

**Table S1. Composition and dosage of herbal ingredients in Yi-qi-hua-yu-jie-du decoction (YJD)**

Scientific name*	Weight
<i>Astragalus mongholicus</i> Bunge	15g
<i>Codonopsis pilosula</i> (Franch.) Nannf.	15g
<i>Citrus × acida</i> Pers.	6g
<i>Pinellia tuberifera</i> Ten.	10g
<i>Poria cocos</i>	10g
<i>Aucklandia costus</i> Falc.	6g
<i>Wurfbainia uliginosa</i> (J.Koenig) Giseke	3g
<i>Atractylodes macrocephala</i> Koidz	10g
<i>Paeonia lactiflora</i> Pall.	10g
<i>Angelica sinensis</i> (Oliv.) Diels	10g
<i>Sparganium stoloniferum</i> (Buch.-Ham. ex Graebn.) Buch.-Ham. ex Juz	10g
<i>Curcuma aromatica</i> Salisb	10g
<i>Salvia chinensis</i> Benth.	30g
<i>Scleromitron angustifolium</i> (Cham. & Schltdl.) Benth.	30g
<i>Glycyrrhiza glabra</i> L.	5g

\* The name has been validated at [www.theplantlist.org](http://www.theplantlist.org).

**Table S2. The content of the marker compounds in YJD**

Name	CAS No	Content in YJD (ng·mg <sup>-1</sup> )
Astragaloside IV	84687-43-4	26.970
Calycosin-7-O-beta-D-glucoside	20633-67-4	144.007
Lobetyolin	136085-37-5	99.204
Atractylenolide III	73030-71-4	32.020
Paeoniflorin	23180-57-6	2305.240
Ferulic acid	1135-24-6	145.685
Succinic acid	110-15-6	682.296
Hesperidin	520-26-3	2239.183
Pachymic acid	29070-92-6	6.024
Dehydrocostus lactone	477-43-0	36.739
Rutin	153-18-4	234.124
Curcumin	458-37-7	50.275
Rosmarinic acid	20283-92-5	95.101
Salvianolic acid B	121521-90-2	496.713
p-Coumaric acid	501-98-4	334.314
Liquiritin	551-15-5	486.248

### **Method S1. Sample preparation and analysis method**

Standard reagents were purchased from Chengdu Must Biotechnology Co (China). Mixed standard working fluids (1–1000 ng/mL) were prepared. Freeze-dried YJD powder (20 mg) was weighed and dissolved in 10 mL double-distilled water. Then, 500  $\mu$ L of the above solution was diluted to 10 mL and the supernatant was collected for LC/MS analysis.

Chromatographic conditions: The Altis-002 (Thermo, USA) and ACE Excel 3 C18-AR column (ACE, UK) were applied to the liquid chromatography analysis. Mobile phase A was 2 mM ammonium formate, and mobile phase B was acetonitrile. The gradient elution program was as follows: 0–10 min 78% A + 22% B, 10–26 min 15% A + 85% B, 26–30 min 78% A + 22% B.

Mass spectrometry conditions: The flow rate of the sheath gas was 50.0 Arb, and that of the auxiliary gas was 10.0 Arb. The ion spray voltages were 3500 V and 2500 V for the positive and negative modes, respectively.

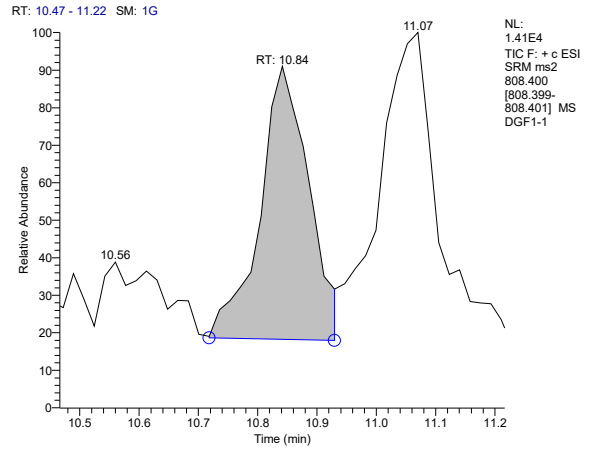
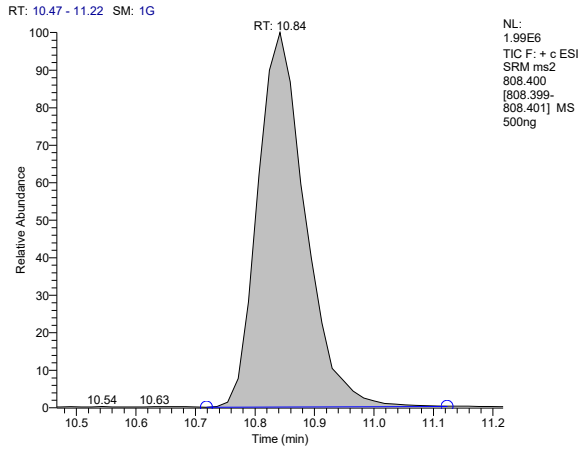
The concentration of each marker compound in YJD was determined based on the standard curve for each reagent and the signal intensity of the corresponding substance in YJD.

