular volume, cardiopulmonary capability, and QOL through stress management. Another important result of exercise is the promotion of the secretion of growth hormones from the pituitary gland.¹ In the field of sports medicine, a variety of training methods are being developed with the aim of promoting the efficient secretion of growth hormones.

In this study, we focused on exercise therapy in healthy subjects using a training aid product that applies appropriate pressure to the muscles (KAATS,[®] manufacturer: Phoenix K.K. ,Tokyo,Japan),²⁻⁴ and compared its effects on the secretion of insulin-like growth factor - I (IGF - I), lactic

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acid, and free fatty acids with those of conventional resistance training.

Materials and Methods

The subjects in our study were healthy volunteers aged from 30 to 40 years old with no serious illnesses or subjective symptoms, who gave their consent to participate in this study. There were 20 subjects (10 men and 10 women). Their mean age was 38.4 ± 5.1 years, height was 165.6 ± 9.2 cm, weight was 63.0 ± 16.0 kg, and body fat percentage was $25.0\% \pm 4.1\%$.

Ten subjects each (5 men and 5 women) were divided into 2 groups: those who used the applied pressure device (AP group) and the control group. The AP group underwent a 15-minute resistance exercise by wearing the device and having a set pressure applied, via a rubber belt, to the proximal part of both upper arms and the proximal part of both thighs. The pressure was applied to both arms and legs simultaneously. The pressure was adjusted to 80mmHg in arms and 100 mmHg in legs. A sports trainer provided instructions on the use of the applied pressure device to the AP group. The control group underwent similar resistance exercises for 15 minutes without having pressure applied to their muscles. The exercises were performed in a group as indicated in written instructions under the supervision of the trainer (time keeper). These tests were conducted on February 11, 2004. The study design is shown in Figure 1.



Figure 1. Study design.

Before the exercise, The subjects' background factors, such as sex, age, height, weight, occupation, and presence of past illness or complications were entered in the Anti-Aging QOL Common Questionnaire: (AAQol),⁵⁻⁷ from which their condition was evaluated (Tables 1 through 3). The subject's physical and mental status was evaluated based on a symptom score derived from the AAQoL. This questionnaire uses an ordinal scale with the following values: 1 = none at all; 2 = almost none; 3 = slightly; 4 = moderately; 5 = severely.

Blood tests were performed 5 times: 15 minutes before exercise, immediately after exercise (0 minutes), then 15 minutes, 60 minutes, and 120 minutes. The setting of these time periods was based on the fact that, in preliminary experiments, the levels of growth hormones, lactic acid, and free fatty acids after exercise peaked at 15 minutes, 0 minutes (immediately after exercise), and 120 minutes after exercise, respectively. Pre-exercise examination items included peripheral blood tests, glutamic-oxaloacetic transaminase; glutamic-pyruvic transaminase; gammaglutamyltranspeptidase alkaline phosphatase, blood urea nitrogen, creatinine, uric acid, total cholesterol, triglycerides, electrolytes (sodium [Na+], potassium [K+], and chlorine [Cl-]), blood glucose, lactic acid, free fatty acids, glycosylated hemoglobin, insulin-like growth factor-I (IGF-I); somatomedin C, dehydroepiandrosterone sulfate (DHEA-s), and thyroid hormones (TSH, free T3, and free T4) cortisol values. Some of these items were measured over time. None of the subjects withdrew from the study.

The t-test, rank sum test, one way analysis of variances (ANOVA), or Kruskal-Wallis ANOVA was used to statistically analyze the findings from the survey and laboratory testing. The standard correlation analysis was also performed. The results were shown in terms of mean \pm standard deviation, and a probability value of less than .05 was considered to be significant.

Muscular resistance training using applied pressure and its effects on the promotion of growth hormone secretion

Table 1. Physical symptom questionnaire.									
Please tell us what	at your symptoms a	are.							
(1 None at all	2 Almost none	3 Slightly	4 Moderately	5 severely)					
Physical sympto	ms				1	2	3	4	5
Tired eyes									
Blurry eyes									
Eye pain									
Stiff shoulders									
Muscular pain/st	iffness								
Palpitations									
Dyspnea									
Tendency to gain	weight								
Weight loss; thin									
Lethargy									
No feeling of goo	od health								
Thirst									
Skin problems									
Anorexia									
Early satiety									
Epigastralgia									
Liable to catch co	olds								
Coughing and sp	utum								
Diarrhea									
Constipation									
Headache									
Dizziness									
Tinnitus									
Lumbago									
Arthralgia									
Edematous									
Easily breaking in	nto a sweat								
Frequent urinatio	n								
Hot flash									
Excessive sensiti	vity to cold								
Others ()								

Table	1. Physical	symptom	auestionr	nair
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Table 2. Mental symptom questionnaire.

(1 None at all	2 Almost none	3 Slightly	4 Moderately	5 severely)					
Physical sympto	oms				1	2	3	4	5
Irritability									
Easily angered/s	hort-tempered								
Loss of motivation	on								
No feeling of hap	opiness								
Nothing to look t	forward in life								
Daily life is not e	enjoyable								
Loss of confiden	ce								

Reluctance to talk with others			
Depressed			
A feeling of uselessness			
Shallow sleep			
Difficulty falling asleep			
Pessimism			
Lapse of memory			
Inability to concentrate			
Inability to solve problems			
Inability to make judgments readily			
Inability to sleep because of worries			
A sense of tension			
Feeling of anxiety for no special reason			
Vague feeling of fear			
Others ()			

Results

Background factors

Table 3 shows the background factors as well as the preexercise data of the subjects in the AP group and the control group. Concerning lifestyle habits such as smoking, drinking alcohol, and exercise, the number of cigarettes smoked was significantly higher in the control group (AP group: 3.0 ± 6.7 cigarettes/day versus control group: 11.5 \pm 10.0 cigarettes/day). No significant differences were seen between the 2 groups in other items. Tables 4 and 5 show the results of the investigation into physical and mental symptoms, based on the Anti-Aging QOL Common Questionnaire. Of the physical symptoms, only dyspnea occurred slightly more often in the control group than in the AP group (P = .0344). No significant differences were seen between the 2 groups in terms of their mental symptoms.

