

CONSTITUENT COMPOSITION AND BIOLOGICAL ACTIVITY OF ESSENTIAL OIL FROM *Artemisia sublessingiana*

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Artemisia sublessingiana Krasch. ex Poljakov (Asteraceae) is a perennial herbaceous plant that grows in eastern, central, southern, and southeastern Kazakhstan [1].

Raw material for the studies was collected in the budding phase during an expedition in the third decade of August 2019 in the vicinity of Zhezqazghan (Karaganda Region). A specimen (2019.08.28.04.01) is preserved in the herbarium of the Biology-Geography Faculty of E. A. Buketov Karaganda University.

Previously, flavonoids (isorhamnetin-3-*O*-rutinoside and 5,7,4'-trihydroxy-6,3'-dimethoxyflavone) and a sesquiterpene lactone (arsubin) [1, 2]; the flavonoids eupatilin, 3',4'-dimethoxyluteolin, 5,7,3'-trihydroxy-6,4',5'-trimethoxyflavone, hispidulin, apigenin, and velutin and the sesquiterpene lactone 8 α ,14-dihydroxy-11,13-dihydromelampolide [3] were isolated from this plant. Compounds isolated by us were tested *in silico* against COVID-19 main protease enzyme (M^{PRO}). Flavonoid compounds turned out to be highly promising with respect to the discovery of drugs for the COVID-19 pandemic [3].

The contents of the valuable constituent santonin in various *Artemisia* species were reported [4]. It was found that *A. sublessingiana* contained ~0.16 g of santonin per 100 g of air-dried raw material. The elemental composition of ash from *A. sublessingiana* and the fatty-acid and amino-acid compositions were previously studied [5].

According to the literature, the essential oil composition of *A. sublessingiana* has not been previously studied. In continuation of the determination of essential oil compositions of *Artemisia* species [6–8], it was studied by us using gas-chromatography–mass-spectrometry (GC-MS).

Essential oil of *A. sublessingiana* was obtained from various plant parts (aerial part, seeds, stems) by steam distillation in a Clevenger apparatus using hexane as a trap [9].

GC-MS analysis of *A. sublessingiana* essential oils was performed under conditions analogous to those in the literature [10] using a Restek Rxi[®]-1ms capillary column (0.25 mm \times 30 m \times 0.25 μ m). Constituents were identified using the NIST 2014 database. Table 1 presents the constituent composition of the *A. sublessingiana* essential oils. The main constituents (> 3.0%) of the essential oils from the aerial part were 3-thujanone (18.9%), chrysanthenone (3.6%), camphor (17.9%), *cis*-chrysanthenyl acetate (31.3%), and nerol acetate (3.1%); from seeds, 1,8-cineol (12.0%), β -thujone (31.8%), 3-thujanone (18.4%), and camphor (14.8%); from stems, 1,8-cineol (11.8%), β -thujone (15.5%), 3-thujanone (34.3%), and camphor (17.2%).

Literature data were used to identify obscure constituents of the essential oil such as artedouglasia oxides A, C, and D and laciniata furanones E, F, and H [11, 12].

The cytotoxic activity of the essential oils from *A. sublessingiana* was studied using *Artemia salina* larvae and the literature method [13]. DMSO was used as the solvent. The antibiotic actinomycin D or staurosporine was used as a control. The experiments found that essential oil obtained from the aerial part at all concentrations exhibited cytotoxicity with lethality of larvae reaching 96%. Essential oil from seeds was cytotoxic at all concentrations with lethality of 75–96%. Essential oil from stems at concentrations of 10 and 5 mg/mL was cytotoxic with lethality of larvae reaching 96% while cytotoxicity was not found at a concentration of 1 mg/mL.

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TABLE 1. Constituent Composition of Essential Oil from *A. sublessingiana*

