Effectiveness of *Cosmos caudatus* in improving sarcopenia indices, dietary intake and physical fitness among older adults with probable sarcopenia and definitive sarcopenia in Kelantan



Dr Divya Vanoh, School of Health Sciences, Universiti Sains Malaysia divyavanoh@usm.my



OVERVIEW



INTRODUCTION

METHODOLOGY

RESULTS & DISCUSSION

CONCLUSION

Population aging





By 2050, the number of older adults will be twice the number of children under age 5 and almost equivalent to the number of children under 12 years.

(United Nations, 2020)



Sarcopenia

Sarcopenia is an age-related disease with a gradual loss of skeletal muscle mass, muscle strength and loss of muscle function (Papadopoulou, 2020)

PREVALENCE OF SARCOPENIA

1% - 30%

4.1% - 11.5%

EWGSOP guideline

AWGS 2019 guideline

(Ramoo et al., 2022)



Physical inactivity

(Schoufour et al., 2021; Vafa et al., 2020)

Sarcopenia



Preventing age-related loss of muscle mass and function through their roles as exogenous antioxidant and anti-inflammatory agents





Cosmos caudatus (C. caudatus)



Locally known as *ulam raja* (kings of *ulam*) (You et al., 2021)

Phytochemical contents



Benefit for muscle health:

- Flavonoids:
 - > Quercetin inhibit inflammatory receptors and their signalling pathway to decrease muscle atrophy
 - > Catechin- improve muscle mass, strength, and body endurance by
 - 1) maintain protein synthesis and degradation balance to slow down the muscle atrophy
 - 2) enhance mitochondrial biogenesis in muscle and provide sufficient energy for physiological activities
- Phenolic acid promote muscle growth and/or reduce muscle wasting while enhance the mitochondrial quality and reduce inflammation and oxidative stress

Antihypertensive, antidiabetic, antiinflammatory and neuroprotective

Rich in polyphenols

(You et al., 2018)

(Le et al., 2014; Li et al., 2020; Nikawa et al., 2021)

PROBLEM STATEMENT



Prevalence in Malaysia:



Iskandar et al. (2021)

- 20.4% probable sarcopenia Ranee et al. (2022), in Klang Valley
- 11.2% probable sarcopenia, 14.5% sarcopenia, 7.1 severe sarcopenia

Foo et al. (2023)



- 12.6% sarcopenia Nazri et al. (2023)
- 41% overall sarcopenia, 22.2% sarcopenia, 18.8 severe sarcopenia

Effects of sarcopenia:

risk of falls, disability, and morbidity

• prolonged hospital admission and/or longterm care placement might be needed

Quality of life, mortality risk

• higher burden to the patients, family members and healthcare systems.

(Foo et al., 2023; Ziaaldini et al., 2017)

Studies of Cosmos caudatus

effect of C. caudatus in humans still obscure.

Cheng et al. (2015)

- acid in the intervention group

You et al. (2021)

stress marker, cognitive function, and mood status

older adults.



• Most of the studies are limited to in-vivo and in-vitro studies but the clinical

A randomized controlled trial among subjects with type 2 diabetes
eight weeks supplementation of *C. caudatus* could improve insulin resistance, C-Reactive Protein (CRP), serum metabolite levels of branched-chain amino

• A 12-week supplementation with C. caudatus among older adults with cognitive impairment reported significant improvement in blood oxidative

• To date, no studies had used *C. caudatus* as an intervention among sarcopenic

STUDY OBJECTIVES



To investigate the effectiveness of supplementation using *Cosmos caudatus* extract in improving sarcopenia indices of older adults with probable sarcopenia and sarcopenia



To investigate the effectiveness of supplementation using *Cosmos* caudatus extract in improving dietary intake of older adults with probable sarcopenia and sarcopenia



To investigate the effectiveness of supplementation using *Cosmos* caudatus extract in improving physical fitness of older adults with probable sarcopenia and sarcopenia



