

SUPPLEMENTARY MATERIAL

Comparative analysis of active ingredients and potential bioactivities of essential oils from *Artemisia argyi* and *A. verlotorum*

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Supplementary material

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Table S1. Composition of the EOs from *Artemisia*.

Compound name	Content (%)				Mass	RI(exp)	RI(lit)	ID ^b
	YP-1 ^a	YP-2 ^a	YP-3 ^a	YP-4 ^a				
neointermedeol	9.69	8.02	4.72	2.79	222.2	1629		MS
caryophyllene	8.73	9.24	1.02	3.8	204.2	1396	1416	RI, MS
caryophyllene oxide	5.93	3.42	17.06	23.7	220.2	1557	1579	RI, MS
<i>cis</i> -chrysanthenol	5.92				152.2	1145	1161	RI, MS
eucalyptol	5.77	11.51			154.1	1013	1030	RI, MS
(<i>-</i>)-4-terpinenol	5.47	5.68			154.2	1159	1182	RI, MS
2-borneol	4.54	10.09	0.52	2.86	154.2	1147	1165	RI, MS
α -terpineol	3.21	3.23			154.2	1172	1187	RI, MS
(<i>-</i>)- <i>cis</i> -verbenol	2.81				152.2	1125	1140	RI, MS
caryophylladienol I	2.42	1.15	3.16	2.42	220.4	1607	1640	RI, MS
<i>cis</i> -carveol	2.35	3.06		0.12	152.2	1199	1226	RI, MS
germacrene D	1.7	1.98			204.2	1456	1478	RI, MS
spainulenol	1.58	1.06			220.4	1552	1574	RI, MS
<i>cis</i> -p-menth-2-en-1-ol	1.49	2.08			154.3	1111	1136	RI, MS
(<i>E</i>)- β -famesene	1.46				204.4	1432	1455	RI, MS
artemisia alcohol	1.38	2.63			154.2	1065	1083	RI, MS
<i>cis</i> -piperitol	1.33	1.4			154.2	1187	1193	RI, MS
chamazulene	1.25	0.53			184.3	1700		MS
humulene	0.97	0.97	0.36	5.87	204.4	1428	1451	RI, MS
β -selinene	0.95	0.82	1.15	1.36	204.4	1461	1483	RI, MS
<i>trans</i> -4-thujanol	0.83	1.45			154.2	1079	1065	RI, MS
γ -terpinene	0.72	0.81			136.2	1040	1057	RI, MS
4-thujanol	0.72				154.3	1048	1068	RI, MS
<i>n</i> -hexadecanoic acid	0.69		1.78		256.4	1933		MS
4(15),5,10(14)-Germacratrien-1-ol	0.68	0.3	1.07	0.73	220.4	1658		MS
phytol	0.57	0.12	1.46	0.31	220.4	2077		RI, MS
(\pm)-phytone	0.51	0.2	3.34	1.8	268.5	1812		RI, MS
selina-4,11-diene	0.49	0.3			218.3	1450	1492	RI, MS
eugenol	0.45	0.84			164.2	1335	1354	RI, MS
bicyclogermacrene	0.43	0.45			204.4	1471	1491	RI, MS
copaene	0.4	0.39			204.4	1352	1375	RI, MS
eremophylene	0.39	0.19			204.4	1564	1507	MS
τ -cadinol	0.34	0.23			222.4	1613	1638	RI, MS
yogomi alcohol	0.33	0.97			154.3	983	1000	RI, MS
γ -pironene	0.31	0.36			136.2	1313	1338	RI, MS
(+)- δ -cadinene	0.3	0.25			204.4	1497	1469	RI, MS
carveol	0.28	0.3			152.2	1209	1200	RI, MS
hexadecanoic acid, methyl ester	0.28				270.5	1893		RI, MS
junenol	0.26	0.2			222.4	1589	1606	RI, MS

(±)-piperitone	0.26	0.21		152.2	1232	1250	RI, MS
perillaldehyde	0.25	0.33		152.2	1252	1268	RI, MS
alloaromadendrene	0.24			204.4	1575	1463	RI, MS
nerolidol	0.23			222.4	1537	1560	RI, MS
(-)carvone	0.23	0.3		150.2	1222	1190	RI, MS
Camphor	0.21			152.2	1263	1277	RI, MS
β -copaene	0.2	0.27		204.4	1403	1429	RI, MS
1-octen-3-ol	0.2	0.34		128.2	963	978	RI, MS
methyl isocostate	0.2			248.4	1754		MS
9,12-octadecadienoic acid, methyl ester	0.18			294.5	2059		MS
α -terpinene	0.18	0.4		136.2	999	1015	RI, MS
linolenic acid, methyl ester	0.18			292.5	2065		MS
o-cymene	0.17			134.2	1007	1020	RI, MS
(-) β -bourbonene	0.17	0.12		204.4	1360	1381	RI, MS
methyleugenol	0.17	0.21		178.2	1382	1400	RI, MS
γ -costol	0.17			220.4	1718		MS
camphene	0.13			136.2	1570	954	RI, MS
cis-chrysanthenol acetate	0.13			194.3	1239	1260	RI, MS
α -bergamotene	0.13		0.84	204.4	1410	1431	RI, MS
epi- β -caryophyllene	0.13		0.2 0.53	204.4	1637		MS
bornyl isovalerate	0.12			238.4	1491	1468	RI, MS
(±)-filiolone	0.12	0.28		150.2	1084	1107	RI, MS
γ -muurolene	0.11		1.15	204.4	1488	1474	RI, MS
geranyl- α -terpinene	0.11			272.5	1919		MS
sclareol oxide	0.11			262.4	1849		MS
(1 α ,3 α ,5 α)-1,5-diethenyl-3-methyl-2-methylene-cyclohexane	0.1			216.2	1676		MS
β -costol		0.21	0.76 0.21	220.4	1737		MS
bornyl acetate		0.28		196.3	1263	1282	RI, MS
isogermacrene D		0.12		218.4	1418	1444	RI, MS
p-cymene		0.41		134.2	1006	1023	RI, MS
isoborneol, isovalerate		0.1		154.3	1490	1520	RI, MS
cis- β -farnesene		0.86	0.42 0.83	204.4	1431		MS
(+)-2-bornanone		5.14		152.2	1124	1132	RI, MS
2,3-dehydro-1,8-cineole		0.28		152.2	971	989	RI, MS
2-ethylidene-6-methyl-3,5-heptadienal		0.16		150.2	1375	1126	MS
1-(1-butynyl) cyclopentanol		0.37		138.2	1167		MS
γ -cadinene		0.14	0.66 0.59	204.2	1618		MS
β -elemene		0.11		204.4	1367	1390	RI, MS
β -bisabolene		0.1	0.35 0.46	204.4	1537	1511	RI, MS
trimethylenenorbornane		0.19		136.2	1135		RI, MS