Table 3. Analysis of background factors in study participants.

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Variable	Unit	AP	Control	P value
Smoking	Cig./day	3.0 ± 6.7	11.5 ± 10.0	0.0390^{\dagger}
Drinking	Go/day x			
(in terms of sake)	day/week	1.8 ± 2.5	3.2 ± 4.7	0.0810
Exercise	Times/week	2.1 ± 2.6	0.5 ± 0.7	0.0810
Height	Cm	165.6 ± 10.5	165.7 ± 8.3	0.9739
Weight	kg	64.1 ± 16.6	62.0 ± 16.2	0.7808
BMI		23.0 ± 3.1	22.3 ± 4.0	0.6598
Body fat %	%	25.9 ± 3.1	$24.1~\pm~4.8$	0.3440
Systolic BP	mmHg	134.5 ± 22.7	132.2 ± 16.2	0.7973
Diastolic BP	mmHg	86.3 ± 12.1	80.2 ± 11.6	0.2650
Fasting blood sugar	mg/dl	95.9 ± 3.3	93.0 ± 15.2	0.5619
Lactic acid	mg/dl	1.06 ± 0.36	1.13 ± 0.31	0.6463
Free fatty acids	mEq/L	0.56 ± 0.20	0.55 ± 0.23	0.9099
HbA1c	mg/dl	4.89 ± 0.26	$4.79~\pm~0.44$	0.5444
Growth hormone	ng/ml	4.28 ± 6.99	6.44 ± 7.44	0.5119
IGF-1	ng/ml	193.9 ± 42.1	188.6 ± 67.2	0.8374
Cortisol	μgdl	18.0 ± 5.6	$17.5~\pm~5.8$	0.8504

DHEA-s	ng/ml	1801.7 ± 505.9	2426.5 ± 1572.1	0.2471
TSH	µgIU/ml	$1.79~\pm~1.13$	1.59 ± 0.83	0.6574
free T4	ng/dl	$0.88~\pm~0.13$	0.99 ± 0.16	0.1102
free T3	pg/ml	$1.38~\pm~0.23$	$1.28~\pm~0.26$	0.4726

BMI, body mass index; BP, blood pressure; IGF-I, insulin-like growth factor-I; HbA1c, glycosylated hemoglobin; TSH, thyroid stimulating hormone; T3, T4, thyroid hormones; DHEA-s, dehydroepiandrosterone sulfate. "One Go" means 180 ml of Sake.

n = 10, mean \pm SD, *t*-test.

Effects of muscle resistance training

Subjects in the AP group had a rise in serum growth hormone levels from 4.28 \pm 6.99 ng/ml before exercise to 8.82 \pm 5.94 ng/ml 15 minutes after finishing resistance training (Table 6). In contrast, subjects in the control group had a decrease in serum growth hormone levels from 6.44 \pm 7.44 ng/ml before training to 4.50 \pm 8.04 ng/ml 15 minutes after finishing resistance exercises. We examined the rate vis- à -vis the pre-exercise values, and found that the AP group showed significantly greater reactive secretion of growth hormones than the control group (P= 0.0449).

The serum lactic acid level before and after exercise changed in the AP group from 1.06 \pm 0.36 mg/dl before exercise to 2.91 \pm 1.55 mg/dl immediately after exercise and 2.14 \pm 0.87 mg/dl 15 minutes after exercise, while that of the control group changed from 1.13 \pm 0.31 mg/dl before exercise to 2.78 \pm 0.99 mg/dl immediately after exercise and 1.58 \pm 0.68 mg/dl 15 minutes after exercise (Table 6). Although the level 15 minutes after exercise had risen significantly over the pre-exercise level in the AP group, no significant differences were seen in the control group. We examined the rate vis- à-vis the pre-exercise value and found that the AP group had a significantly higher value of lactic acid 15 minutes after completion of exercise than the control group (P = 0.0259).

The serum free fatty acid levels before and after exercise changed in the AP group from 0.56 \pm 0.20 mEq/L before

exercise to 0.62 ± 0.26 mEq/L 60 minutes after exercise and 0.78 ± 0.30 mEq/L 120 minutes after exercise (Table 6). The levels of the AP group 60 and 120 minutes after exercise were both significantly higher than the preexercise levels (P < 0.01). The level changed in the control group, from 0.55 ± 0.23 mEq/L before exercise to $0.58 \pm$ 0.23 mEq/L 60 minutes after exercise and 0.67 ± 0.22 mEq/L 120 minutes after exercise. No significant rise in free fatty acid levels was seen in the control group. Similarly, a comparison between the AP group and the control group showed no significant differences in free fatty acid levels (P = 0.3750).

The serum IGF-I level before and after exercise changed in the AP group from 193.9 \pm 42.1 ng/ml before exercise to 195.1 \pm 58.9 ng/ml 60 minutes after exercise and 189.2 \pm 43.4 ng/ml 120 minutes after exercise (Table 6). The level changed in the control group, from 188.6 \pm 67.2 ng/ml before exercise to 185.6 \pm 76.4 ng/ml 60 minutes after exercise and 181.7 \pm 60.9 ng/ml 120 minutes after exercise. No significant changes in IGF-I were seen in either the AP group or the control group.

Figure 2 shows levels of growth hormone, lactic acid, and free fatty acids before and after muscle stress training. No adverse reactions or events were seen in either the AP group or the control group throughout the course of stress training.