METHODOLOGY



Study design

A 12-week double blind placebo controlled randomized trial



Inclusion criteria

- Older adults age 60 years and above
- Has probable sarcopenia and definitive sarcopenia diagnosed using AWGS 2019 guidelines
- Non-smoking

Not taking any other vitamin, herbal or traditional medications

Similar pattern of fruits and vegetables consumption

hospitalised / on tube feeding (\mathbf{X})

Sample size

- Calculated using the formula by Chan (2003)
- 92 subjects were recruited with postintervention standard deviation of 7.3, 90% power, and 95% confidence interval with an additional drop-out rate of 30%.
- The mean and standard deviation chosen were based on the study by Takeuchi et al. (2018)

Exclusion criteria

Older adults who are undergoing regular hemodialysis, bedridden,

(X) Older adults with severe sarcopenia (meeting all 3 criteria)

Older adults with chronic kidney diseases, diarrhea, chronic constipation or gastrointestinal diseases such as inflammatory bowel disease, irritable bowel syndrome, haemorrhoid, diverticulitis

Older adults on antibiotics for the past 30 days, corticosteroid, immunosuppressants, warfarin therapy or on medications affecting intestinal motility such as laxatives, antidepressants, opioid. anticholinergic, prebiotic and probiotic during the study period

Older adults living in a long-term care facility

Randomization and Allocation

The randomization sequence of the subjects were generated using a randomization website (https://www.randomization.com) by simple randomization. The villages were randomized to either the control or the treatment group with a 1:1 allocation ratio.

Intervention

The treatment group will receive 500 mg of *C. caudatus* extract, while the control group will receive a similar dose of maltodextrin daily for 12 weeks taken before breakfast, daily with health diet counselling for both groups. -Maltodextrin chosen due to no therapeutic effect

Adherence & Compliance

- Weekly telephone calls were made to monitor adherence and address subjects' concern (eg: presence of any side effects).
- Compliance were monitored by the researcher by asking the subjects to return the supplement bottle every month.
- Subjects were given a small diary to mark the chart daily after consuming the supplements

STUDY FLOWCHART



Excluded (n = 195)

- No sarcopenia (n = 49)
- Severe sarcopenia (n = 36)
- Smoking (n = 39)
- Other supplement intake (n = 14)
- Decline to participate (n = 29)
- Unable to contact (n = 23)
- Health problem (n = 5)



STUDY OUTCOMES



Changes in Sarcopenia Indices

• Diagnosed using Asian Working Group for Sarcopenia (AWGS)

Muscle mass

- Measured using Karada Scan Omron Body Composition Monitor
- SMI: (muscle mass/height2)
- Low muscle mass: SMI values of <7 kg/m² for men and <5.7 kg/m² for women

Muscle strength

- Hand grip strength was measured using hydraulic hand dynamometer
- Poor handgrip strength<28 kg for men and <18 kg for women

Skeletal Muscle Index (SMI)	Handgrip strength	SPPB scor
Normal	Normal	Normal
Normal	Normal	Low
Normal	Low	Low
Normal	Low	Normal
Low	Normal	Normal
Low	Normal	Low
Low	Low	Normal
Low	Low	Low

Physical performance

- Assessed using SPPB
- Consisted of three major assessments: the gait speed, balance tests and a chair stand test
- Poor physical performance: score ≤9

re Sarcopenia Status No sarcopenia Probable sarcopenia Probable sarcopenia Probable sarcopenia Probable sarcopenia Sarcopenia Sarcopenia

(Chen et al., 2020)

STUDY OUTCOMES

Changes in Dietary Intake

- Assessed using Dietary History Questionnaire (DHQ)
- Taken for seven days via interview administered method
- Data was collected using household cooking utensils such as teaspoon, dessertspoon, teacup, various sizes of bowls, rice scoop, and plates.
- Intake was analysed using Nutritionist Pro Software and was compared with the Malaysian Recommended Nutrient Intake (RNI) 2017.