Constituent	RII	Essential oil content, %		
		stems	aerial part	seeds
7-Methyl-3,4-octadiene	843	0.2		0.1
Ethyl 3-methylbutanoate	848	0.1	0.2	0.1
Tricyclene	915	0.2	0.1	0.1
α -Pinene	925	0.2	1.2	0.3
Camphene	939	2.2	1.8	2.3
Sabinene	963	0.7	0.2	0.8
β -Pinene	966	0.2	0.1	0.2
Mesitylene	983		0.2	
Pseudocumene	983	0.1		
Unk. 1	985	0.1		
α -Terpinene	1008	0.2		0.2
1,2,4-Trimethylbenzene	1010		0.2	
α -Cymene	1016	0.7	0.4	0.8
α -Limonene	1020		0.1	
β -Phellandrene	1021			0.1
1,8-Cineol	1027	11.8	1.1	12.0
γ -Terpinene	1051	0.4	0.1	0.3
<i>cis</i> -Sabinene hydrate	1068			0.1
Unk. 2	1085		0.1	
Filifolone	1097		0.6	0.1
β-Thujone	1106	15.5	1.5	31.8
3-Thujanone	1115	34.3	18.9	18.4
Chrysanthenone	1118	0.6	3.6	0.6
α -Campholenal	1121		0.1	
<i>cis</i> -2-Menthenol	1121			0.5
<i>trans</i> -Pinocarveol	1135		0.9	0.3
<i>cis</i> -Sabinol	1136			0.1
Camphor	1141	17.2	17.9	14.8
Unk. 3	1144	0.5		
Pinocarvone	1153	0.5	0.9	0.3
<i>cis</i> -Chrysanthenol	1157	0.5	0.3	0.3
Isothujol	1159	0.1		
1,3,4-Trimethyl-3-cyclohexenyl-1-carboxaldehyde	1160	0.1		
Borneol	1163	1.5	0.3	0.9
Terpinen-4-ol	1171	1.2	0.4	0.9
<i>p</i> -Cymen-8-ol	1180	0.2	0.2	0.1
α -Terpineol	1187			0.4
Myrtenal	1190	0.2	0.2	0.1
Myrtenol	1193	0.6	0.2	
3-Methylbut-3-enyl (<i>E</i>)-2-methylbut-2-enoate	1195		0.1	
<i>trans</i> -Piperitol	1201			0.2
Verbenone	1203	0.1		
Unk. 4	1207	0.3		
<i>p</i> -Cymenol	1231	0.1		
Ciminal	1235	0.2		0.1
Carvone	1241	0.3	0.3	0.2
Piperitone oxide	1249	0.1		
Piperitone	1253	0.6		0.2
<i>cis</i>-Chrysanthenol acetate	1254		31.3	
α -Citral	1265		0.2	
Bornyl acetate	1280	0.1	0.5	
Unk. 5	1281		0.1	
<i>trans</i> -Sabinyl acetate	1289	0.8	0.5	0.5
Methyl <i>cis</i> -cinnamate	1297		0.2	
Thymol	1299			0.1

TABLE 1 (continued)

Constituent	RII	Essential oil content, %		
		stems	aerial part	seeds
Carvacrol	1302	0.2		
Unk. 6	1309	0.1	0.7	0.1
Myrtenyl acetate	1318		0.2	
Hexyl tiglate	1329		0.2	
<i>trans</i> -Dihydrocarvyl acetate	1332			0.1
α -Terpinyl acetate	1341			0.1
<i>cis</i> -Chrysanthenyl propionate	1346		0.1	
Nerol acetate	1379		3.1	
2-(Acetylmethyl)-3-carene	1389		0.5	
<i>cis</i> -Jasmone	1392	0.3		0.2
2-Ethylidene-6-methyl-3,5-heptadienal	1394	0.1	0.8	0.2
Isocaryophyllene	1403			0.1
Aristolene	1454	0.1	0.2	0.1
<i>cis</i> -Muurool-3,5-diene	1459			0.1
Germacrene D	1460		0.3	
2-Isopropenyl-4a,8-dimethyl-1,2,3,4,4a,5,6,7-octahydronaphthalene	1462		0.3	
γ -Gurjunene	1473			0.1
Artedouglasia oxide C	1499	0.2		0.1
Laciniata furanone G	1502	0.2		0.2
Artedouglasia oxide A	1513	0.3		0.1
Laciniata furanone F	1519	0.3		0.1
Laciniata furanone E	1528	0.2		0.2
Laciniata furanone H	1539	0.2		0.1
Spathulenol	1562	0.2	0.2	0.2
Artedouglasia oxide D	1565	0.2		
2-Phenylethyl tiglate	1572		0.5	
Isobutyl pentanoate	1581			0.2
α -Cadinol	1646		0.1	
(1 <i>R</i> ,7 <i>S</i> , <i>E</i>)-7-Isopropyl-4,10-dimethylenecyclodec-5-enol	1676		0.1	
Squalene	2803			0.4
Total		95.3	92.3	91.0

TABLE 2. Antiradical Activity of Essential Oils at Various Concentrations, %

Sample	Concentration, mg/mL				
	0.1	0.25	0.5	0.75	1.0
Butylhydroxyanisole (BHA)	80.82 \pm 4.30	81.23 \pm 2.22	82.30 \pm 3.17	83.08 \pm 2.33	83.88 \pm 2.01
<i>A. sublessingiana</i> (seeds)	20.32 \pm 2.03	20.18 \pm 3.37	20.19 \pm 2.32	22.45 \pm 4.64	32.68 \pm 3.08
<i>A. sublessingiana</i> (stems)	11.96 \pm 3.27	11.99 \pm 3.54	13.22 \pm 3.68	14.31 \pm 3.72	18.55 \pm 3.19
<i>A. sublessingiana</i> (aerial part)	10.38 \pm 4.17	10.08 \pm 3.18	11.31 \pm 3.69	11.98 \pm 2.17	16.03 \pm 2.04

Antiradical activity was determined by the literature method [14, 15]. The tested essential oils showed low antioxidant activity as compared to the standard (butylhydroxyanisole) (Table 2).

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