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Symptoms	AP	Control	P value
Tired eyes	$2.4~\pm~0.7$	2.9 ± 0.9	0.1753
Blurry eyes	$2.0~\pm~0.8$	$2.2~\pm~0.9$	0.6132
Eye pain	$1.5~\pm~0.5$	$1.7~\pm~0.5$	0.3880
Stiff shoulders	3.1 ± 1.3	2.4 ± 1.2	0.2199
Muscular pain/stiffness	$2.4~\pm~0.7$	$2.2~\pm~1.0$	0.6182
Palpitations	$1.5~\pm~0.5$	$1.9~\pm~0.7$	0.1800

Table 4. Scores from the physical symptom questionnaire

Dyspnea	1.3 ± 0.5	$2.1~\pm~1.0$	0.0344^{\dagger}
Tendency to gain weight	$2.7~\pm~0.9$	2.5 ± 1.1	0.6652
Weight loss; thin	1.4 ± 0.7	2.1 ± 1.1	0.1068
Lethargy	$2.0~\pm~0.5$	$2.3~\pm~0.8$	0.3306
No feeling of good health	1.9 ± 0.3	$2.1~\pm~1.0$	0.5520
Thirst	$2.0~\pm~0.7$	$2.1~\pm~1.0$	0.7947
Skin problems	$2.2~\pm~1.2$	$2.4~\pm~1.2$	0.6410
Anorexia	$1.7~\pm~0.5$	$1.3~\pm~0.5$	0.0806
Early satiety	$1.6~\pm~0.8$	$2.0~\pm~0.8$	0.2546
Epigastralgia	1.8 ± 0.7	$1.6~\pm~0.7$	0.5560
Liable to catch colds	1.9 ± 0.6	$2.2~\pm~0.6$	0.2790
Coughing and sputum	$2.1~\pm~0.8$	$2.2~\pm~0.8$	0.7730
Diarrhea	$1.7~\pm~0.7$	$2.2~\pm~0.8$	0.1451
Constipation	1.9 ± 0.9	1.9 ± 1.3	0.9999
Headache	1.9 ± 0.9	$1.8~\pm~0.6$	0.7730
Dizziness	1.3 ± 0.5	$1.6~\pm~0.7$	0.2790
Tinnitus	$1.4~\pm~0.5$	$1.6~\pm~0.5$	0.3979
Lumbago	$2.4~\pm~1.0$	$2.1~\pm~0.7$	0.4453
Arthralgia	$1.7~\pm~0.8$	$1.7~\pm~0.7$	0.9999
Edematous	1.5 ± 0.7	$1.8~\pm~0.8$	0.3823
Easily breaking into a sweat	2.0 ± 1.3	$2.3~\pm~1.3$	0.6215
Polakisuria	1.9 ± 0.7	$1.7~\pm~0.7$	0.5350
Hot flash	$1.5~\pm~0.7$	$1.8~\pm~0.9$	0.4240
Excessive sensitivity to cold	$2.0~\pm~0.9$	2.1 ± 1.1	0.8297

This questionnaire uses an ordinal scale with the following values: 1 = none at all; 2 = almost none; 3 = slightly;

4 =moderately; 5 =severely.

n = 10, mean \pm SD, *t*-test

Table 5. Scores from the mental symptom questionnair

Table 5. Seores from the mental symptom	questionnane.		
Symptom	AP	Control	P value
Irritability	$2.7~\pm~0.8$	2.5 ± 1.1	0.6470
Easily angered	$2.9~\pm~0.7$	$2.3~\pm~0.8$	0.1033
Loss of motivation	1.9 ± 0.6	$2.1~\pm~0.9$	0.5520
No feeling of happiness	1.8 ± 0.6	$1.5~\pm~0.5$	0.2643
Nothing to look forward in life	$1.7~\pm~0.7$	1.4 ± 0.5	0.2790
Daily life is not enjoyable	$2.0~\pm~0.8$	$1.7~\pm~0.5$	0.3306
Loss of confidence	1.9 ± 0.6	$1.8~\pm~0.4$	0.6601
Reluctance to talk with others	$1.5~\pm~0.7$	$2.1~\pm~1.2$	0.1892
Depressed	$1.6~\pm~0.7$	$1.5~\pm~0.5$	0.7222
A sense of uselessness	$1.7~\pm~0.7$	1.9 ± 0.6	0.4825
Shallow sleep	1.3 ± 0.7	$2.2~\pm~1.2$	0.0575
Difficulty falling asleep	1.3 ± 0.7	$2.1~\pm~1.2$	0.0822
Pessimism	$2.0~\pm~0.7$	$2.1~\pm~1.2$	0.8201
Lapse of memory	$2.6~\pm~0.7$	2.8 ± 1.0	0.6182
Inability to concentrate	$2.1~\pm~0.6$	$2.2~\pm~0.6$	0.7142
Inability to solve problems	$2.1~\pm~0.6$	$2.1~\pm~0.7$	0.9999
Inability to make judgments readily	$2.0~\pm~0.7$	$2.3~\pm~0.8$	0.3823
Inability to sleep because of worries	$1.6~\pm~0.7$	1.8 ± 0.6	0.5109

A sense of tension	2.2 ± 0.8	$2.2~\pm~0.9$	0.9999
Feeling of anxiety for no special reason	$1.6~\pm~0.4$	$1.8~\pm~0.4$	0.3553
A vague feeling of fear	$1.4~\pm~0.5$	$1.5~\pm~0.5$	0.6733

This questionnaire uses an ordinal scale with the following values: 1 = none at all; 2 = almost none; 3 = slightly;

4 = moderately; 5 = severely.

n = 10, mean \pm SD, *t*-test.

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Table 6.	Changes	ın	measured	variables	over time.