Changes in Physical Fitness

Comprises of SPPB, handgrip test, back scratch test, 2-minute step test, timed-up and go test and chair sit and reach test

Back scratch test

- To assess upper body flexibility in the shoulder joint and shoulder arch on both the left and right sides
- The subject will be required to place one hand behind the head, reaching over the shoulder and one hand up the middle of the back.
- The distance in cm between the extended middle fingers will be recorded.
- If the fingertips touch, then the score will be 0.

2-minute step test

• To assess aerobic capacity thus evaluating subjects' level of functional fitness.

• The subject will be required to stand next to a wall while a mark is placed on the wall at the level that corresponds to the midway between the patella and iliac crest (top of the hip bone).

• The number of times the right knee reaches the point midway between the patella and iliac crest within 2 minutes will be recorded.

STUDY OUTCOMES

Changes in Physical Fitness

Comprises of SPPB, handgrip test, back scratch test, 2-minute step test, and chair sit and reach test \bullet

Timed-up and go test

- For balance and mobility
- Subject get up from chair, walk for 3 meters, turn around and walk back to chair to sit
- Score recorded as the fastest time to complete

- hip.

Chair sit and reach test

• To assess lower body flexibility

• The subject will be required to sit on a chair and one foot must remain flat on the floor. The other leg is required to be extended forward with the knee straight. Subject must use their hands to reach towards their toes by bending at the

• The distance is measured between the tip of the fingertips and the toes (cm)



STATISTICAL ANALYSIS

BASELINE

Independent-t-test Chi-Square test

- Missing data were handled using multiple imputation method

 - group over time were analyzed using generalized linear model repeated measure analysis.
- Differences between treatment & control • Effect size computed using Cohen's d

INTERVENTION

analysed using intention-to-treat • Data method



RESULTS & DISCUSSION



Baseline Findings

Variable	Intervention (n=47)	Control (n=45)
	Demographic	
Age, years	67.68(5.49)	70.49(6.90)
Gender		
Men	29(61.7)	22(48.9)
Women	18(38.3)	23(51.1)
Education years, years	10.26(3.30)	4.38(4.15)
Income, MYR	1720.21(1516.42)	830.32(494.07)
Marital status		
Married	34(72.3)	17(37.8)
Not married	3(6.4)	14(31.1)
Divorced	10(21.3)	14(31.1)
Working status (past)		
Government	24(51.1)	3(6.7)
Private	5(10.6)	6(13.3)
Own job	10(21.3)	23(51.1)
Not working	8(17.0)	13(28.9)
Living status		
Husband/Wife only	14(29.8)	10(22.2)
Husband/Wife and children	21(44.7)	17(37.8)
Children only	9(19.1)	13(28.9)
Relatives	1(2.1)	1(2.2)
Alone	2(4.3)	3(6.7)
Others	0(0.0)	1(2.2)

*Significant using Independent-t-test/Chi-Square test



p-value

0.033* 0.216

< 0.001* < 0.001* 0.001*

< 0.001*

0.216

• Participants in the intervention group: younger, had higher education & better income (p<0.05)

he intervention group, better muscle strength, muscle performance, TUG, 2-min step test & gait Baseline Findings Participants