chrysanthenone	0.35		150.2	1373	1127	RI, MS
alloaromadendrene oxide	0.23	0.5	220.4	1593	1462	RI, MS
α -himachalene	0.15		204.4	1572	954	RI, MS
ledane			204.4	1512	1497	RI, MS
palustrol		0.97	222.4	1540	1566	RI, MS
ledol		2.52	222.2	1575	1568	RI, MS
himbacol		3.56	222.4	1565	1589	RI, MS
mustakone		1.84	218.3	1650		RI, MS
(+)- α -cyperone			218.3	1593	1739	RI, MS
(\pm)- α -curcumene		1.87	202.3	1457	1480	RI, MS
α -gurjunene		3.82	204.4	1384	1407	RI, MS
(+)- α -calacorene		0.56	200.3	1516	1539	RI, MS
(-) α -muurolene			204.4	1474	1496	RI, MS
neoisolongifolene			220.4	1630		MS
aromandendrene	0.1	0.83	204.4	1636		MS
α -copaene			204.4	1351	1375	RI, MS
δ -cadinene			204.4	1497	1511	RI, MS
1,5-cadinadiene			204.4	1618	1495	RI, MS
eudesma-4,11-diene			204.4	1449		RI, MS
β -calacorene	0.19	0.47	200.3	1535	1560	RI, MS
isoledene		0.15	204.4	1466	1377	RI, MS
(-) -calamenene			202.3	1380	1518	RI, MS
α -neocallitropsene	1.04	0.48	206.4	1654		MS
cis-2,2-dimethyl-3-(3-methyl-2-butenyl)-6-methylene		0.36	158.2	1667		MS
cyclohexanemethanol						
(+)- β -eudesmol	6.67	3.52	222.4	1622		MS
isospathulenol	3.77	2.5	220.4	1600	1630	RI, MS
caryophylladienol II	1.25	1.06	220.4	1603	1633	RI, MS
τ -muurolol	0.17	1.04	222.4	1613	1640	RI, MS
isoaromadendrene epoxide			220.4	1662		MS
terpinen-4-ol			154.3	1156	1175	RI, MS
(-) -myrtenol			152.2	1174	1213	RI, MS
(-) -trans-pinocarveol			152.2	1117	1137	RI, MS
linalool			154.3	1081	1097	RI, MS
<i>l</i> -camphor			152.2	1122	1139	RI, MS
(1 <i>S</i>)-(--) -verbenone			150.2	1187	1204	RI, MS
4-methylene-5-hexenal			110.2	1085	897	RI, MS
2,5-dimethyl-3-methylene-1,5-heptadiene	0.31		136.2	1591	915	RI, MS
9-(1-methylethylidene)-1,5-cycloundecadiene		0.31	204.4	1736		MS
(-) -xanthorrhizol		0.63	218.3	1723		MS

1,1,7,7a-tetramethyl-1a,2,6,7,7a,7b-hexahydro-1 <i>H</i> -cyclopropa[a]-naphthalene	0.21	202.3	1761		MS
α -muurolene	0.2	204.4	1420	1440	RI, MS
<i>trans</i> - α -bergamotene	0.6	204.4	1469	1438	RI, MS
8,9-dehydro-neoisolongifolene	0.17	202.3	1795		MS
1-ethenyl-1-methyl-2,4-bis(1-methylethenyl)-cyclohexane	0.14	204.4	1367	1389	RI, MS
valeren-4,7(11)-diene	0.16	204.4	1756		MS
(-)isocaryophyllene	0.37	204.4	1689		MS
pentamethylcyclopentadiene	0.53	136.2	1726		MS
isophytol	0.11	296.3	1913		MS
manool oxide	0.15	290.5	1978		MS
m-anisalcohol	3.36	138.2	1180	1198	RI, MS
chlorpyrifos	0.42	350.6	1957		MS
4-methylene-6-(1-propenylidene)cyclooctene	0.11	160.3	1825		MS
monoterpenoids	38.71	31.29	1.05	4.4	
sesquiterpenoids	38.42	52.01	63.56	77.72	
diterpenoids	0.11		0.26		
aromatics	2.03	1.58	3.36		
aliphatics	3.11	1.03	6.68	2.83	
others			0.42		
total	82.38	85.91	75.32	84.95	

^a Percent calculated from MS data.; ^bIdentification method: MS, identified on the basis of computer matching of the mass spectra with those of the Wiley and Mass Finder libraries and comparison with literature data.

Table S2. Common components of EO_s.

Common components of GAAEOs and LAAEOs	Common components of GAVEOs and LAVEOs	Common components of GAVEOs and LAVEOs
(-)-4-terpinenol	(+)- α -calacorene	neointermedeol
(-)-carvone	(+)- β -eudesmol	caryophyllene
(-)- β -bourbonene	(\pm)-phytione	caryophyllene oxide
(\pm)-filifolone	(\pm)- α -curcumene	2-borneol
(\pm)-phytione	2-borneol	caryophylladienol I
(\pm)-piperitone	aromandendrene	humulene
1-octen-3-ol	caryophylladienol I	β -selinene
2-borneol	caryophylladienol II	4(15),5,10(14)- Germacratrien-1-Ol
(+)- δ -cadinene	caryophyllene	phytol
artemisia alcohol	caryophyllene oxide	(\pm)-phytione
bicyclogermacrene	<i>cis</i> - β -farnesene	
carveol	<i>epi</i> - β -caryophyllene	
caryophylladienol I	4(15),5,10(14)- Germacratrien-1-Ol	
caryophyllene	himbacol	
caryophyllene oxide	humulene	
chamazulene	isoledene	
<i>cis</i> -carveol	isospathulenol	
<i>cis</i> -piperitol	ledol	
<i>cis</i> -p-menth-2-en-1-ol	mustakone	
copaene	neointermedeol	
eremophylene	palustrol	
eucalyptol	phytol	
eugenol	α -gurjunene	
4(15),5,10(14)- Germacratrien-1-Ol	α -neocallitropsene	
germacrene D	β -bisabolene	
humulene	β -calacorene	
junenol	β -costol	
methyleneugenol	β -selinene	
neointermedeol	γ -cadinene	
perillaldehyde	τ -muurolol	
phytol		
selina-4,11-dien		
spainulenol		

trans-4-thujanol

yogomi alcoho

α -terpinene

α -terpineol

β -copaene

β -selinene

γ -piuronene

γ -terpinene

τ -cadinol

Table S3. Numbering correspondence table for Figure 4.