Variable	Unit	AP	Control	P value
Growth hormones				
Pre-exercise value	ng/ml	$4.28~\pm~6.99$	$6.44~\pm~7.44$	
15 min. after exercise	ng/ml	$8.82~\pm~5.94$	$4.50~\pm~8.04$	0.0449
Lactic acid				
Pre-exercise value	mg/dl	$1.06~\pm~0.36$	$1.13~\pm~0.31$	
Right after exercise	mg/dl	$2.91 \pm 1.55^{\ddagger}$	$2.78~\pm~0.99^{\ddagger}$	0.8254
15 min. after exercise	mg/dl	$2.14~\pm~0.87^{\ddagger}$	$1.58~\pm~0.68$	0.0259
IGF-I				
Pre-exercise value	ng/ml	193.9 ± 42.1	188.6 ± 67.2	
60 min. after exercise	ng/ml	195.1 ± 58.9	185.6 ± 76.4	0.7586
120 min. after exercise	ng/ml	189.2 ± 43.4	181.7 ± 60.9	0.7548
Free fatty acid				
Pre-exercise value	mEq/L	$0.56~\pm~0.20$	$0.55~\pm~0.23$	
60 min. after exercise	mEq/L	$0.62~\pm~0.26^{\ddagger}$	$0.58~\pm~0.23$	0.7334
120 min. after exercise	mEq/L	$0.78~\pm~0.30^{\ddagger}$	$0.67~\pm~0.22$	0.3750

n = 10, mean \pm SD

[‡]P < 0.01, paired *t*-test (vis- à -vis pre-exercise values).

P value: Examination of the difference with pre-exercise values or of rates; AP group vs. control group



Figure 2. Changes over time of the levels of growth hormone, lactic acid, and free fatty acids before and after muscle stress training. The pre-exercise value for each item was set as 1, and the subsequent changes are represented as a proportion of their pre-exercise values. Parentheses show the mean pre-exercise values. GH, growth hormone; FFA, free fatty acid; AP, applied pressure group. [†]P < 0.05, t-test, AP group vs. control group. [‡]P < 0.01, paired *t*-test, vis- à -vis pre-exercise values.

Analysis by sex

Tables 7 and 8 show the results of a sex-specific study of exercise-induced reactive secretion of growth hormones. Among male subjects, growth hormone levels rose significantly in the AP group, from 0.56 ± 0.50 ng/ml before exercise to 12.54 ± 3.92 ng/ml 15 minutes after exercise (P = 0.0027), but the levels in the control group changed from 1.04 ± 1.51 ng/ml to 7.48 ± 11.01 ng/ml, also a significant change (Figure 3). Meanwhile, female subjects showed patterns of change different from those of the male subjects. First, concentrations of the growth hormone before exercise were 0.80 ± 1.09 ng/ml in men

and 9.90 \pm 7.71 ng /ml in women, showing that women had a significantly higher value than men (P < 0.05). Moreover, with female subjects, while the AP group saw no significant changes between the pre-exercise growth hormone level of 8.00 \pm 8.66 ng/ml and 5.10 \pm 5.43 ng/ml 15 minutes after exercise, the control group saw the levels drop significantly, from 11.84 \pm 7.03 ng/ml to 1.52 \pm 1.35 ng/ml (P = 0.0163). We examined the rate vis- àvis the pre-exercise value and found that the control group secreted significantly fewer growth hormones than the AP group (P = 0.0256).



Figure 3. Growth hormone levels before and after exercise according to sex of subject. GH, growth hormone. ${}^{\ddagger}P < 0.01$, paired t-test, vis- à -vis pre-exercise values

In male subjects, the serum lactic acid level changed before and after exercise in the AP group, from 1.24 ± 0.43 mg/dl before exercise, to 3.70 ± 1.77 mg/dl immediately after exercise and 2.48 ± 1.9 mg/dl 15 minutes after exercise (Table 7). In the AP group, the values immediately after exercise and 15 minutes after exercise both rose significantly over the pre-exercise values (P = 0.0087 and P = 0.0079, respectively). No significant differences were seen in the changes in lactic acid values between the AP group and the control group.

In female subjects, the serum lactic acid levels changed before and after exercise in the AP group, from 0.88 \pm 0.13 mg/dl prior to exercise, to 2.12 \pm 0.84 mg/dl immediately after exercise and 1.80 \pm 0.46mg/dl 15 minutes after exercise, while in the control group, the levels

changed from 1.06 ± 0.41 mg/dl prior to exercise to 2.56 ± 0.65 mg/dl immediately after exercise and 1.38 ± 0.37 mg/dl 15 minutes after exercise (Table 8). In the AP group, the values immediately after exercise and 15 minutes after exercise both rose significantly over the pre-exercise values (P = 0.0150 and P = 0.0084, respectively). No significant differences were seen in the changes in lactic acid values between the AP group and the control group.

In male subjects, the serum free fatty acid levels changed before and after exercise in the AP group, from 0.61 \pm 0.15 mEq/L before exercise, to 0.63 \pm 0.32 mEq/L 60 minutes after exercise and 0.83 \pm 0.35 mEq/L 120 minutes after exercise (Table 7). The AP group saw no significant changes between the pre-exercise values and the values 60 minutes after exercise and 120 minutes after exercise. In

the control group, the levels changed from 0.58 ± 0.22 mEq/L prior to exercise, to 0.57 ± 0.28 mEq/L 60 minutes after exercise and 0.65 ± 0.26 mEq/L 120 minutes after exercise. No significant rises in free fatty acid values were seen in the control group. No significant differences were seen in the changes in free fatty acid values between the AP group and the control group.

In female subjects, the serum free fatty acid levels changed before and after exercise in the AP group from 0.51 \pm 0.24 mEq/L prior to exercise to 0.61 \pm 0.21 mEq/L 60 minutes after exercise and 0.73 \pm 0.28 mEq/L

120 minutes after exercise (Table 8). In the AP group, the values 60 minutes and 120 minutes after exercise had risen significantly over pre-exercise values (both P = 0.0009). In the control group, the levels changed from 0.53 \pm 0.26 mEq/L before exercise to 0.59 \pm 0.21 mEq/L 60 minutes after exercise and 0.69 \pm 0.20 mEq/L 120 minutes after exercise. The value of the control group 120 minutes after exercise rose significantly over the pre-exercise level (P = 0.0224). No significant differences were seen in the changes in free fatty acid values between the AP group and the control group.

