

Supporting materials for

A Review of The Functional Activities of Chia Seed and Mechanisms of Action Related to Molecular Targets

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Table S1

Composition of chia seed and the health promoting effects

Composition	Major component	Extract method	Processing	Functional properties	Way of action	
CSO	ALA ¹⁻³	cold-pressing ¹	microwave ³	anti-tumor ⁸	contributes to the expression of nuclear transcription factors ¹⁴	
		heat-pressing ¹⁻²	roasting ⁶	modulate blood pressure	normalize the operation of calcium and sodium ion channels ⁹	
		enzyme assisted solvent ³	encapsulation ⁷	regulate glucose metabolism ¹⁰	adjust glycolysis ¹⁰	
		microwave assisted solvent ³		immunomodulation ¹¹	contribute to proliferation of immunoglobulin E and thymocytes ¹⁵	
		ultrasonic assisted solvent ⁴		reinforcement of blood lipid regulation ¹²	reduce the production of LDL and increase the content of HDL ¹⁶	
		supercritical fluid ⁵		relieve metabolic syndrome ¹³	improve insulin guidance and delivery ¹⁷	
		Soxhlet extraction ²				
		oligosaccharides ¹⁸	water ¹⁹	delivery ²²⁻²³	moisturize intestines ²³	
-----					cholesterol-lowering ²⁸	
-----			ion solution ²⁰		regulate blood sugar ²⁴	
-----					hypoglycemic ²⁴	

		ultrasonic assist ²¹	anti-bacteria ²⁵⁻²⁷	inhibiting cell transduction ²⁹
CPI	albumin, globulin, prolammin, glutelin ³⁰⁻³²	alkali-solution& acid-isolation ³³⁻³⁴ ultra-pure water ³⁵ 1.0mol/L NaCl ³⁶ 0.2% NaOH ³³	hydrolysis ³⁷⁻⁴⁰ increase satiety ⁴¹	anti-hypertension ⁴⁰ ACE activity inhibition ⁴⁰ blocking elevation of GLP-1 level caused by DPP-IV ⁴¹ 5-LOX, COX1, COX2, iNOS inhibition ⁴¹ relieve atherosclerosis ⁴¹⁻⁴² reduces gene expression in adipogenesis ⁴¹⁻⁴²
Phenol & flavonoids	rosmarinic acid, caffeic acid, salicylic acid, flavonoids, myricetin, quercetin ⁴¹⁻⁴⁴	alkaline ⁴⁵ concentrate methanol-ethanol 70% (v/v) ⁴² mixtures of moderately polar solvents ⁴⁶⁻⁴⁷ water: acetone (1: 1) ⁴⁸ water: ethanol: acetone (1:2:3, v/v/v) ⁴⁶	maintain homeostasis in the blood environment ⁴³ anti-oxidation ⁴¹⁻⁴³ reduce fat absorption ⁴³ DNA oxidation protection ⁴³	curbing the activity of pancreatic lipase and α-glucosidase ⁴³ free radicals (DPPH, ABTS, OH ⁻) scavenging β-carotene bleaching inhibition ⁴¹⁻⁴³ inhibitory of pancreatic lipase activity ⁴³ free radical scavenging ⁴¹⁻⁴³

Abbreviations: CSO, chia seed oil; CSM, chia seed gum, CPI, protein isolation; ALA, alpha linolenic acid; LDL, low density lipoprotein; ACE, angiotensin converting enzyme; GLP-1, glucagon peptide-1; DPP-IV, dipeptidyl peptidase IV; 5-LOX, 5-lipoxygenase; COX1, cyclooxygenase-1; COX2, cyclooxygenase-2; iNOS, inducible nitric oxide synthase; DPPH, 2,2-Diphenyl-1-(2,4,6-trinitrophenyl) hydrazyl; ABTS, 2, 2'-azino-bis (3-ethylbenzothiazoline-6-sulfonic acid);