Weight, kg	64.21(11.66)	61.58(8.49)	0.1	.09				
Height, cm	157.50(8.12)	156.77(7.18)	0.3	24				
BMI, kg/cm2	25.93(4.51)	25.12(3.52)	0.1	.68	Participo	Participants in the control group:		
Waist circumference, cm	88.73(10.92)	88.52(10.16)		62	low calf circumference & mu			
Hip circumference, cm	94.43(8.79)	94.74(6.24)	0.3	80		muss(p<0.03)		
MUAC, cm	28.96(3.40)	28.14(2.90)	0.0	87				
Calf circumference, cm	34.69(3.69)	33.46(2.71)	0.03	30*				
Muscle mass (%)	27.63(3.90)	26.17(3.52)	0.03	37*				
Fat mass (%)	30.95(7.27)	33.22(6.62)	0.0	58				
		Variable		Interv	ention (n=47)	Control (n=45)	p-value	
				Ρ	hysical fitness			
		Muscle strength, kg		25.02(6.89)		21.24(7.50)	0.007*	
 Participants in the interve 	ntion group: better	SPPB, score		9.62(1.81)		8.59(2.54)	0.005*	
muscle strength, SPPB scc	pre, TUG, 2-min step	Back scratch, cm		-11	.83(10.11)	-13.01(10.67)	0.301	
test & gait speed (p<0.05)		TUG, seconds		11	24(2.75)	13.86(3.46)	< 0.001*	
		2-min step test, ste	eps	60.	92(34.71)	49.29(25.70)	0.028*	
		Chair-sit-and reach,	cm	-1.58(13.18)		-4.23(11.79)	0.154	
Significant using Independent-t-test		Gait speed, second	ds	4.	74(1.19)	6.03(1.55)	<0.001	



Participants in the intervention group: better muscle strength muscle performance, TUG, 2-min step test & gait speed (p<0.05) Baseline Findings

Variable	Intervention (n=47)	Control (n=45)	p-value						
Dietary Intake (macronutrient)									
Energy (kcal)	1391.40(379.94)	1497.64(507.57)	0.131						
Protein, g	57.59(18.36)	64.66(28.21)	0.078						
Carbohydrate, g	198.46(63.75)	203.98(68.76)	0.345						
Fat, g	40.51(13.57)	46.61(23.54)	0.065						
Dietary cholesterol, mg	207.01(146.64)	242.20(196.15)	0.165						
Saturated fat, g	9.97(4.76)	11.15(8.66)	0.208						
MUFA, g	7.30(3.89)	7.40(4.86)	0.455						
PUFA,g	4.03(2.27)	4.13(2.76)	0.423						
Fiber,g	3.42(1.91)	3.15(1.60)	0.230						
Sugar, g	32.45(24.24)	30.58(25.23)	0.359						

• No significant differences in macronutrient intake between both groups

*Significant using Independent-t-test



Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)
 Baseline Findings

Variable	Intervention (n=47)	
	Dietary Intake (vitamin)	
Vitamin A, RE	553.90(305.27)	
Vitamin C, mg	68.24(47.79)	
Vitamin D, µg	2.34(2.15)	
Vitamin E, mg	2.62(1.40)	
Thiamin, mg	0.65(0.23)	
Riboflavin, mg	1.14(0.45)	
Niacin, mg	11.52(4.28)	
Pyridoxine, mg	0.71(0.45)	
Folate, μg	56.23(26.34)	
Cobalamin, µg	1.87(2.91)	
Biotin, μg	3.22(2.13)	
Pantothenic acid, mg	0.29(0.52)	
Vitamin K, µg	9.35(13.92)	

• No significant differences in vitamin intake between both groups

*Significant using Independent-t-test



p-value
0.268
0.325
0.103
0.466
0.282
0.194
0.470
0.108
0.206
0.212
0.274
0.113
0.095

Participants in the intervention group, better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05) Baseline Findings

Variable	Intervention (n=47)	Control (n=45)	p-value
	Dietary Intake (mineral)		
Sodium, mg	3141.83(1447.45)	3331.54(2447.35)	0.325
Potassium, mg	1313.82(674.74)	1395.54(604.95)	0.232
Calcium, mg	412.23(197.07)	471.46(352.91)	0.160
Iron, mg	10.94(5.19)	11.06(4.86)	0.452
Phosphorus, mg	812.11(714.03)	869.38(373.17)	0.216
Magnesium, mg	84.63(43.88)	84.41(59.25)	0.492
Zinc, mg	2.95(1.60)	2.38(1.48)	0.039
Copper, mg	0.37(0.19)	0.35(0.21)	0.290
Manganese, mg	0.40(0.48)	0.39(0.47)	0.983
Selenium, µg	25.38(16.12)	18.85(21.83)	0.498
Molybdenum, μg	1.47(1.81)	1.34(1.23)	0.369