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Variable	Unit	AP	Control	P value
Growth hormones				
Pre-exercise value	ng/ml	$0.56~\pm~0.50$	1.04 ± 1.51	
15 min. after exercise	ng/ml	$12.54 \pm 3.92^{\ddagger}$	7.48 ± 11.01	0.0654
Lactic acid				
Pre-exercise value	mg/dl	$1.24~\pm~0.43$	$1.20~\pm~0.20$	
Right after exercise	mg/dl	$3.70~{\pm}~1.77^{\ddagger}$	$3.00~\pm~1.28^{\dagger}$	0.2310
15 min. after exercise	mg/dl	$2.48~{\pm}~1.09^{\ddagger}$	$1.78~\pm~0.89$	0.1210
IGF-I				
Pre-exercise value	ng/ml	203.3 ± 60.8	201.1 ± 86.7	
60 min. after exercise	ng/ml	221.8 ± 77.4	211.5 ± 101.5	0.3016
120 min. after exercise	ng/ml	197.6 ± 61.5	194.3 ± 85.1	0.4434
Free fatty acid				
Pre-exercise value	mEq/L	$0.61~\pm~0.15$	$0.58~\pm~0.22$	
60 min. after exercise	mEq/L	$0.63~\pm~0.32$	$0.57~\pm~0.28$	0.2461
120 min. after exercise	mEq/L	$0.83~\pm~0.35$	$0.65~\pm~0.26$	0.2461

n = 5, mean \pm SD. [†]P < 0.05, [‡]P < 0.01, paired *t*-test vis- à-vis pre-exercise values. P value: Examination of the difference with pre-exercise values or of rates; AP group vs. control group

Table 8. Changes over	time in femal	e subjects.
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e	5			
Variable	Unit	AP	Control	P value
Growth hormones				
Pre-exercise value	ng/ml	$8.00~\pm~8.66$	11.84 ± 7.03	
15 min. after exercise	ng/ml	$5.10~\pm~5.43$	$1.52\pm1.35^{\dagger}$	0.0256
Lactic acid				
Pre-exercise value	mg/dl	$0.88~\pm~0.13$	$1.06~\pm~0.41$	
Right after exercise	mg/dl	$2.12~\pm~0.84^{\dagger}$	$2.56~\pm~0.65^{\ddagger}$	0.3067
15 min. after exercise	mg/dl	$1.80~\pm~0.46^{\ddagger}$	1.38 ± 0.37	0.0593
IGF-I				
Pre-exercise value	ng/ml	$184.4~\pm~8.0$	176.2 ± 47.5	
60 min. after exercise	ng/ml	168.4 ± 5.3	159.6 ± 34.0	0.4722
120 min. after exercise	ng/ml	180.8 ± 16.6	169.1 ± 26.8	0.3958
Free fatty acid				
Pre-exercise value	mEq/L	$0.51~\pm~0.24$	$0.53~\pm~0.26$	

60 min. after exercise	mEq/L	$0.61\pm0.21^{\ddagger}$	0.59 ± 0.21	0.2044
120 min. after exercise	mEq/L	$0.73~\pm~0.28^{\ddagger}$	$0.69~\pm~0.20^{\dagger}$	0.2044

n = 5, mean \pm SD. [†]P < 0.05, [‡]P < 0.01, paired *t*-test vis- à -vis pre-exercise values); P value: Examination of the difference with pre-exercise values or of rates; AP group vs. control group

Study of sex differences

Because sex differences were seen in exercise-induced reactive secretion of growth hormones, we investigated sex differences in terms of background factors, mental symptoms, and pre-exercise examination results (Table 9). The results revealed the following lifestyle habits and background factors to be subject to sex differences: frequency of exercise (men: 2.1 ± 2.62 times/week; women: 0.5 ± 0.7 times/week, P = 0.0250), height (men: 173.4 ± 5.0 cm, women: 157.9 ± 4.2 cm, P = 0.0000),

weight (men: 74.8 \pm 14.6 kg, women: 51.2 \pm 4.2 kg, P = 0.0001), systolic pressure (men: 144.4 \pm 17.6 mmHg, women: 122.3 \pm 14.2 mmHg, (P = 0.0062), and diastolic pressure (men: 90.5 \pm 12.2 mmHg, women: 76.0 \pm 6.0 mmHg, P = 0.0034). Blood examination items that showed sex differences were growth hormone (men: 0.80 \pm 1.09 ng/ml, women: 9.92 \pm 7.71 ng/ml, P = 0.0016), DHEA-s (men: 2955.7 \pm 1082.7 ng/ml, women: 1272.5 \pm 445.2 ng/ml, P = 0.0002), and thyroid hormone T3 (men: 1.52 \pm 0.26 pg/ml, women: 1.33 \pm 0.18 pg/ml, P = 0.0335).