Table S2

Micronutrients and their contents in CS

Component	Name	Content(mg/100g)	Ref.	Name	Content	Ref.
Phenolic acid	polyphenols	51.1-97.5	49-50	rosmarinic acid	92.67	50
	protocatechuic acid	74.71	50	gallic acid	1.15	50
	chlorogenic acid	0.468-23.5	49-50	caffeic acid	0.3-15.6	50-52
Flavonoid	myricetin	0.95	49, 52	quercetin	0.6-26.8	49, 52
	kaempferol	0.017-50.9	50, 52			
Isoflavone	daidzin	0.66	52	genistein	0.51	49
	genistin	0.34	52	glycerin	0.14	49
	glycitein	0.05	49			
Vitamin	vitamin A	54(IU)	52	thiamine	0.62	52
	riboflavin	0.17	49,52	nicotinic acid	8.83	52,53
	folic acid	0.049	52	ascorbic acid	1.6	52
	vitamin E	0.5-42.7	50,52			
Mineral	sodium	16-22.37	6,53	magnesium	335-403	6,49
	phosphorus	407-870	49, 53	potassium	407-870	6,49

calcium	580-829.21	49	manganese	5.74	53
iron	7.72-24.4	49, 52	cobalt	0.035	49
copper	0.924-1.45	52	zinc	4.58-9.11	53
selenium	0.055.2-0.078	49,52	molybdenum	0.2	6,52

Note: IU, international unit

Table S3. Amino acid sequence and the active site of CPI compositions

Origin	Amino acid sequence	Access	Pathway/substance	Biological activity	Ref.
Total protein of chia seed	YACLKV KLKKNL LPVFGLAAEGNVTYLH CNSPGPHDVALDQ RMVLPEYELLYE	<i>in silico</i> digestion digestion	- Mmp-2, PPAR- γ , adiponectin PPAR- α NF- κ B	antibacterial mediate adipocyte differentiation, relieve inflammation prevent adipogenesis, inhibition inflammation	54 41 55
Ribulose bisphosphate Carboxylase	VPILIQIRPFPEPKPQPWVDVF VMVNVPYPAIPSLPLVEVKVTIH PGVGVLVWVA	SGID	DPP-4, α -glucosidase	antidiabetic activity	56-57
Oleosin	SPSLPLPHSHSWVDVFVG VKVSVTVIAIAVL				
Fatty acid (F.A) desaturase 3 isoform 2	PAVPLIAIHILRPMNPQPTPW PYSFSHSKVDVFVHKVTVWW YSLPLSWSYVEVGVAVL				
F.A desaturase 3 isoform 1	PAVPLIRPMPEPNPQPTPWPSFS HSKVDVKVTWWVYIASLPLIHL SWSYVEVGVLVA				
F.A desaturase 7 isoform 1	VRINIRPFPGSHVEVFVKVMVN VTWWVYIHLPKMPQWPWSFSW SYVGVTAVLPL				
F.A desaturase 8	PAILPFPEPNSHSYVEVFVKVMV QVTWWVYVASLVRIHINIQPMPQ PWSWVGPKVLSFPL				
F.A desaturase 2 isoform 2	PASPIAPPGIHPNPYSHSWVDVEV GVNVTVWSLSFSYVKVLVQVYVA				