• No significant differences in mineral intake between both groups

*Significant using Independent-t-test



Dietary Under-Over Reporting

Category		Groups									
	Intervention (%)										
	Baseline	6 th week	12 th week	Baseline 6 th week		12 th week					
Under	76.6	70.0	64.4	71.1	55.3	48.6					
Normal	19.1	22.5	28.9	20.0	28.9	35.1					
Over-report	4.3	7.5	6.7	8.9	15.8	16.2					

Compliance of the Study Groups

Response rate: Intervention: 95.74%, Control: 84.44% Supplement adherence rate: Intervention: 93.14% , Control: 72%

• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

Results: Comparison of changes in sarcopenia indices in CC and placebo arm from baseline to the end of follow-up (12 weeks)

Parameter	Treatment (n=47)	Control (n=45)	Gr	oup effe	ct	Time effect			Interaction effect		
			Partial eta	Power	p-value	Partial eta	Power	p-value	Partial eta square	Power	p-value
			square			square					
		Sar	copenia Ir	ndices (p	rimary ou	tcome)					
Balance test,			0.006	0.103	0.497	0.102	0.972	<0.001*	0.027	0.408	0.126
score											
Baseline	3.68(0.81)	3.40(1.16)									
6 th week	3.56(1.01)	3.51(1.08)									
12 th week	3.93(0.36)	3.51(0.97)									
Chair stand			0.030	0.341	0.120	0.017	0.272	0.266	0.000	0.054	0.963
test, sec											
Baseline	14.05(3.40)	15.04(3.69)									
6 th week	13.89(3.37)	14.56(3.27)									
12 th week	13.69(3.16)	14.20(2.81)									
Muscle			0.008	0.121	0.434	0.014	0.249	0.321	0.028	0.440	0.108
strength, kg											
Baseline	25.02(6.89)	21.24(7.49)									
6 th week	24.62(6.87)	23.35(6.43)									
12 th week	25.26(6.57)	23.78(6.98)									

• No significant differences in sarcopenia indices between both groups

• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

Results: Comparison of changes in sarcopenia indices in CC and placebo arm from baseline to the end of follow-up (12 weeks)

Parameter	Treatment (n=47)	Control (n=45)	Gr	oup effe	ct	Time effect			Interaction effect		
			Partial	Power	p-value	Partial	Power	p-value	Partial eta	Power	p-value
			eta			eta			square		
			square			square					
		Sar	copenia Ir	ndices (pr	rimary ou	tcome)	0 0 7 0	0.000	0.005	0 5 4 0	0.070
Fat mass, %			0.002	0.068	0.689	0.002	0.073	0.826	0.035	0.512	0.070
Baseline	30.95(7.27)	33.23(6.44)									
6 th week	31.63(7.36)	31.41(5.86)									
12 th week	32.27(6.69)	31.89(6.41)									
SMI, kg/m2			0.005	0.096	0.526	0.008	0.160	0.482	0.021	0.313	0.186
Baseline	7.08(1.18)	6.51(0.73)									
6 th week	6.98(1.08)	6.67(0.87)									
12 th week	6.82(1.08)	6.68(0.97)									
Calf			0.010	0.147	0.364	0.028	0.464	0.102	0.011	0.196	0.405
Circumference											
,cm	34.69(3.69)	33.43(2.59)									
Baseline	3/1 30(3 65)	33 /1(2 53)									
6 th week		33.41(2.33)									
12 th week	34.18(3./1)	33.04(2.29)									

• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

Results: Comparison of changes in physical fitness in CC and placebo arm from baseline to the end of follow-up (12 weeks)

