

Received: 2021/07/28, Revised: 2021/08/31,
Accepted: 2021/09/03, Published: 2021/09/30

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Effects of marine oligomeric polyphenols on body composition and physical ability of elderly individuals with sarcopenia: a pilot study

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[Purpose] We aimed to identify the effects of marine oligomeric polyphenol (MOP) intake in elderly individuals with sarcopenia.

[Methods] Older adults (aged 65 years or older) were recruited based on the diagnostic criterion for sarcopenia and were randomly assigned to the MOP intake group (n=10) or the placebo (PBO) intake group (n=10). To determine the effect of MOP intake received for four weeks, the pre- and post-intake body composition (weight, skeletal muscle mass, and bone density) and senior fitness tests were assessed.

[Results] Our results showed there were significant differences in the skeletal muscle mass ($p=0.039$), bone density ($p=0.020$), fat-free mass index ($p=0.026$), and 2.4 m up and go test ($p=0.001$) between pre-test and post-test. There was a significant difference between the pre-test and post-test and an interaction effect for the one-leg stand test ($p=0.010$ and $p=0.049$, respectively). However, there were no significant differences in body fat percentage, calf circumference, grip strength, or the chair rise test.

[Conclusion] Some variables exhibited significant differences in the pre- and post-assessments, and there was an interaction effect for the one-leg stand. However, this was insufficient to prove the effectiveness of MOP intake in improving sarcopenia. Therefore, additional studies are essential to examine the effects of MOP intake and exercise intervention on the body composition and fitness of patients over a longer period.

[Key words] Sarcopenia, marine oligomeric polyphenols, body composition, senior fitness test

INTRODUCTION

In Korea, the proportion of adults aged 65 years or older was 7.2% in 2000 and 14.3% in 2018. This proportion is expected to exceed 20% by 2025¹.

In terms of health problems among older adults, 86.7% have had one or more chronic illnesses, and 30.8% have had minor disabilities, making daily life difficult. Thus, the health problems of older adults are considered an important social issue. Furthermore, 10.5% of the elderly experience difficulties in their daily lives, indicating the severity of health problems of older adults. This is caused by biological changes associated with aging. In particular, skeletal muscle mass (SMM) reduction and bone density weakening are substantial with increasing age².

Sarcopenia refers to a degenerative reduction in muscle mass and strength caused by aging. It is accompanied by gait speed reduction and a decline in physical functions³. Sarcopenia is strongly associated with senescence⁴. It was recently registered in the International Classification of Disease (2016) as ICD-10-CM code M62.84. Improvement plans are necessary since social costs due to sarcopenia are increasing⁵. The causal mechanisms of sarcopenia have shown that oxidative stress, pro-inflammatory cytokines, and sex steroids are the major factors contributing to the development of sarcopenia⁶⁻⁸.

Aging affects the catabolism of muscle proteins. Based on the oxidative stress theory of aging, aging is caused by the accumulation of oxidative damage to proteins by reactive oxygen (ROS) and nitrogen species⁹. This damage to proteins plays a major role in muscle atrophy via the ubiquitin-proteasome pathway¹⁰.

Food intake helps prevent and alleviate sarcopenia, and essential amino acids promote protein synthesis¹¹. Furthermore, exercise and appropriate protein and energy intake are necessary to prevent and manage sarcopenia, and the provision of protein could help reduce muscle loss. Substances such as leucine, amino acids, and creatine can increase strength¹².

Several studies have reported the beneficial effects of marine polyphenols in reducing inflammation-related conditions, achieved by its action

on enzymatic and non-enzymatic inflammatory targets¹³. Seaweeds are widely used in food, nutraceutical, pharmaceutical, and industrial companies because of their advantages to human health¹⁴. Currently, marine algal polyphenols have been shown to have beneficial effects on muscle cells and can induce muscle cell growth; however, additional *in vivo* tests are required¹⁵. Moreover, seaweed is of importance because it assures soils and crops are free from chemicals; thus, it is advantageous for human health¹⁴.