NO.	compounds	NO.	compounds	NO.	targets	NO.	pathway
C1	γ -pironene	C15	β -calacorene	T1	NTRK3	P1	PI3K-Akt signaling pathway
C2	α -cymene	C16	pentamethylcy clopentadiene	T2	NOS2	P2	Pathways of neurodegeneration - multiple diseases
C3	neointerme deol	C17	m-anisalcohol	T3	NFKB1	P3	Pathways in cancer
C4	methyleneugenol	C18	himbacol	T4	CNR2	P4	Neuroactive ligand-receptor interaction
C5	junenol	C19	caryophyllene oxide	T5	CHRM4	P5	MicroRNAs in cancer
C6	eugenol	C20	alloaromadendrene oxide	T6	AR	P6	Metabolic pathways
C7	eucalyptol	C21	2-borneol	T7	APOBEC 3A	P7	Chemical carcinogenesis-receptor activation
C8	<i>cis</i> -carveol	C22	1-ethenyl-1-methyl-2,4-bis(1-methylethenyl)-cyclohexane	T8	APEX1	P8	cAMP signaling pathway
C9	chamazulene	C23	α -himachalene	T9	ADRA2C	P9	Calcium signaling pathway
C10	1-octen-3-ol	C24	bornyl acetate	T10	ACHE	P10	Alzheimer disease
C11	β -elemene	C25	9-(1-methylethylidene)-1,5-cycloundecadiene	T11	HTR1E		
C12	trimethylenenorbornane	C26	(1S)-(-)-verbenone	T12	NR3C2		
C13	p-cymene			T13	KDM1A		
C14	β -selinene			T14	GPR55		

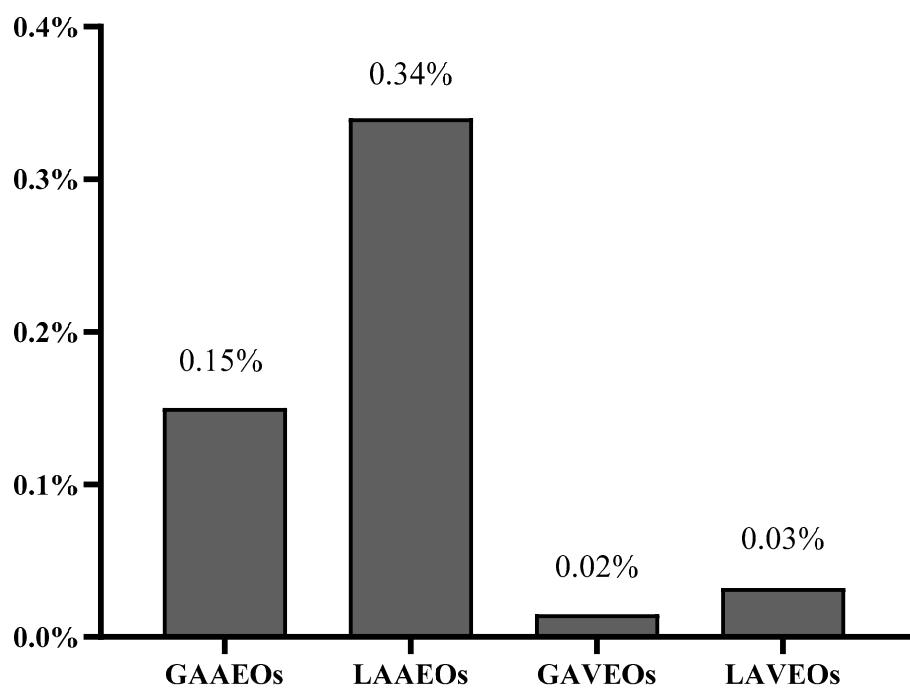
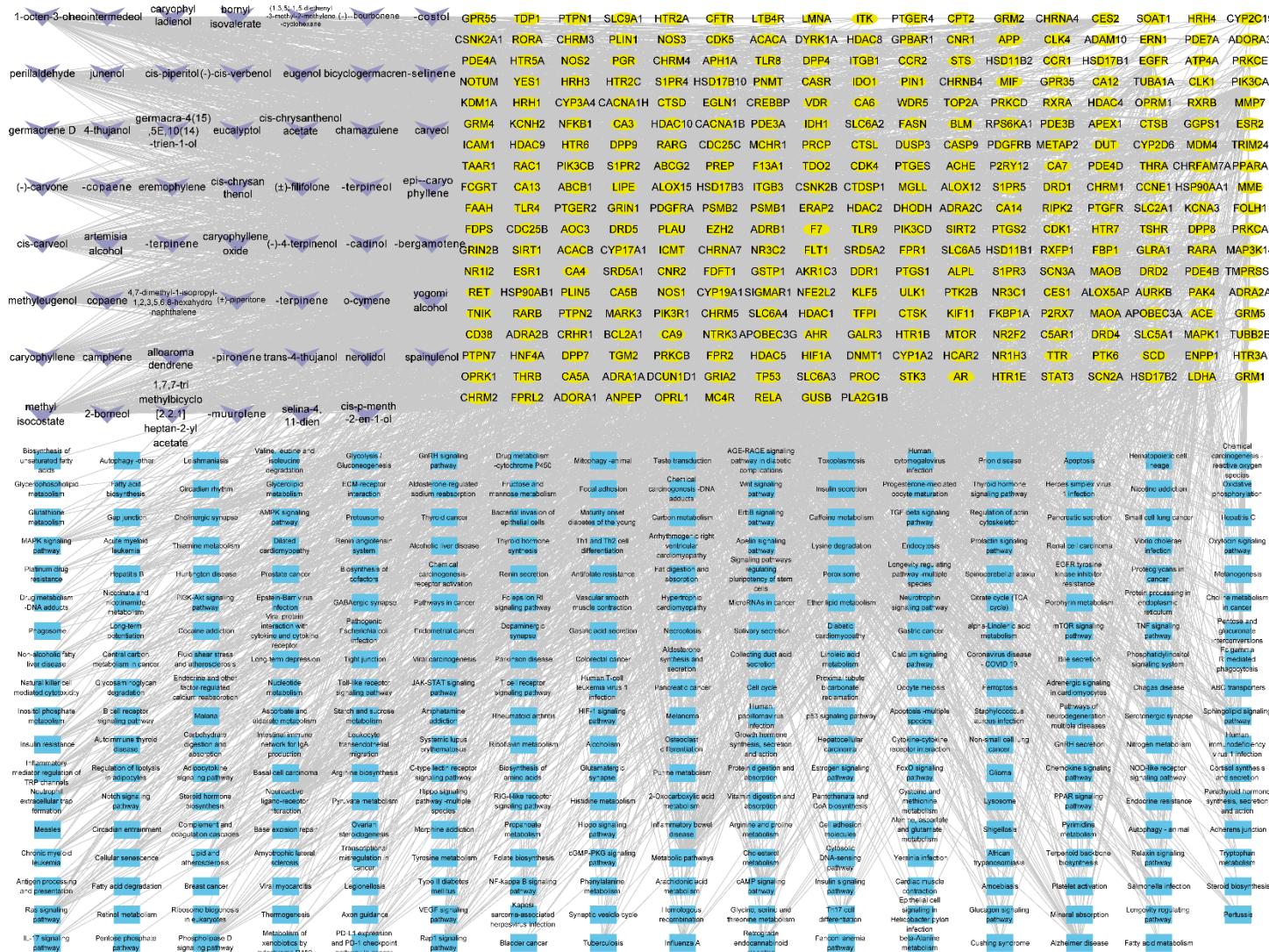
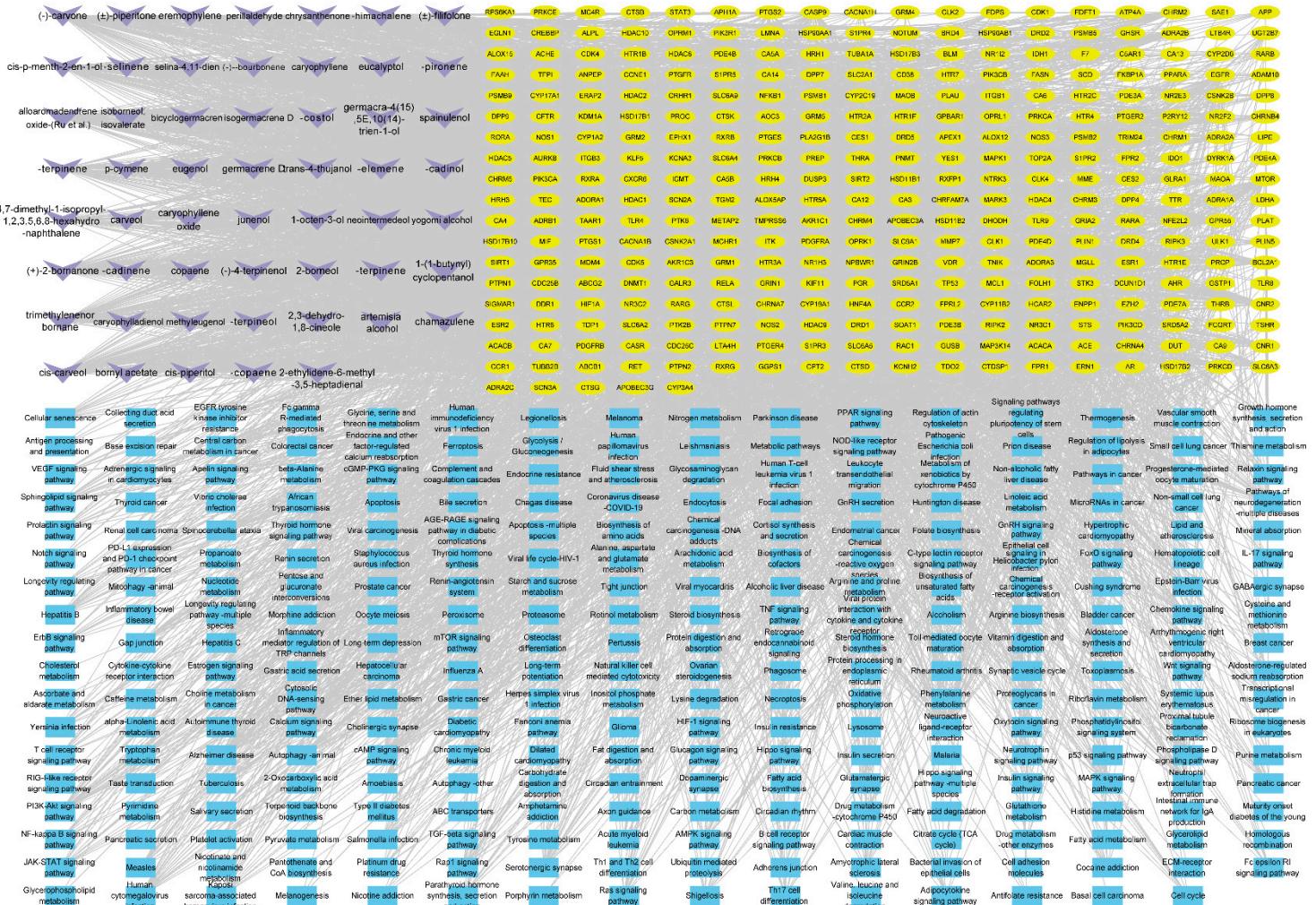