Parameter	Treatment (n=47)	Control (n=45)	Gr	oup effe	ct	Time effect			Interaction effect		
			Partial	Power	p-value	Partial	Power	p-value	Partial eta	Power	p-value
			square			square			Square		
			Phy	ysical fi	tness						
Back scratch,											
cm	-11.83(10.11)	-13.01(10.67)	0.000	0.052	0.889	0.009	0.172	0.483	0.013	0.211	0.392
Baseline	-10.56(8.96)	-11.17(8.90)									
6 th week	-8.07(8.16)	-10.49(8.46)									
12 th week											
Chair sit and			0.029	0.334	0.125	0.005	0.110	0.684	0.009	0.165	0.490
reach, cm	-1.58(13.18)	-4.17(11.04)									
Baseline	-2.85(12.18)	-6.31(12.69)									
6 th week	-1.35(5.20)	-5.52(9.82)									
12 th week											

• No significant differences in physical fitness between both groups

• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

Results: Comparison of changes in dietary intake in CC and placebo arm from baseline to the end of follow-up (12 weeks)

Parameter	Treatment (n=47)	Control (n=45)	Gı	Group effect Time effect		;t	Interaction effect				
			Partial eta square	Power	p-value	Partial eta square	Power	p-value	Partial eta square	Power	p-value
Dietary Intake											
Fat, g			0.050	0.522	0.045*	0.008	0.155	0.514	0.000	0.050	0.999
Baseline	40.51(13.57)	46.61(23.54)									
6 th week	37.75(11.01)	44.99(20.38)									
12 th week	39.11(13.79)	43.98(15.63)									
Cholesterol, mg			0.534	0.051	0.042*	0.009	0.168	0.495	0.000	0.051	0.996
Baseline	207.01(146.64)	242.20(196.15)									
6 th week	208.59(129.01)	261.07(195.16)									
12 th week	210.59(137.68)	237.43(171.17)									

No interaction effect in macro and micronutrient intake were observed between baseline till 12th week follow-up for both groups

-Significant group effect for fat & cholesterol were observed at each time point

• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05) DISCUSSION

This study demonstrates that 12-week supplementation with *Cosmos caudatus* had no significant changes in sarcopenia indices, physical fitness and dietary intake among subjects with probable & definitive sarcopenia in Kelantan



• Findings from systematic review showed that intervention based on antioxidant supplementation in combination with vitamin D and protein, as well as increasing fruits & vegetables intake improved muscle strength and physical function

• (Besora-Moreno et al 2022)







• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

DISCUSSION





(*Ref: Park et al (2017*)

• Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

DISCUSSION





(*Ref: Park et al (2017*)

Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05) SUMMARY OF PAST INTERVENTION STUDIES

Author (year), Country	Intervention	Duration	Subjects	Main findings
Harper et al (2020), USA	Exercise & 500 mg Resveratrol, Exercise & 1000mg resveratrol, Exercise & placebo (3 arms)	12 weeks	Older adults with physical function limitations (n=60)	-Exercise + 1000 mg: 6 min walk test score was higher than other
Mafi et al (2019), Iran	Resistance training (RT) & Epicatechin (EP), RT, EP, Placebo (4 arms) Dose EP: 1 mg pure epicatechin per 1 kg BW	8 weeks	Men with sarcopenia (n=62)	-EP+RT: improvement in leg press, chest press as compared to other groups
Kim et al (2023)	Exercise + tea catechin, exercise, tea catechin, health education (4arms) Dose: 350ml/day of tea catechin	3 months	Women with sarcopenia (aged 75 yrs and older)	Combined intervention of exercise + tea catechin: improved leg muscle mass and walking



CONCLUSION

Cosmos caudatus supplement alone provides less benefit in improving health outcomes among sarcopenic older adults.



Resistance exercise (2-3x/week) Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)
 Current Work

CC supplement

IGF-P13K-AKT-mTOR

Gut microbiota

This study is supported by the Ministry of Higher Education Malaysia for the Fundamental Research Grant Scheme (FRGS) with project code: FRGS/1/2022/SKK06/USM/02/1.