	PLPHVFIL
F.A desaturase 2 isoform 1	PASPIAPPGIHIQPKPQQYSHSWVD VEVGVNQVWVSLPHSFSYVKVYY APLVFVLIL
Monoacylglycerol	VAPAIAIQIRPKPMPTSFSKSYVEHV
Acyltransferase	KVLVTSLVRPEPLVFVG
Eukaryotic translation	VPLIMIRIWPFPGPQSHVDVFVG
Initiation factor	HVYIAPLPEIHILIQPNVLVQSLVNSK
S-adenosylmethionine	SLIHIMIRPFPHSKVMVTPAPEIAVRP
Decarboxylase	GVDVFVASFSYVEVYILVL
Tubulin beta chain	VASLVRPLILINPTPYVHVVKVNVQVT VYIQIRPGSKVDVFVGVEVL
Peptidyl-prolyl cis-trans	PHPKPMPTSVDVKVMVQVTI
Isomerase	AVRVEVFVGAIL
Serine threonineprotein	VAPAIAVRIHIRPGPMPNVDVM
Phosphatase	VQVWVYPLVEVFVKVTL
Elongation factor 1-alpha	PAVRPPLIHIMPMPESFSHVDVQVYY PIAPLPTSKVTVAPKVLPGVEVGVK
Glyceraldehyde-3-Phosphate	PAIAVRPLIHILPEPFPMFSKSYVLV
Dehydrogenase	SVTINVFVHVVKVNVAVEVGVD
Actin	PAVPVRIHIRIWPPEPGPHPKVVDVE VFVHVVKVMVWPLIMIQPSSKVAI ASLVGVLIL
Clathrin adaptor complex	PPVAPAIPPLINIQPFPGPQSWVDV HILIMIRPTSF SYVFVGVKVVWVY VEVLVNRVT
α -tubulin 7 protease	VASKVDVFVHVG

FtsH protease	IAPGSHVDVEVHVLMVNILVASKVKVT				
7S globulin	PAIPASPIASLPLIQIRIWPQPHQSYVEVK				
	VMVNWWVRPFPTSFVDFVPEINSKVLILVA				
11S globulin	SLPLP E PFP G PKP M TPY V D V EVG V MVQV	<i>in silico</i>	molecular docking	ACE, DPP-4 activity inhibition, renin inhibitor, antioxidant,	58
	YPAILIQIRWSYVFPHQAVAVLVR				
TAQEPTIRF		<i>in silico</i>	molecular docking	ACE, DPP-4 activity inhibition, antithrombosis	58
PGLTIGDTIPNL		size exclusion	inhibit the activities of collagenase, hyaluronidase, tyrosinase	ACE, DPP-4 activity inhibition	59
LSLPNYHPNPRL		chromatography	and elastase	ACE activity inhibition, regulation of glucose absorption	
LIVSPLAGRL				ACE, DPP-4 activity inhibition, glucose uptake modulation, activate proteolysis mediated by ubiquitin	
				skin anti-aging	
IVSPLAGRL				ACE, DPP-4 activity inhibition, glucose uptake modulation, activate	
APHWYTN		size exclusion	inhibit the activities of collagenase, hyaluronidase, tyrosinase	proteolysis mediated by ubiquitin	59
DQNPRSF		chromatography	and elastase	skin anti-aging	
GDAHWAY		/	antibacterial		
GDAHWTY					
GDAHWVY					
GFEWITF					
KKLKRVYV					
KADVPGKK		/		antibacterial	59
KGDVIAIR				combined with CCR2	56-60
LKQGDVIAIR		simulated gastrointestinal digestion		antiphlogistic	
GNIFRGL					
FQKGIIQVVNHK					
IGTPGKGIL					
QKGDVIAIR					
KQGDVIAIR					

-----	QKGDVIAIRA				
	KQGDVIAIR				
	DVPGLKK				
	FAFFEFELLFAFFT	simulated gastrointestinal	combined with ICAM1	relieve inflammation	
	LPGPPATF	digestion			56-57

Abbreviations: Mmmp-2, Matrix metalloproteinases-2; SGID, simulated gastrointestinal digestion; PPAR- γ , peroxisome proliferator activated receptor gamma; PPAR- α , peroxisome proliferator activated receptor alpha; NF- κ B, markers nuclear factor κ B; DPP-IV, dipeptidyl peptidase IV; ACE, angiotensin converting enzyme; CCR2, C-C motif chemokine receptor 2; ICAM, intercellular cell adhesion molecule.

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