**Fig. S1 The secondary mass spectra of the marker compounds in the standard reagent and YJD.**

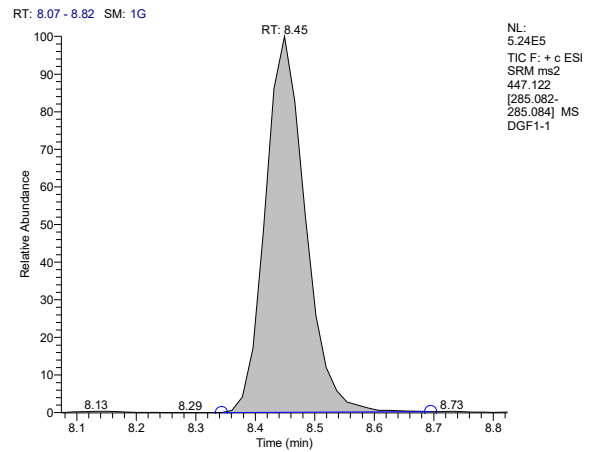
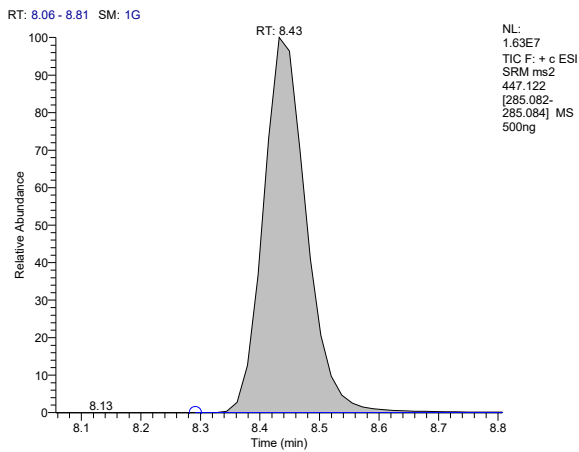
**Standard reagent**

**YJD**

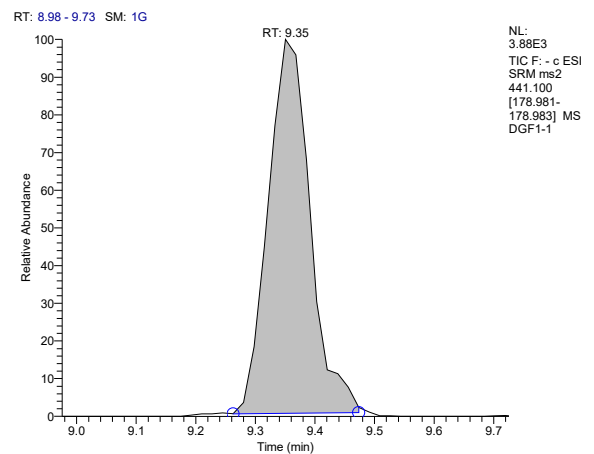
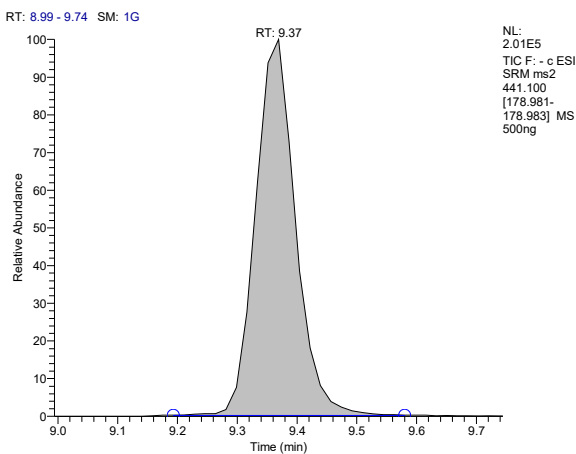
**1. Astragaloside IV**