Acknowledgement

This study is supported by the Fundamental Research Grant Scheme (FRGS): FRGS/1/20202/SKK06/USM/02/1 from the Ministry of Higher Education Malaysia

Participants in the intervention group: better muscle strength, muscle performance, TUG, 2-min step test & gait speed (p<0.05)

References

- 1. Rafi M, Hayati F, Umar AH, Septaningsih DA, Rachmatiah T. LC-HRMS-based metabolomics to evaluate the phytochemical profile and antioxidant capacity of Cosmos caudatus with different extraction methods and solvents. Arabian Journal of Chemistry. 2023 Sep 1;16(9):105065.
- 2. Moshawih S, Cheema MS, Ahmad Z, Zakaria ZA, Hakim MN. A comprehensive review on Cosmos caudatus (Ulam raja): pharmacology, ethnopharmacology, and phytochemistry. International Research Journal of Education and Sciences. 2017;1(1):14-31.
- 3. Bagherniya M, Mahdavi A, Shokri-Mashhadi N, Banach M, Von Haehling S, Johnston TP, Sahebkar A. The beneficial therapeutic effects of plant-derived natural products for the treatment of sarcopenia. Journal of Cachexia, Sarcopenia and Muscle. 2022 Dec;13(6):2772-90.
- 4. Park SS, Kwon ES, Kwon KS. Molecular mechanisms and therapeutic interventions in sarcopenia. Osteoporosis and sarcopenia. 2017 Sep 1;3(3):117-22.
- 5. Harper SA, Bassler JR, Peramsetty S, Yang Y, Roberts LM, Drummer D, Mankowski RT, Leeuwenburgh C, Ricart K, Patel RP, Bamman MM. Resveratrol and exercise combined to treat functional limitations in late life: a pilot randomized controlled trial. Experimental gerontology. 2021 Jan 1;143:11111.
- 6. Mafi F, Biglari S, Afousi AG, Gaeini AA. Improvement in skeletal muscle strength and plasma levels of follistatin and myostatin induced by an 8-week resistance training and epicatechin supplementation in sarcopenic older adults. J Aging Phys Act 2019;27:384–391.
- 7. Kim H, Suzuki T, Saito K, Yoshida H, Kojima N, Kim M, et al. Effects of exercise and tea catechins on muscle mass, strength and walking ability in community-dwelling elderly Japanese sarcopenic women: a randomized controlled trial. *Geriatr Gerontol Int* 2013;13:458–465.
- 8. Wiedmer P, Jung T, Castro JP, Pomatto LC, Sun PY, Davies KJ, Grune T. Sarcopenia-Molecular mechanisms and open questions. Ageing research reviews. 2021 Jan 1;65:101200.
- 9. Nazri NS, Vanoh D, Leng SK. Determinants of Poor Diet Quality among Elderly with Low Socioeconomic Status. Jurnal Gizi dan Pangan. 2023 Nov 30;18(3):147-56. 10. Iskandar I, Joanny A, Azizan A, Justine M. The Prevalence of Sarcopenia and Its Impact on Quality of Life in Elderly Residing in the Community. Malaysian Journal of Medicine & Health Sciences. 2021 Jun 2;17(3).
- 11. Ranee R, Shahar S, You YX, Ajit Singh DK, Mohamed Sakian NI. Prevalence and Risk Factors of Sarcopenia Among Community Dwelling Older Adults in Klang Valley. Malaysian Journal of Medicine & Health Sciences. 2022 Jan 1;18(1).
- 12. Nazri NS, Vanoh D, Leng SK. Malnutrition, low diet quality and its risk factors among older adults with low socio-economic status: a scoping review. Nutrition Research Reviews. 2021 Jun;34(1):107-16.
- 13. Papadopoulou SK. Sarcopenia: a contemporary health problem among older adult populations. Nutrients. 2020 May 1;12(5):1293. 14. Le NH, Kim CS, Park T et al. 2014. Quercetin protects against obesity-induced skeletal muscle inflammation and atrophy. Mediators Inflammation doi: 10.1155/2014/834294.
- 15. Li P, Liu A, Xiong W et al. 2020. Catechins enhance skeletal muscle performance. Critical Reviews in Food Science and Nutrition 60(3):515–528. 16. Nikawa T, Ulla A, Sakakibara I. 2021. Polyphenols and their effects on muscle atrophy and muscle health. Molecules 26(4887).