Table 9. Sex differences seen in background factors

Variable	Unit	Men	Women	P value
Smoking	Cig./day	3.0 ± 6.7	11.5 ± 10.0	0.0726
Drinking	Go/day x			
(in terms of sake)	day/week	1.8 ± 2.5	3.2 ± 4.7	0.0706
Exercises	Times/week	2.1 ± 2.6	$0.5~\pm~0.7$	0.0250^{\dagger}
Age	years	$38.6~\pm~5.1$	$38.2~\pm~5.3$	0.8653
Height	cm	$173.4~\pm~5.0$	$157.9~\pm~4.2$	0.0000^{\ddagger}
Weight	kg	$74.8~\pm~14.6$	$51.2~\pm~4.2$	0.0001 [‡]
BMI		$24.7~\pm~3.8$	$20.6~\pm~1.4$	0.0040^{\ddagger}
Body fat %	%	$23.7~\pm~4.8$	$26.3~\pm~2.8$	0.1463
Systolic BP	mmHg	144.4 ± 17.6	122.3 ± 14.2	0.0062^{\ddagger}
Diastolic BP	mmHg	$90.5~\pm~12.2$	$76.0~\pm~6.0$	0.0034^{\ddagger}
Fasting blood sugar	mg/dl	$95.7~\pm~15.0$	$93.2~\pm~4.0$	0.6175
Lactic acid	mg/dl	$1.22~\pm~0.32$	$0.97~\pm~0.30$	0.0887
Free fatty acid	mEq/L	$0.60~\pm~0.18$	$0.52~\pm~0.24$	0.4241
HbA1c	mg/dl	$4.88~\pm~0.44$	$4.80~\pm~0.26$	0.6284
Growth hormone	ng/ml	$0.80~\pm~1.09$	$9.92~\pm~7.71$	0.0016^{\ddagger}
IGF-I	ng/ml	$202.2~\pm~70.6$	$180.3~\pm~32.4$	0.3848
Cortisol	μ g/dl	$18.1~\pm~6.0$	$17.4~\pm~5.5$	0.8024
DHEA-s	ng/ml	2955.7 ± 1082.7	1272.5 ± 445.2	0.0002^{\ddagger}
TSH	ngIU/ml	$1.57~\pm~0.96$	$1.81~\pm~1.01$	0.5941
free T4	ng/dl	$0.93 ~\pm~ 0.178$	$0.94~\pm~0.13$	0.9091
free T3	pg/ml	$1.52~\pm~0.26$	$1.33~\pm~0.18$	0.0335^{\dagger}

BMI, body mass index; BP, blood pressure; IGF-I, insulin-like growth factor-I; HbA1c, glycosylated hemoglobin; TSH, thyroid stimulating hormone; T3, T4, thyroid hormones; DHEA-s, dehydroepiandrosterone sulfate.

n = 10, mean \pm SD, *t*-test.

"One GO" means 180 ml of Sake.

Table 10 and 11 show the results of the survey on mental symptoms using the Anti-Aging QOL Common Questionnaire. Regarding physical symptoms, men had higher scores than women for "diarrhea" (P < 0.01) and

"easily breaking into a sweat" (P < 0.01), while women had slightly higher scores than men for "constipation" (P < 0.05) and "headache" (P < 0.05). No sex differences were seen in mental symptoms.

Table 10. Sex differences seen in	physical	symptoms
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Symptoms	Men	Women	P value
Tired eyes	$2.8~\pm~0.6$	2.5 ± 1.0	0.4240
Blurry eyes	$2.2~\pm~0.8$	$2.0~\pm~0.9$	0.6132
Eye pain	1.6 ± 0.5	$1.6~\pm~0.5$	0.9999
Stiff shoulders	2.6 ± 1.4	2.9 ± 1.2	0.6055
Muscular pain/stiffness	$2.4~\pm~0.8$	$2.2~\pm~0.9$	0.6182
Palpitations	1.6 ± 0.5	$1.8~\pm~0.8$	0.5109
Dyspnea	1.6 ± 1.0	$1.8~\pm~0.8$	0.6182
Tendency to gain weight	2.6 ± 1.2	$2.6~\pm~0.8$	0.9999
Weight loss; thin	$2.0~\pm~0.9$	1.5 ± 1.0	0.2581
Lethargy	$2.4~\pm~0.7$	$1.9~\pm~0.6$	0.0962
No feeling of good health	$2.1~\pm~0.88$	$1.9~\pm~0.6$	0.5520
Thirst	$2.4~\pm~0.8$	$1.7~\pm~0.7$	0.0553
Skin problems	2.4 ± 1.2	$2.2~\pm~0.6$	0.6410
Anorexia	1.6 ± 0.5	1.4 ± 0.5	0.3979
Early satiety	$1.9~\pm~0.7$	$1.7~\pm~0.8$	0.5743
Epigastralgia	$1.8~\pm~0.8$	$1.6~\pm~0.7$	0.5560
Liable to catch colds	$2.0~\pm~0.7$	$2.1~\pm~0.6$	0.7222
Coughing and sputum	2.3 ± 0.7	$2.0~\pm~0.8$	0.3823
Diarrhea	2.4 ± 0.7	1.5 ± 0.5	0.0044^{\ddagger}
Constipation	1.4 ± 0.7	2.4 ± 1.2	0.0327^{\dagger}
Headache	1.5 ± 0.5	$2.2~\pm~0.8$	0.0314^{\dagger}
Dizziness	$1.5~\pm~0.7$	1.4 ± 0.5	0.7222
Tinnitus	$1.5~\pm~0.5$	1.5 ± 0.5	0.9999
Lumbago	$2.2~\pm~0.9$	$2.3~\pm~0.8$	0.8006
Arthralgia	$1.9~\pm~0.9$	1.5 ± 0.5	0.2317
Edematous	$1.9~\pm~0.7$	$1.4~\pm~0.7$	0.1373
Easily breaking into a sweat	2.9 ± 1.4	$1.4~\pm~0.7$	0.0064^{\ddagger}
Polakisuria	1.9 ± 0.7	$1.7~\pm~0.7$	0.5350
Hot flash	1.9 ± 1.0	1.4 ± 0.5	0.1753
Excessive sensitivity to cold	$1.4~\pm~0.5$	$2.7~\pm~1.0$	0.0013 [‡]

This questionnaire uses an ordinal scale with the following values: 1 = none at all; 2 = almost none; 3 = slightly;

4 =moderately; 5 =severely.

n = 10, mean \pm SD, *t*-test.