Figure S1. The Yields of EOs of *Artemisia*.

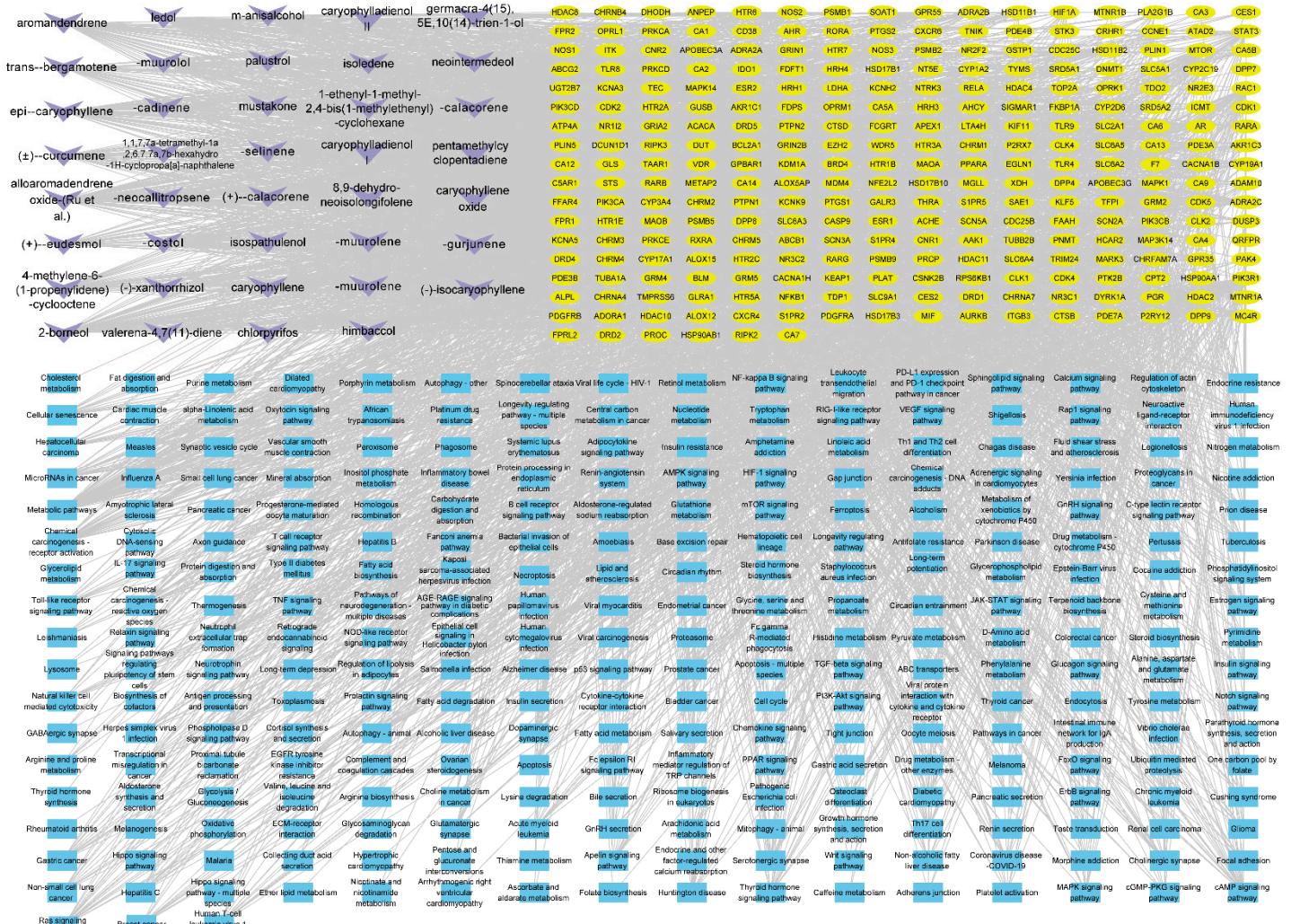
GAAEOs



LAAEOs



GAVEOs



LAVEOs

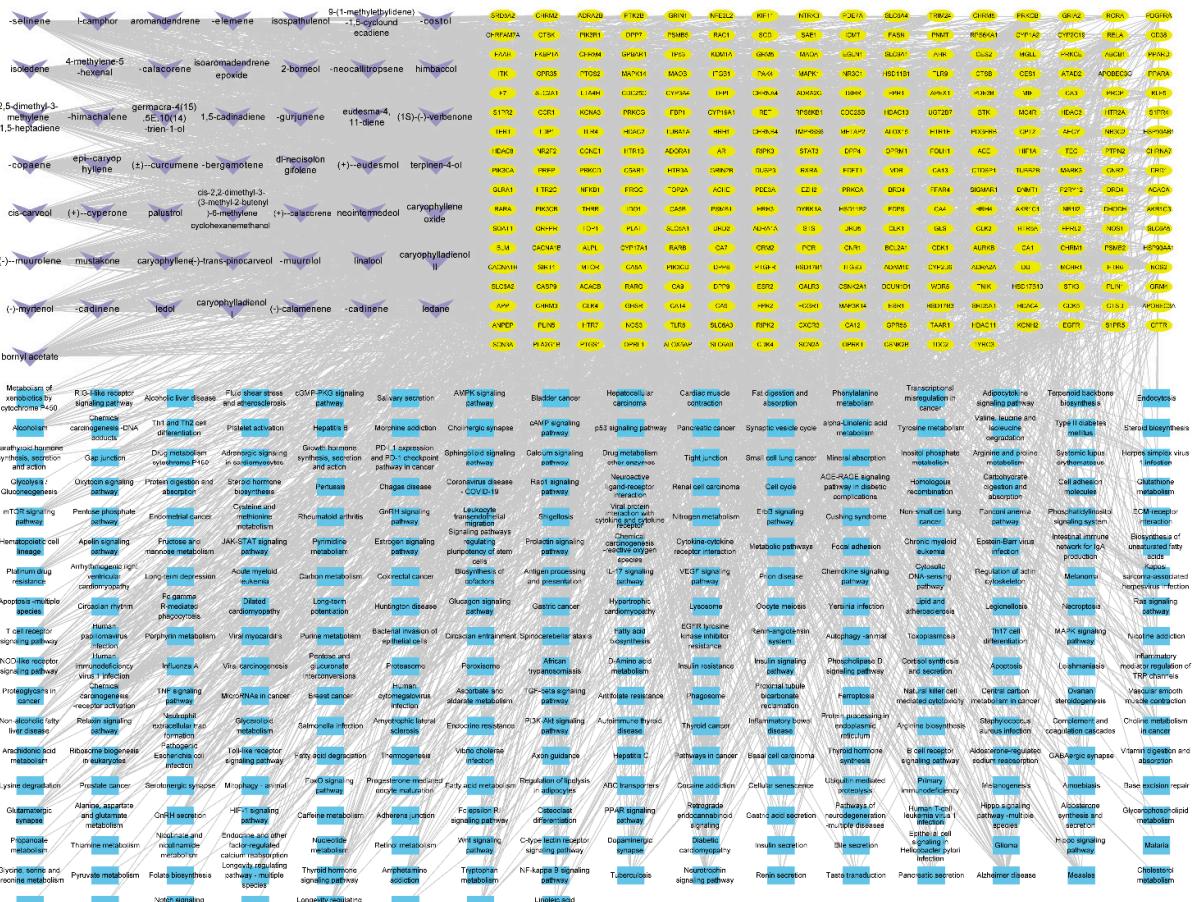


Figure S2. Ingredient-Target-pathway network of EOs. The ingredient represented by the purple V. The targets were represented by the yellow ellipse. The pathways were represented by the blue rectangle. The active ingredients were represented by the node, and the edge links the targets and the active ingredients.