The antioxidant properties of food containing polyphenols include the inhibition of oxidative stress. According to a study by Fernando et al.¹⁶, polyphenolic compounds isolated from marine algae exhibited various beneficial biological properties. Many polyphenols are used in beauty, health, and medical products after their extraction from land-based plants and marine plants, such as those of Phaeophyceae and Rhodophyceae families. In particular, phlorotannins, which are polyphenols extracted from members of Phaeophyceae, are based on various phloroglucinol compounds and have phenyl and phenoxy with low, medium, and high molecular weights¹⁷. Phlorotannins have various biological effects, such as antioxidant, anticancer, antidiabetic, antiviral, and antihypertensive effects. Additionally, they suppress the levels of matrix metalloproteinase enzymes in cancer cells and have anti-allergy functions and protective effects against radiation¹⁸. Recently, numerous studies have reported polyphenols as effective bioactive molecules that attenuate muscle atrophy and enhance muscle health¹⁹. However, the studies focused on marine oligomeric polyphenols (MOPs) for the sarcopenic elderly are limited.

Therefore, the goal of this study was to investigate the effects of the intake of food with MOPs extracted from Phaeophyceae in older adults with sarcopenia by observing the changes in body composition (body fat percentage, body mass index [BMI; %BF]), SMM, bone density, calf circumference, and senior fitness test indices (grip strength, chair rise test, gait speed, and balance). We also provide supportive methods to improve sarcopenia in older adults.

METHODS

Participants

Older adults aged 65 years or older who were registered at the K Senior Welfare Service Center in Seoul were recruited for this study. They participated in similar programs at the same center and had similar living patterns. A total of 26 patients who did not undergo internal or orthopedic surgery or treatment within the last 6 months and did not have limitations regarding participation in this study according to expert consultation through a medical interview were included. An explanatory statement was written if there were abnormal symptoms or a patient requested to drop out of the study, and the patient was excluded (n=6). We excluded participants who had uncontrollable chronic diseases. The remaining patients were randomly assigned to the MOP intake group (n=10; men=2, women=8) or the placebo (PBO) intake group (n=10; men=2, women=8).

All patients were informed about the purpose of this study, the ingredients of foods, duration of intake, and measurements. Voluntary consent was obtained before the study. The characteristics of the study participants are shown in Table 1. This study was approved by the Korea National Sport University Institutional Review Board and was conducted in accordance with the 1975 Declaration of Helsinki guidelines (IRB number: 1263-202006-HR-037-03).

Table 1. Patient baseline characteristics (n=20).

	Age (years)	Height (cm)	Weight (kg)	ALM
MOP	74.63±9.30	154.44±13.39	59.66±12.37	14.56
PBO	73.38±10.66	154.80±6.99	60.56±9.48	15.01
p	0.806	0.947	0.873	0.682

MOP: Marine oligomeric polyphenol, PBO: Placebo, ALM: Appendicular Lean Mass (sum of lean mass from both arms and legs).

Criteria for sarcopenia

Sarcopenia refers to a decline in muscle mass, strength, and physical performance³. As a standard, appendicular lean mass (ALM; the sum of lean mass from both arms and legs) suggested by Studenski and colleagues²⁰ was used: If the ALM value was below 15.02 kg for women or 19.75 kg for men, a patient was diagnosed with sarcopenia.

Measurements and variables

Body composition

The Inbody770 (InBody Co. Korea Measurement, Korea) measured the SMM, BMI, and body fat percentage. In accordance with the ACSM guidelines²¹, all measurements were made once the patients changed into comfortable clothes. Additionally, food and fluid intake and strenuous physical activities were limited within 2 h of measurement.

Measurements using dual-energy X-ray absorptiometry

To identify the changes in bone density (t-score) and the fat-free mass index (%FFMI), dual-energy X-ray absorptiometry (DEXA; Lunar Prodigy, GE Medical Systems, Waukesha, USA) was used. DEXA has high reliability for measuring changes in partial and overall body composition²².