Table 11. Sex differences se	en in mental	symptoms
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	5 1		
Symptoms	Men	Women	P value
Irritability	3.0 ± 0.9	$2.2~\pm~0.8$	0.0544
Easily angered	$2.9~\pm~0.9$	$2.3~\pm~0.7$	0.1033
Loss of motivation	$2.1~\pm~0.9$	$1.9~\pm~0.6$	0.5520

No feeling of happiness	$1.6~\pm~0.7$	$1.7~\pm~0.5$	0.7142
Nothing to look forward in life	$1.5~\pm~0.5$	$1.6~\pm~0.7$	0.7222
Daily life is not enjoyable	$1.7~\pm~0.7$	$2.0~\pm~0.7$	0.3306
Loss of confidence	$1.8~\pm~0.6$	1.9 ± 0.3	0.6601
Reluctance to talk with others	1.9 ± 1.3	$1.7~\pm~0.7$	0.6685
Depressed	$1.5~\pm~0.7$	$1.6~\pm~0.5$	0.7222
A sense of uselessness	$1.9~\pm~0.7$	$1.7~\pm~0.5$	0.4825
Shallow sleep	1.8 ± 1.3	$1.7~\pm~0.8$	0.8409
Difficulty falling asleep	$1.7~\pm~1.3$	$1.7~\pm~0.6$	0.9999
Pessimism	$2.3~\pm~1.2$	$1.8~\pm~0.7$	0.2468
Lapse of memory	$2.9~\pm~1.0$	$2.5~\pm~0.7$	0.3136
Inability to concentrate	$2.3~\pm~0.5$	$2.0~\pm~0.7$	0.2643
Inability to solve problems	$2.2~\pm~0.6$	$2.0~\pm~0.7$	0.5001
Inability to make judgments readily	$2.1~\pm~0.7$	$2.2~\pm~0.8$	0.7730
Inability to sleep because of worries	$1.5~\pm~0.5$	$1.9~\pm~0.7$	0.1800
A sense of tension	$2.5~\pm~0.9$	$1.9~\pm~0.7$	0.1091
Feeling of anxiety for no special reason	$1.7~\pm~0.5$	$1.7~\pm~0.5$	0.9999
Vague feeling of fear	$1.3~\pm~0.5$	1.6 ± 0.5	0.1964

This questionnaire uses an ordinal scale with the following values: 1 = none at all; 2 = almost none; 3 = slightly;

4 =moderately; 5 =severely.

n = 10, mean \pm SD, *t*-test

Factors that influence reactive secretion of growth hormones

The correlation between individual subjects' growth hormone secretion rates (value 15 minutes after exercise ÷ pre-exercise value) and each background factor was studied. By sex, men had significantly higher secretion rates than women (r = 0.5362, P < 0.05) which correlated significantly with height (r = 0.5653, P < 0.01), weight (r= 0.5071, P < 0.05), systolic pressure (r = 0.4899, P < 0.05) 0.05), and diastolic pressure (r = 0.5486, P < 0.005). Analysis of the growth hormone secretion rate's common logarithm value showed that, by sex, men had a significantly higher secretion rate (r = 0.7244, P < 0.01), and that it correlated significantly with height (r = 0.6950, P < 0.01), weight (r = 0.5157, P < 0.05), systolic pressure (r = 0.5434, P < 0.005), serum DHEA-s level (r = 0.5862, P < 0.005)P < 0.01), and pre-exercise growth hormone value (r = -0.6902, *P* < 0.01).

Next, we studied the correlation between reactive secretion of growth hormones and physical/mental symptoms as revealed by the Anti-Aging QOL Common Questionnaire. The only item that showed a significant correlation with a higher secretion rate of growth hormones was "irritability," a mental symptom (r = 0.5121, P < 0.05). Conversely, items associated with inhibition of growth hormone secretion were physical symptoms such as "headache" (r = -0.4762, P < 0.05) and "excessive sensitivity to cold" (r = -0.4661, P < 0.05) and the mental symptom of "no feeling of happiness" (r = -0.4779, P < 0.05). Analysis of the growth hormone secretion rate's common logarithmic value showed that none of these items promoted growth hormone secretion, while items such as "headache" (r = -0.5315, P < 0.05), "excessive sensitivity to cold" (r = -0.6070, P < 0.01), and "vague feeling of fear" (r = -0.4792, P < 0.05) tended to inhibit such secretion.

Factors that influence free fatty acid production

The correlation between individual subjects' free fatty acid production rates (value 120 minutes after exercise \div pre-exercise value) and each background factor was studied. The results showed free fatty acid production rates to be inversely correlated with T4 (r = -0.6453, P < 0.01) and T3 (r = -0.6686, P < 0.01). In other words, the lower the thyroid hormone value, the higher the postexercise free fatty acid production rate tended to be. Moreover, pre-exercise free fatty acid values were inversely correlated with HbA1c (r = -0.5032, P < 0.05), so the higher the HbA1c level, the lower the free fatty acids value tended to be.

Discussion

It is known that, because growth hormone metabolites measurable in the urine increase with resistance training, exercise induces the reactive secretion of growth hormones. This fact has been used by sports medicine as a benchmark for developing training programs and methods. However, there are as yet no reports on the kinetics of post-exercise secretion of growth hormones, lactic acid, and free fatty acids in the Japanese population.

The results of our test suggest that 15-minute resistance training induces the production of lactic acid and the secretion of growth hormones, activates fat metabolism that promotes fat decomposition, and brings about the tendency for serum free fatty acid levels to rise. We also found that, by improving this resistance training and conducting training by applying pressure to the muscles, we were able to significantly boost the reactive secretion of growth hormones. Because changes in free fatty acids also increase significantly, we believe that the metabolism of fat cells is activated.

Sex differences

We studied the sex differences between lifestyle habits and other background factors, mental symptoms and preexercise examination data, and found that a characteristic feature seen with our subjects was that the men exercised significantly more than did the women, and tended to smoke and drink less than their female counterparts, showing that they were more conscious of the need to take care of their health. Although male and female subjects were the same ages, the "age" of DHEA-s hormone was significantly younger for men than for women, and the age of IGF-I hormone tended to be younger in men than in women. These may reflect the differences in their lifestyle habits.