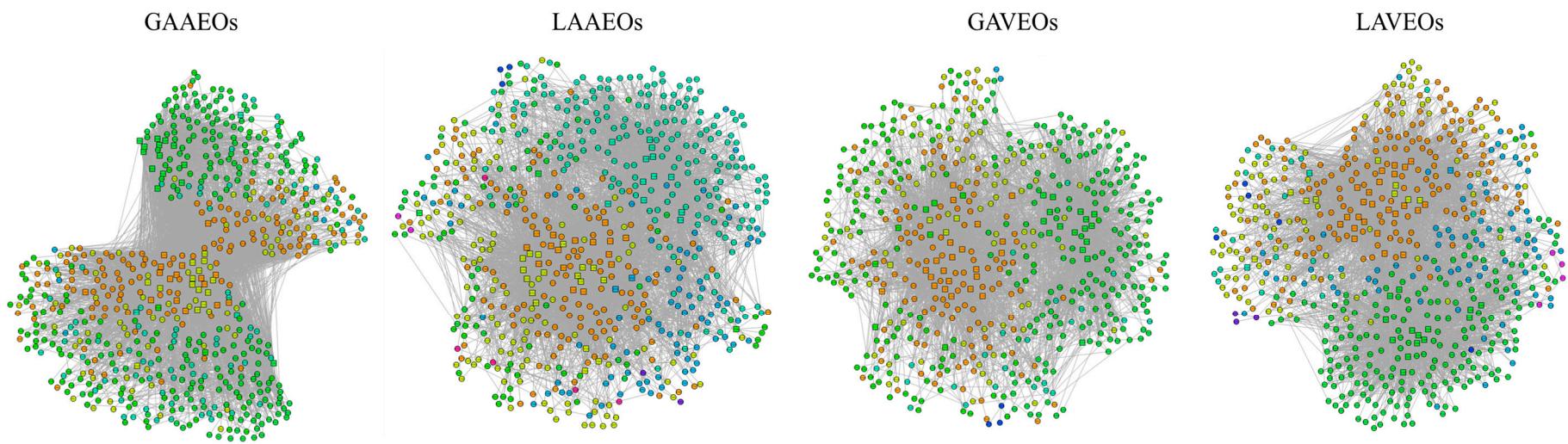
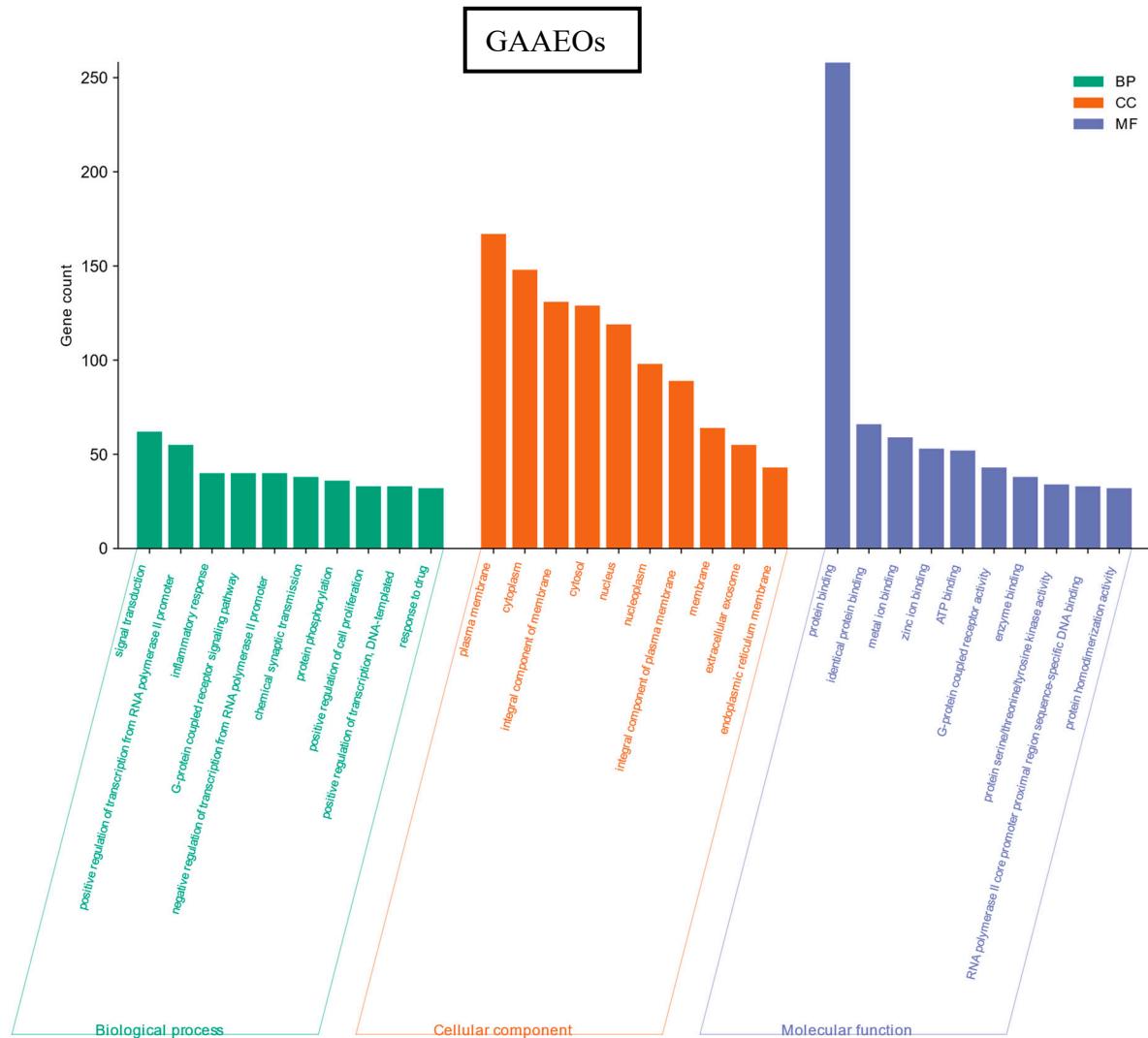