**2. Calycosin-7-O-beta-D-glucoside**

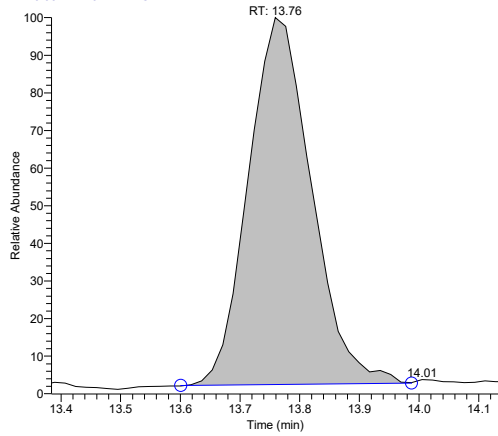


**3. Lobetyolin**



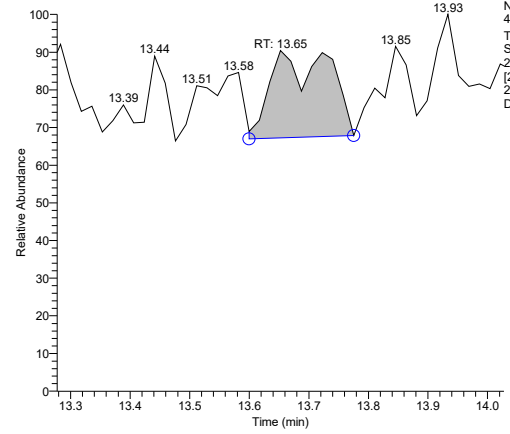
## 4. Atractylenolide III

RT: 13.38 - 14.13 SM: 1G



NL: 1.09E5  
TIC F: + c ESI  
SRM ms2 249.100  
[231.011-231.013] MS  
500ng

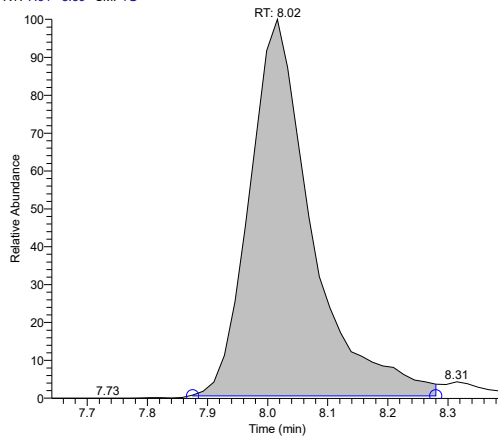
RT: 13.28 - 14.03 SM: 1G



NL: 4.66E3  
TIC F: + c ESI  
SRM ms2 249.100  
[231.011-231.013] MS  
DGF1-1

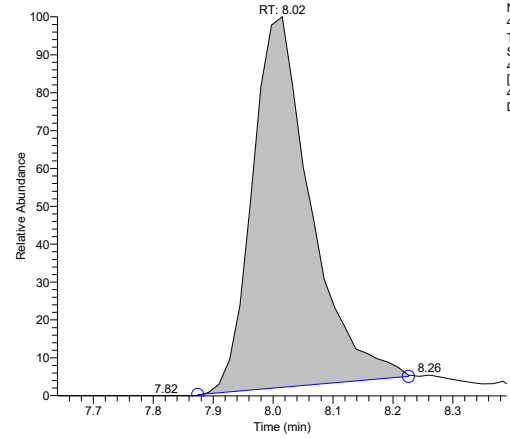
## 5. Paeoniflorin

RT: 7.64 - 8.39 SM: 1G



NL: 8.37E4  
TIC F: - c ESI  
SRM ms2 479.050  
[449.124-449.126] MS  
500ng

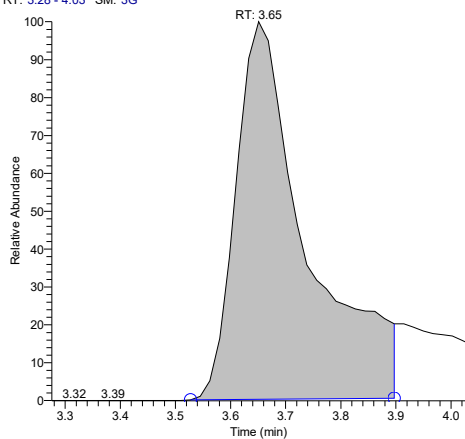
RT: 7.64 - 8.39 SM: 1G



NL: 4.38E4  
TIC F: - c ESI  
SRM ms2 479.050  
[449.124-449.126] MS  