Anthropometry

For calf circumference, the thickest part of the calf was measured with a measuring tape while the patient sat straight with their knees bent at 90° angles¹⁵.

Grip strength

A dynamometer (Takei, Japan) was used to measure grip strength. The patient sat up straight in a chair and bent their arm approximately 15° away from the body. Then, the patient held the dynamometer such that the second joint of the fingers formed a right angle, and the measurement was taken. Measurements were taken twice; first for the left and then the right hand. The higher value was used²³.

Chair rise test

The chair rise test is a method used to measure lower

body muscle power in older adults. The patient was asked to sit on a 45 cm-tall chair. The patient held his or her arms against their chest, and the measurement was taken five times without rest at maximum speed when the patient was ready to sit and rise from the chair²⁴.

The 2.4 m up and go test

The purpose of the 2.4 m up and go test was to assess agility and dynamic balance. A chair was placed against the wall, and a cone was placed 2.4 m ahead of the chair. The patient sat in the middle of the chair with his or her back straightened. The patient placed their hands on their thighs, and their feet were placed flat on the ground. When the “go” signal was given, the patient rose from the chair, circled the cone, and walked back to sit down as quickly as possible. The time spent to accomplish this task was measured. The shortest time duration among the two attempts was recorded²⁵.

One-leg stand test

To measure static balance, a one-leg stand test was performed. This method can qualitatively and easily measure balance²⁶. In this study, the patients lifted their leg behind themselves and bent the respective knee at 90° with their eyes open. The time taken for the lifted foot of the patient to touch the floor was measured twice. The longest duration was considered for this study.

Method of intake

Patients were instructed to dissolve one spoon (0.7 g) of Mannas™ (developed by BotaMedi, Inc.), which contained MOP, in 500 ml of water. They were instructed to drink the solution on an empty stomach after waking up and before sleeping each day for 4 weeks. Mannas powder products

consist of 99% dextrin + 1% MOP. The PBO drink was made with dextrin to have a similar flavor and taste. The PBO group was instructed in a similar manner. To determine whether the participants consumed the products each day, both the welfare center employees and the research team performed phone monitoring. The dose was determined according to a study wherein the best dose to improve endurance was obtained by ingesting 72 mg (dissolved form 180 mL ionized beverage) in a single dose of “ECP,” which substantially is the same ingredient as MOP²⁷.

Data analysis and statistical methods

Descriptive statistics (mean, standard deviation) were calculated for all data from this study using SPSS version 21.0. A two-way analysis of variance with repeated measures was performed to evaluate changes in body composition and fitness over time and between groups according to food intake. The difference between the periods was determined using the Wilcoxon test, and the difference between the groups was evaluated using the Mann-Whitney U test. The statistical significance level for all tests was set at $\alpha=0.05$.

RESULTS

Changes in the body composition due to MOP intake

Body composition

When changes in the body composition were analyzed after MOP intake, there was a statistically significant increase in the SMM between the pre-test and post-test period ($F=5.184$, $p=0.039$) (Table 2). There were no significant

Table 2. Changes in the BIA.