- 17. Besora-Moreno M, Llauradó E, Valls RM, Tarro L, Pedret A, Solà R. Antioxidant-rich foods, antioxidant supplements, and sarcopenia in old-young adults 55 years old: A systematic review and meta-analysis of observational studies and randomized controlled trials. Clinical Nutrition. 2022 Oct 1;41(10):2308-24.
- 18. Lohne-Seiler H, Kolle E, Anderssen SA, Hansen BH. Musculoskeletal fitness and balance in older individuals (65-85 years) and its association with steps per day: a cross sectional study. BMC geriatrics. 2016 Dec;16:1-1.
- 19. Bohannon RW, Crouch RH. Two-minute step test of exercise capacity: systematic review of procedures, performance, and clinimetric properties. Journal of Geriatric Physical Therapy. 2019 Apr 1;42(2):105-12.
- 20. Mayorga-Vega D, Bocanegra-Parrilla R, Ornelas M, Viciana J. Criterion-related validity of the distance-and time-based walk/run field tests for estimating cardiorespiratory fitness: a systematic review and meta-analysis. PloS one. 2016 Mar 17;11(3):e0151671.
- 21. Chen LK, Woo J, Assantachai P, Auyeung TW, Chou MY, Iijima K, Jang HC, Kang L, Kim M, Kim S, Kojima T. Asian Working Group for Sarcopenia: 2019 consensus update on sarcopenia diagnosis and treatment. Journal of the American Medical Directors Association. 2020 Mar 1;21(3):300-7.
- 22. You YX, Shahar S, Rajab NF, Haron H, Yahya HM, Mohamad M, Din NC, Maskat MY. Effects of 12 weeks Cosmos caudatus supplement among older adults with mild cognitive impairment: a randomized, double-blind and placebo-controlled trial. Nutrients. 2021 Jan 29;13(2):434.
- 23. Cheng SH, Barakatun-Nisak MY, Anthony J, Ismail A. Potential medicinal benefits of Cosmos caudatus (Ulam Raja): A scoping review. Journal of research in medical sciences. 2015 Oct 1;20(10):1000-6.
- 24. Foo LH, Wen YS, Kadir AA. Assessments of sarcopenia and its associated factors in community-dwelling middle-aged and older Chinese adults in Kelantan, Malaysia. Scientific Reports. 2023 May 9;13(1):7498.
- 25. Ziaaldini MM, Marzetti E, Picca A, Murlasits Z. Biochemical pathways of sarcopenia and their modulation by physical exercise: a narrative review. Frontiers in medicine. 2017 Oct 4;4:167.
- 26. Vafa M, Abiri B, Dehghani M. The association of food intake and physical activity with body composition, muscle strength, and muscle function in postmenopausal women. Clinical and Preclinical Models for Maximizing Healthspan: Methods and Protocols. 2020:363-71.
- 27. Ramoo K, Hairi NN, Yahya A, Choo WY, Hairi FM, Ismail N, Peramalah D, Kandiben S, Ali ZM, Ahmad N, Abdul Razak I. Sarcopenia and All-Cause Mortality Risk in Community-Dwelling Rural Malaysian Older Adults. Asia Pacific Journal of Public Health. 2024 Mar;36(2-3):225-31.
- 28. Schoufour JD, Tieland M, Barazzoni R, Ben Allouch S, Bie JV, Boirie Y, Cruz-Jentoft AJ, Eglseer D, Topinková E, Visser B, Voortman T. The relevance of diet, physical activity, exercise, and persuasive technology in the prevention and treatment of sarcopenic obesity in older adults. Frontiers in nutrition. 2021 May 24;8:661449.