DGF1-1

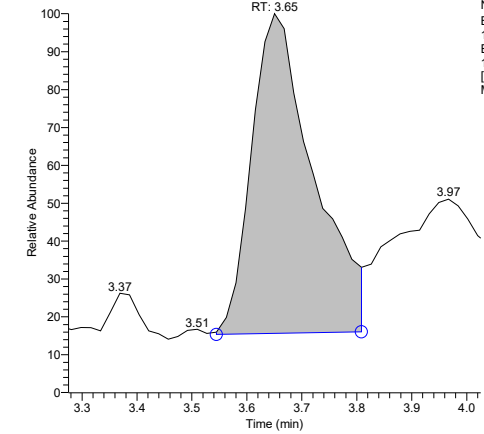
## 6. Ferulic acid

RT: 3.28 - 4.03 SM: 3G



NL: 1.10E6  
Base Peak m/z= 133.48-134.48 F: - c  
ESI SRM ms2 192.950  
[133.987-133.989] MS  
500ng

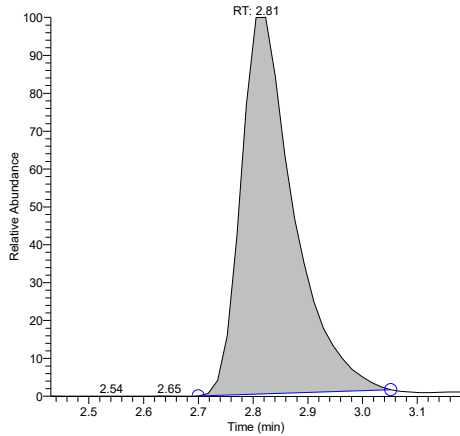
RT: 3.28 - 4.03 SM: 3G



NL: 4.00E4  
Base Peak m/z= 133.48-134.48 F: - c  
ESI SRM ms2 192.950  
[133.987-133.989] MS  
DGF1-1

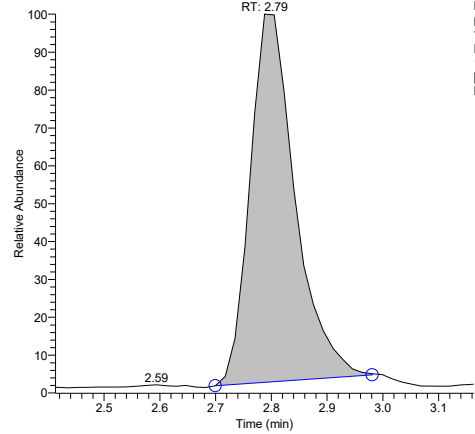
## 7. Succinic acid

RT: 2.43 - 3.18 SM: 3G



NL: 9.00E5  
Base Peak m/z=  
72.50-73.50 F: - c ESI  
SRM ms2 117.150  
[72.999-73.001] MS  
ICIS 500ng

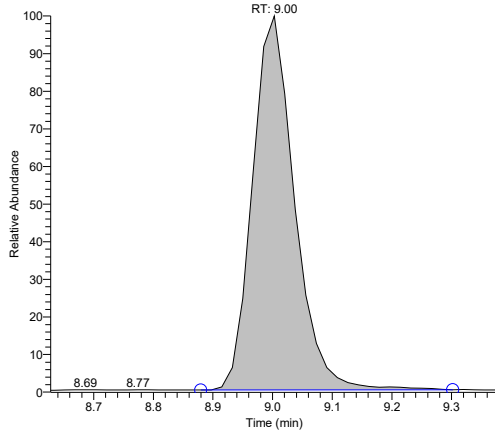
RT: 2.41 - 3.16 SM: 3G



NL: 1.62E5  
Base Peak m/z=  
72.50-73.50 F: - c  
ESI SRM ms2  
117.150  
[72.999-73.001] MS  
DGF1-1