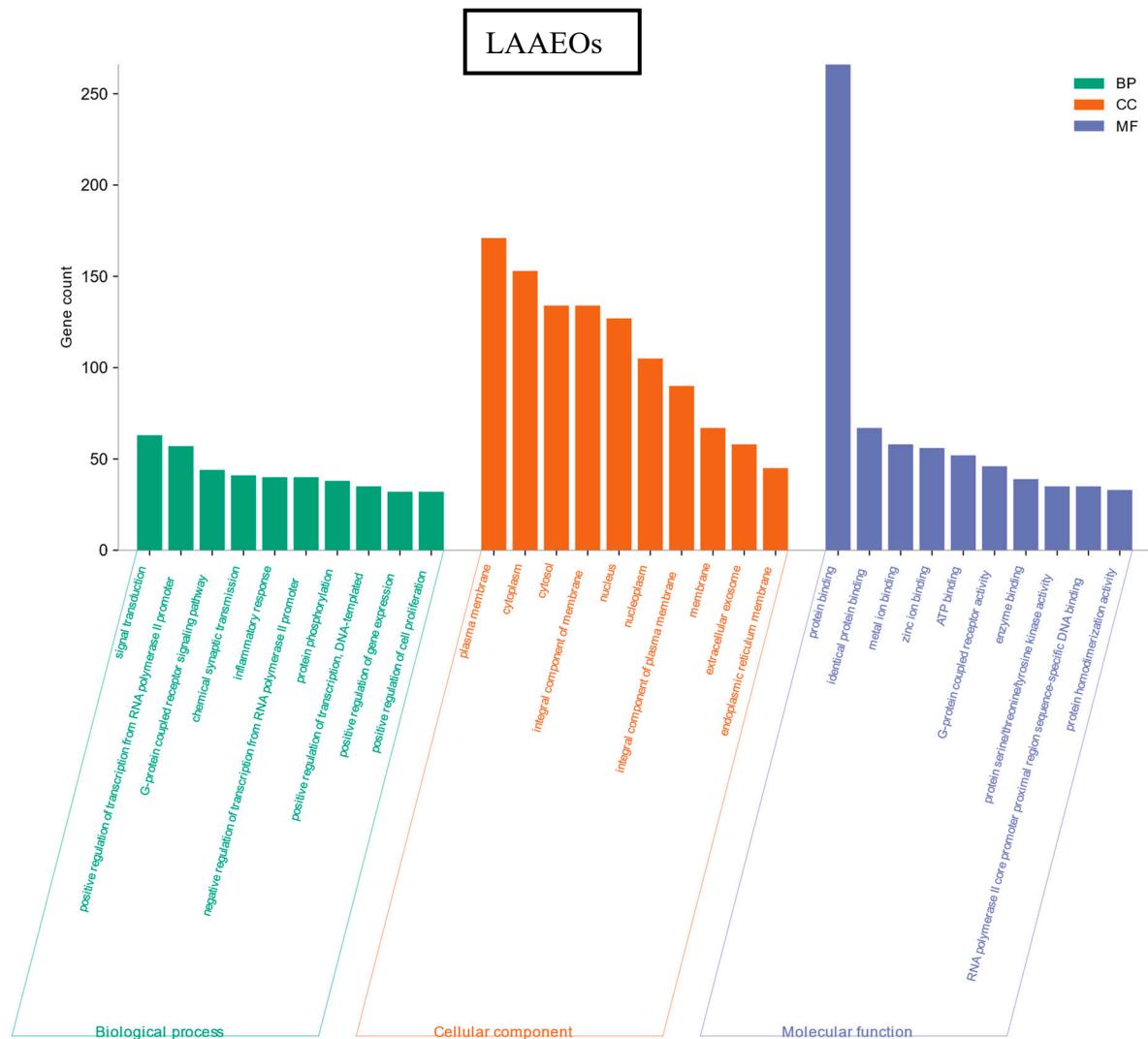
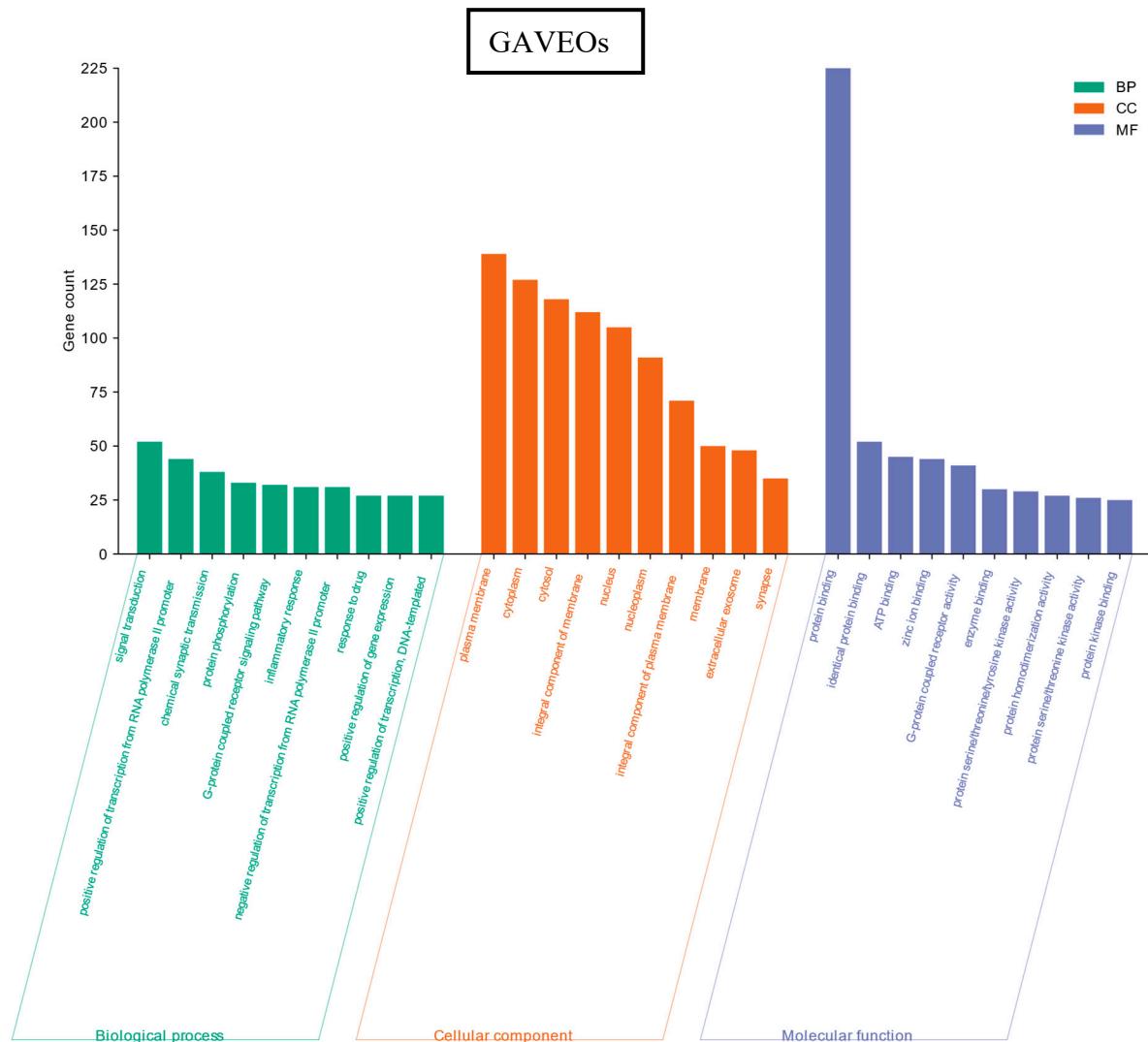