A difference was seen between men and women in the kinetics of secretion of growth hormones. First, women had significantly higher pre-exercise values than men, and while no significant differences were seen in the IGF-I value, which is an indicator of growth hormone secretion, women tended to have lower values than men. There are as yet no reports providing evidence that women have higher growth hormone levels than similarly aged men, and thus the problem is how to interpret our findings. Our clinical trial was conducted at an athletics field on the grounds of Heisei International University campus. There is a possibility that taking part in a study at a site so suggestive of exercise may already have stepped up the secretion of growth hormones in some female subjects. Because the secretion of growth hormones is temporary, by the time exercise training began, the subjects may already have entered a period of inactivity or insensitivity. Needless to say, reactivity may fundamentally differ between men and women. For example, in women, postexercise growth hormone secretion may not peak 15 minutes after exercise: the peak may be postponed until even later. Regarding changes in free fatty acid levels, while men showed no significant differences, women saw their levels rise significantly. Therefore, because the effects of AP with respect to breaking down fatty acids were observed even in female subjects, women may find resistance training more useful in breaking down fat than will men.

Factors that influence reactive secretion of growth hormones

To investigate the factors that influence the reactive secretion of growth hormones, we analyzed the correlation between various background factors and the secretion rate of growth hormones before and after exercise. An interesting finding was obtained. We found that male sex, height, weight, blood pressure, and serum DHEA-s value were factors that enhanced the reactive secretion of growth hormones. It goes without saying that height and weight were analyzed with a BMI within the range of 21 to 24. DHEA, a hormone whose level decreases with age, is drawing interest in the field of anti-aging medicine. Our results showed that this hormone steps up the reactive secretion of growth hormones, which was an extremely interesting finding.

Our analysis also showed that an individual's mental status affected the reactive secretion of growth hormones. That is to say, an individual in a negative mental state, such as having "no feeling of happiness" or "a vague feeling of fear" is liable to show lower levels of secretion of growth hormones even after going through the same round of exercises as the others. These symptoms may be related to excessive stress or depressive tendencies. Our results showed that the more intense the subjects' headache, the less likely it was that growth hormones would be secreted. However, because individuals participating in this study were healthy, the "headache" in this case was mostly muscle contraction-type headache, so tension and stress may also have been involved.

Effects of exercise therapy

There is evidence that exercise has numerous positive effects on a person's physical and mental status. It also makes the body lighter and alleviates stiff shoulders, and helps make a person more friendly and at ease with others.¹ In other words, it is useful, both physically and mentally, to help make a person more active and sociable. Besides these, exercise improves menopausal symptoms, stimulates the brain, reinvigorates mental activity, and enhances memory, thought, and concentration.

Thanks to mechanization and the development of transportation systems, modern life requires little physical effort. The proportion of physically inactive people is clearly on the rise. On average, from the age of 40 years, muscular atrophy is said to progress at a rate of 1% each year through daily living alone⁸. Once a person becomes bedridden, moreover, muscular atrophy accelerates rapidly to a rate of 1% once every 2 days. Because women have a smaller muscle volume than men to begin with, they should engage themselves more actively in muscular training. Ordinarily, people do not use all their articular range of motion in their daily activities. Not using the joints promotes articular contracture, as is evident in periarthritis scapulohumeralis, which is more commonly referred to as age-related frozen or stiff shoulder. This symptom can be prevented by stretching exercises. To prevent aging, today's people should incorporate, in their daily lifestyles, a wellbalanced program of aerobic exercises, muscle training, and flexibility exercises.9

Some of the effects of exercise include the prevention of obesity, prevention of increased insulin resistance, burning of calories taken in, releasing stress, and enhancement of the quality of sleep. From the anti-aging medicine perspective, we wish to emphasize its effects on promoting the secretion of growth hormones and production of IGF-I. There are also reports that exercise is effective against arthralgia, lower back pain, and hypertension. Exercise is therefore expected to greatly contribute to enhancing a person's quality of life.

It has been shown that individuals aged 35 to 40 years can extend their healthy lifespan by 1 full year by incorporating moderate exercise into their daily lifestyle.¹⁰ If people who had been outstanding athletes while in school give up sports after graduation, the benefits of physical exercise eventually wear off. It is important, therefore, that sports and exercise should be continued throughout life. It is also vital not to overdo or overexert oneself. Overly strenuous exercises that imitate those of professional athletes do more physical harm than good.

Instructions/guidance on exercise

Individuals older than 40 years who are new to exercise therapy should be instructed to start with mild exercise. They should be encouraged to walk (an aerobic exercise) for 30 minutes 4 times a week, undergo 20-minute muscle training twice a week, and 10-minute flexibility exercises (stretching) every day. The load of aerobic exercises may be increased if necessary, based on the individual's progress in the program. The maximum may be about 1 hour per day of jogging or swimming. Competitive marathons that last 2 hours or more apply too much stress. Every muscular training menu should include exercise of the quadriceps of the thigh because these are where agerelated atrophy is the most conspicuous.⁸ These muscles play an important role in the movement of raising the thighs when standing up and walking. Pressurized training using KAATS is regarded as a part of the muscle training program. Although this is an efficient training over short periods, its safety in the elderly has not yet been confirmed. Therefore, it is not suited to older people. The program is useful for people who are young, middle-aged, or in the prime of life and who wish to boost their health or to avoid becoming bedridden.

Factors that make an elderly person become bedridden are, first, reduction in bone density, and second, reduction in muscular strength. This is often triggered by compression fractures of the lumbar spine and the femoral neck due to falling and other accidents. To minimize the number of elderly people who require nursing care, exercise and training to prevent falls and to avoid injury even after a fall are likely to become increasingly important.¹¹

Conclusion

Once people exceed the age of 30 years, if nothing is done, the volume of fat-free muscles normally decreases by 1% each year as a result of aging. Their fat tissues show a tendency to increase. In terms of anti-aging therapy, exercise therapy is extremely significant in that it promotes the secretion of growth hormones, fortifies the muscles, and consumes extra fat energy. If growth hormones can be stimulated to be secreted even more efficiently through pressurized training, this would constitute an even more effective anti-aging therapy.

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