	Group	Pre-test	Post-test	F	Significance
Weight (kg)	MOP	59.66±12.37	60.03±12.94	2.691	T: 0.123
	PBO	60.56±9.48	61.01±9.61	0.031	t×g: 0.862
SMM (kg)	MOP	20.98±5.56	21.31±5.73	0.028	G: 0.869
	PBO	21.21±2.39	21.33±2.63	5.184	T: 0.039*
BMI	MOP	24.86±3.19	24.99±3.53	1.296	t×g: 0.274
	PBO	25.39±4.46	25.58±4.47	0.003	G: 0.955
%BF	MOP	33.76±9.38	33.15±9.63	1.707	T: 0.212
	PBO	33.28±11.01	33.56±10.85	0.068	t×g: 0.798
Body water (L)	MOP	28.94±6.82	29.30±7.04	0.08	G: 0.782
	PBO	29.20±3.23	29.31±3.44	0.222	T: 0.645
Intracellular fluid (L)	MOP	17.61±4.26	17.88±4.37	1.703	t×g: 0.213
	PBO	17.80±1.83	17.89±2.02	0	G: 0.994
Extracellular fluid (L)	MOP	11.33±2.56	11.43±2.67	3.137	T: 0.098
	PBO	11.40±1.41	11.43±1.42	0.869	t×g: 0.367
Intracellular fluid (L)	MOP	17.61±4.26	17.88±4.37	0.003	G: 0.960
	PBO	17.80±1.83	17.89±2.02	5.777	T: 0.031*
Extracellular fluid (L)	MOP	11.33±2.56	11.43±2.67	1.444	t × g: 0.249
	PBO	11.40±1.41	11.43±1.42	0.004	G: 0.953
Intracellular fluid (L)	MOP	17.61±4.26	17.88±4.37	0.916	T: 0.355
	PBO	17.80±1.83	17.89±2.02	0.330	t × g: 0.575
Extracellular fluid (L)	MOP	11.33±2.56	11.43±2.67	0.001	G: 0.972
	PBO	11.40±1.41	11.43±1.42		

Values are expressed as the mean ± standard deviation. * $p<0.05$, T=time, G=group, MOP: Marine oligomeric polyphenol, PBO: placebo, BIA: bioelectric impedance analysis, t × g=interaction of time and group.

differences in weight, BMI, or body fat percentage between the groups. Furthermore, there was a statistically significant increase in the intracellular fluid between the pre-test and post-test period ($F=5.777, p=0.031$). There were no significant differences in body water or extracellular fluid.

Dual-energy X-ray absorptiometry Bone density (t-score)

Changes in the body composition based on MOP intake exhibited a statistically significant increase between the pre-test and post-test period ($F=6.929, p=0.020$) (Table 3). There were no group or interaction effects.

%FFMI

The %FFMI exhibited a statistically significant increase between the pre-test and post-test period ($F=6.206, p=0.026$) (Table 3). There were no group or interaction effects.

Lean body mass (kg)

Lean body mass exhibited a statistically significant increase between the pre-test and post-test period ($F=7.154, p=0.018$) (Table 3). There were no group or interaction effects.

Calf circumference

The calf circumference exhibited non-significant differences for time, group, and interaction effects (Table 4).

Grip strength

Grip strength exhibited non-significant differences for time, group, and interaction effect (Table 5).

The 2.4 m up and go test

The 2.4 m up and go test showed a statistically significant difference between the pre-test and post-test period ($F=16.537, p=0.001$) (Table 5). There were no group or interaction effects.

Chair rise test

The chair rise test outcomes showed no statistically significant differences for time, group, and interaction effects (Table 5).

One-leg stand test

The one-leg stand test exhibited statistically significant difference between the pre-test and post-test period ($F=8.836, p=0.01$) and interaction effects ($F=4.651, p=0.049$) (Table 5). No group effects were observed.

Table 3. Changes in DEXA measurement outcomes.

Group	Pre-test	Post-test	F	Significance
t-score MOP	-0.95±1.45	-0.84±1.42	6.929	T: 0.020*
			0.566	t × g: 0.464
t-score PBO	-0.99±0.79	-0.93±0.74	0.012	G: 0.915
			6.206	T: 0.026*
%FFMI MOP	16.29±1.31	16.77±1.02	0.231	t × g: 0.638
			0.139	G: 0.715
%FFMI PBO	16.55±0.69	16.88±1.19	7.154	T: 0.018*
			1.812	t × g: 0.200
Lean Body Mass (kg) MOP	37.50±8.35	38.22±7.91	0.002	G: 0.965
Lean Body Mass (kg) PBO	37.88±3.74	38.12±3.67		

Values are expressed as the mean ± standard deviation. * $p<0.05$, T=time, G=group, DEXA: dual-energy X-ray absorptiometry, t × g=interaction of time and group.