Figure S3. R cluster analysis. The related ingredients, targets and pathways information resulted in a data set which was converted to an igraph graph with “igraph” software package. Different communities were marked in different colors by a R_rainbow, and nodes with similar colors have higher associativity.







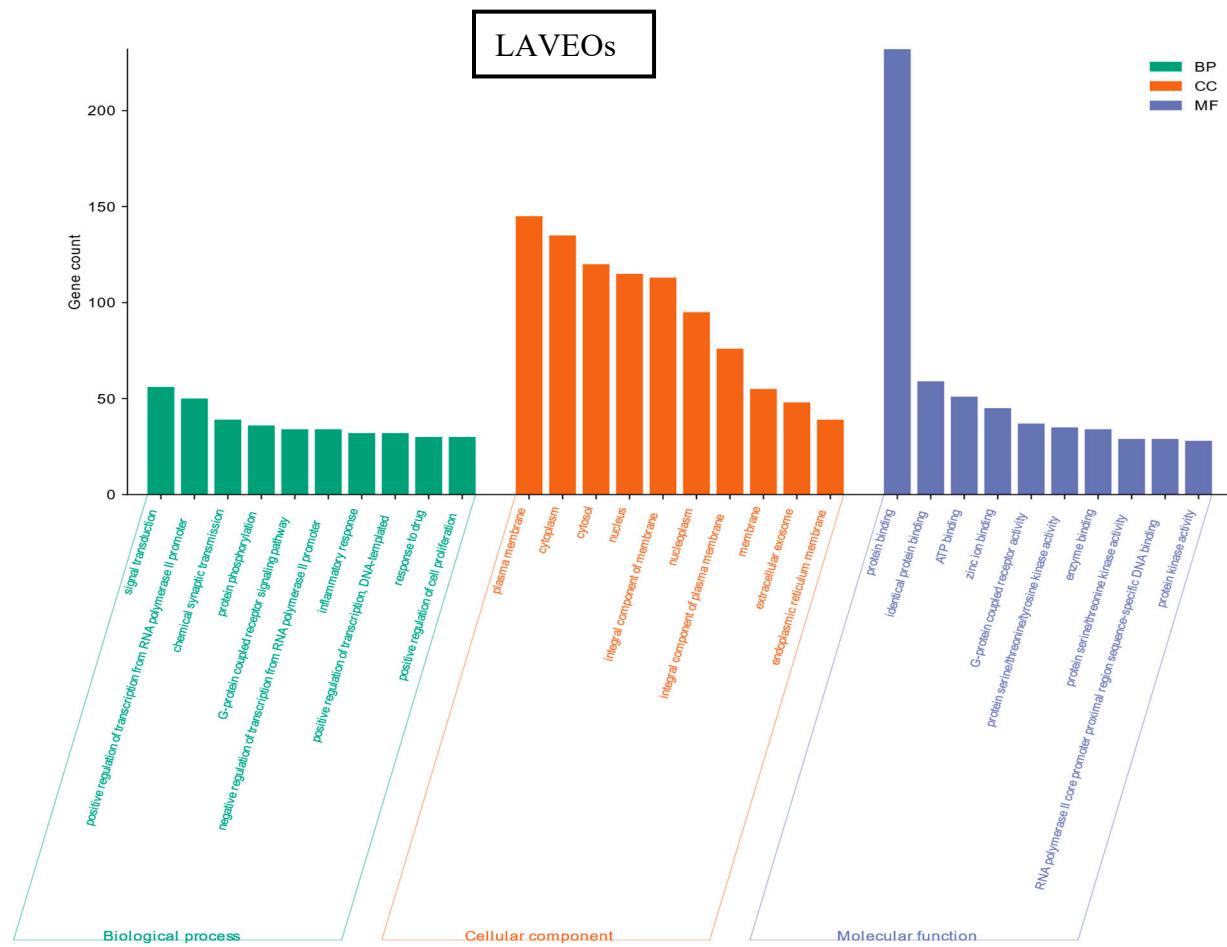


Figure S4. Gene Ontology (GO) analysis. Main Gene Ontology terms enriched by major hubs from cluster Profiler. The top 10 terms of BP, MM, and CC measured by adjusted value were selected to demonstrate.

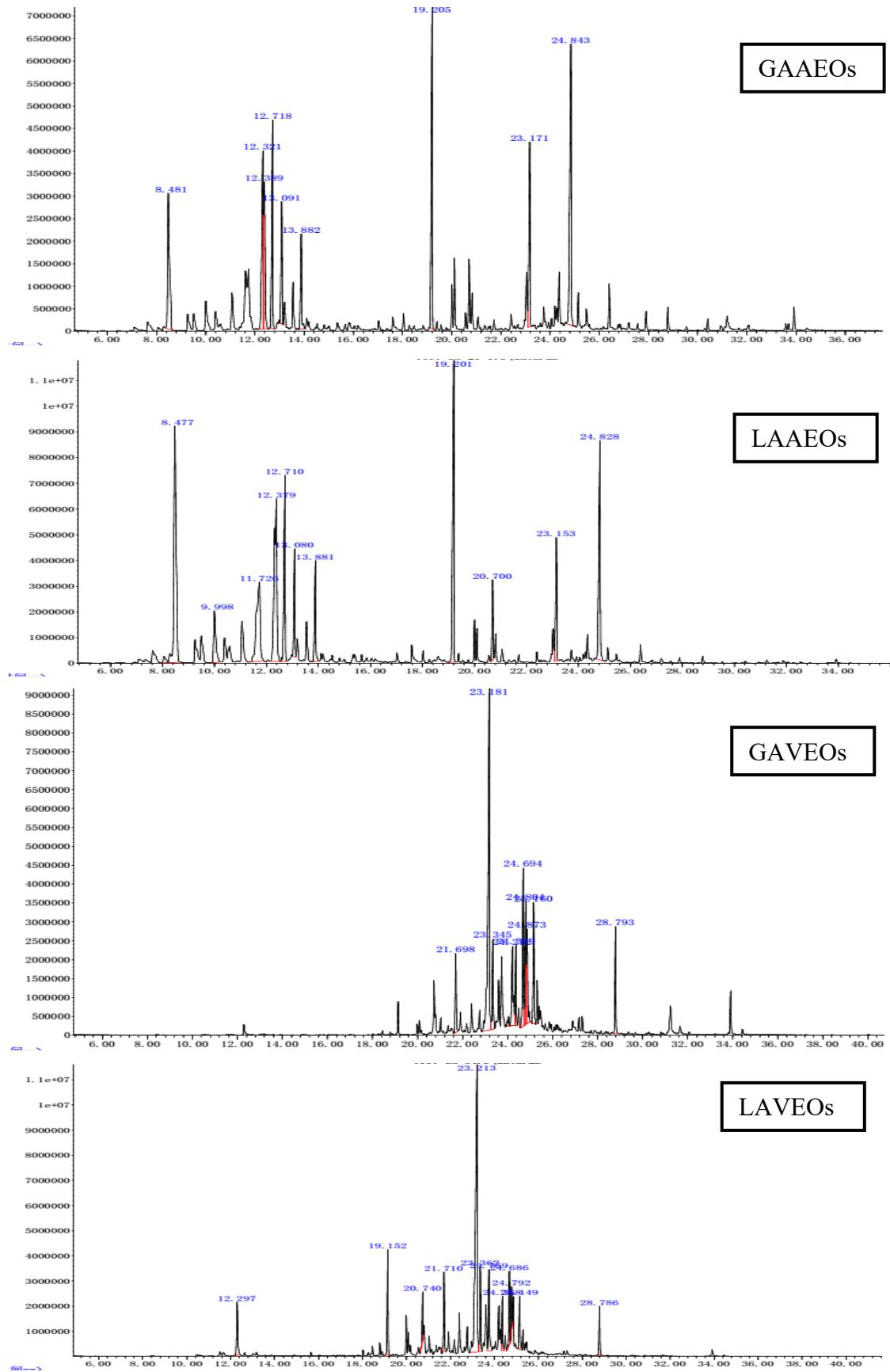


Figure S5 Total Ions Chromatograms (TIC) of EOs from *Artemisia*.