Table 4. Changes in the calf circumference.

Group	Pre-test	Post-test	F	Significance	
Calf circumference Left	MOP	33.94±3.51	34.05±4.23	0.488	T: 0.496
				0.054	t × g: 0.819
Calf circumference Right	PBO	34.13±2.70	34.35±2.73	0.022	G: 0.885
				0.001	T: 0.978
Calf circumference Right	MOP	34.31±3.89	34.20±4.40	0.286	t × g: 0.601
				0.005	G: 0.947

Values are expressed as the mean ± standard deviation. * $p<0.05$, T=time, G=group, t × g=interaction of time and group.

Table 5. Changes in senior fitness test measurements.

	Group	Pre-test	Post-test	F	Significance
Left Grip Strength	MOP	19.99±7.36	20.71±7.29	0.120	T: 0.734
	PBO	20.80±3.59	20.48±4.32	0.827	t × g: 0.379
Right Grip Strength	MOP	20.79±6.81	21.54±8.36	0.01	g: 0.922
	PBO	22.49±1.82	22.88±2.87	0.714	T: 0.412
Chair rise test MOP		14.63±5.01	15.75±3.92	0.072	t × g: 0.792
				0.306	g: 0.589
2.4m up& go MOP				1.583	T: 0.229
				0.063	t × g: 0.805
2.4m up& go PBO				0.057	G: 0.815
				16.537	T: 0.001**
One-leg stand MOP				1.066	t × g: 0.319
				0.138	G: 0.716
One-leg stand PBO				8.836	T: 0.010*
				4.651	t × g: 0.049*
				0.013	G: 0.773

Values are expressed as the mean ± standard deviation. T=time, G=group, t × g=interaction of time and group.

DISCUSSION

Sarcopenia has recently been recognized as a geriatric syndrome characterized by musculoskeletal reduction, strength decrease, and a decline in physical function caused by aging²⁸. It is considered a serious problem for older adults because it increases the risk of mobility disorders and falls and affects resulting mortality rates²⁹. It is also a cause of major illnesses, such as diabetes, arthritis, cancer, and stroke³⁰. Therefore, this study was conducted to test the effects of MOP intake on body composition and fitness of older adults (aged 65 years or older) with sarcopenia.

Marine polyphenols present in Phaeophyceae members have been reported to positively affect skeletal muscle by inhibiting the activity of alpha glucosidase, which decomposes consumed sugar into glucose, aids in muscle glucose uptake, and activates insulin signaling-related protein to affect glucose homeostasis. Furthermore, it inhibits the production of inflammatory cytokines in musculoskeletal cells³¹.

This study did not show a significant interaction effect; however, differences between the pre-test and post-test period in body composition factors, including SMM, bone density (t-score), %FFMI, and intracellular fluid, were significant. Various plant polyphenols have been reported to have beneficial effects on muscle mass preservation by reversing mitochondrial dysfunction originating from oxidative stress-induced muscle wasting¹⁹. Further investigation of various polyphenols on muscle mass preservation is also suggested in a study³².

In a previous study, marine polyphenols extracted from *Ecklonia cava*, belonging to family Phaeophyceae, inhibited the formation of fat cells to impede the accumulation of fat tissues and decrease the production of triglycerides, thereby

decreasing weight³³. A previous study on obese patients reported that sea polyphenol (*Ecklonia cava* extract) intake resulted in a decrease in BMI, an increase in mRNA, and a decrease in lipid peroxidation-related genes, thereby having a positive effect on obesity-related metabolic disorders³⁴. However, in this study, body composition did not show the beneficial effects of MOP, unlike previous research.

Furthermore, concerning changes in bone density after MOP intake, *Ecklonia cava* extract has been reported to alleviate bone loss, inhibit bone resorption, and increase the level of collagen to prevent post-menopausal bone loss³⁵. In the present study, in addition to the results of the body composition variables, bone density did not show significant interaction effects.

Protein catabolism and muscle wasting occur because of aging and inflammation in the body, and cell volume decreases because of insufficient cell nutrition, with the volume of intracellular fluid decreasing as a result³⁶. This study showed that MOP intake resulted in a significant increase in intracellular fluid over time. The anti-inflammatory function of the marine polyphenols³⁷ inhibits muscle catabolism to affect the increase in intracellular fluid. Furthermore, the marine polyphenols consumed in the current study are phenolic compounds that contain highly hydrophilic phlorotannins because of the molecular bonding between water and hydrogen¹¹. Thus, intracellular fluid was expected to increase, but the results only showed a significant effect by time, which was insufficient for an interaction effect.

Because sarcopenia is associated with decrease in skeletal muscles, decrease in strength, or decline in physical function, its symptoms can be assessed by simultaneously performing body composition and fitness tests²⁸. Grip strength is an indicator that can objectively measure body

strength³⁸. The thresholds for diagnosing sarcopenia in East Asians are <26 kg for men and <18 kg for women²⁸. Gait speed, which combines agility and dynamic balance, is important for preventing falls³⁹. Lower body strength and muscle power can be measured by performing the chair rise test, which is strongly related to daily life function and dysfunction⁴⁰. The results of senior fitness test¹⁹ to determine the fitness and physical functions of the older adults in this study showed no significant differences between time or groups for grip strength or the chair rise test. There were significant differences by time in the 2.4 m up and go test, which measured dynamic balance. There were significant differences between the time duration and groups in the one-leg stand test, which assessed static balance. Thus, MOP intake affected static balance.

For older adults, a decrease in SMM can decrease bone thickness, create thinner cortices, and lower bending strength (lower section modulus), which directly affect balance and increase the risk of falls⁴¹. The SMM in this study showed a significant increase between pre-test and post-test periods in the MOP intake group, but there was no interaction effect. According to a previous study on mice, muscle hypertrophy with the polyphenol-rich fraction (E80) was promoted when reloading exercises were applied to the unloaded muscle. However, during the unloading period, E80 did not prevent muscle atrophy. This suggests that exercise combined with polyphenol intake could promote or increase muscle hypertrophy⁴².

A previous study reported that polyphenols activate physiological processes, including mitochondrial biogenesis in skeletal muscle⁴³, thereby improving muscle-related fitness factors, such as agility and balance, and aerobic functions⁴⁴. The consumption of oxygen in muscles increases during exercise. It generates ROS in the tissues, and senescent muscle appears to be more susceptible to oxidative stress during exercise because of age-related ultrastructural and biochemical changes that facilitate ROS formation⁴⁵. The ability of polyphenols to remove ROS enables patients with sarcopenia to continue physical exercise⁴⁶. The MOP consumed in this study was also a strong antioxidant. Thus, elimination of the activated oxygen, which is overproduced because of exercise⁴¹, enables continued exercise. Although some variables, such as SMM, increased with the MOPs, we could not track all the nutritional intake of the participants, and this could be a limitation of this study.

The measurements and analyses to examine the changes in the body composition based on MOP intake did not show significant effects with respect to SMM, bone density (t-score), and the %FFMI. These results indicated that MOP intake might require further research to identify its effects on body composition.

In the senior fitness test conducted to examine changes in fitness, there was a significant difference between time duration for the 2.4 m up and go test. There were significant differences between time duration and groups in the one-leg stand test, which indicates static balance. Thus, MOP intake may be effective in improving the static balance. Although the analyses did not show interaction effects for muscle

mass increase, marine algal polyphenols belonging to phlorotannin regulate myogenesis by downregulating Smad signaling, a negative regulator, and upregulating insulin-like insulin growth factor-1 signaling, a positive regulator¹⁵. This might have affected the one-leg stand test, which is related to muscle function.

Additional studies on the effect of MOP intake and exercise intervention on body composition and fitness of older adults are necessary in the future.

ACKNOWLEDGEMENTS

This study is funded by BotaMedi, Inc.

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