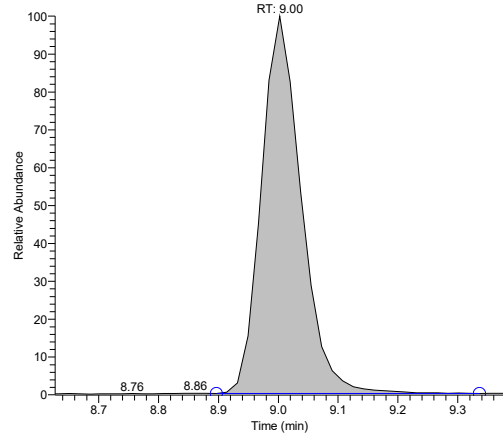
## 8. Hesperidin

RT: 8.63 - 9.38 SM: 1G



NL:  
2.72E6  
TIC F: - c ESI  
SRM ms2  
609.100  
[301.053-  
301.055] MS  
500ng

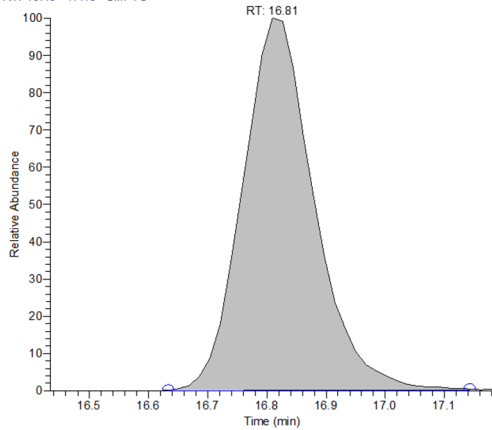
RT: 8.63 - 9.38 SM: 1G



NL:  
1.42E6  
TIC F: - c ESI  
SRM ms2  
609.100  
[301.053-  
301.055] MS  
DGF1-1

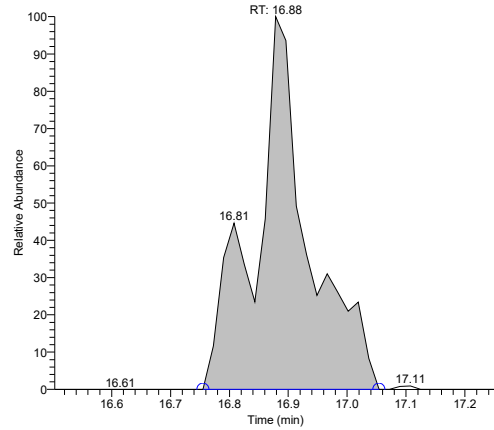
## 9. Pachymic acid

RT: 16.43 - 17.18 SM: 1G



NL:  
5.44E5  
TIC F: - c ESI  
SRM ms2  
527.350  
[465.320-  
465.322] MS  
500ng

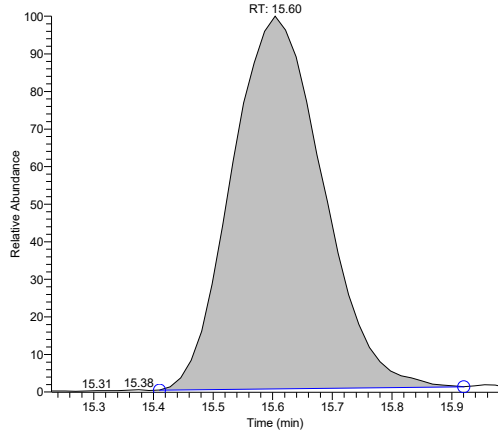
RT: 16.50 - 17.25 SM: 1G



NL:  
5.44E2  
TIC F: - c ESI  
SRM ms2  
527.350  
[465.320-  
465.322] MS  
DGF1-1

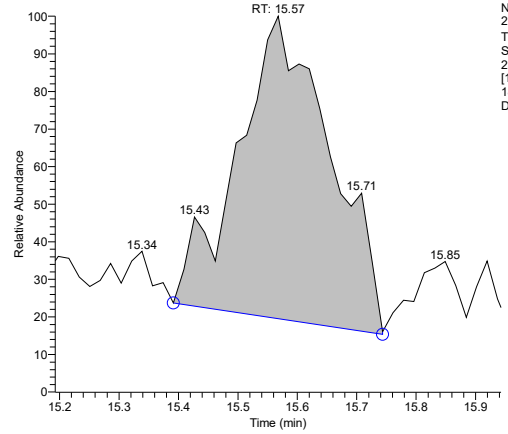
## 10. Dehydrocostus lactone

RT: 15.23 - 15.98 SM: 1G



NL: 1.82E5  
TIC F: + c ESI  
SRM ms2  
231.100  
[185.154-  
185.156] MS  
500ng

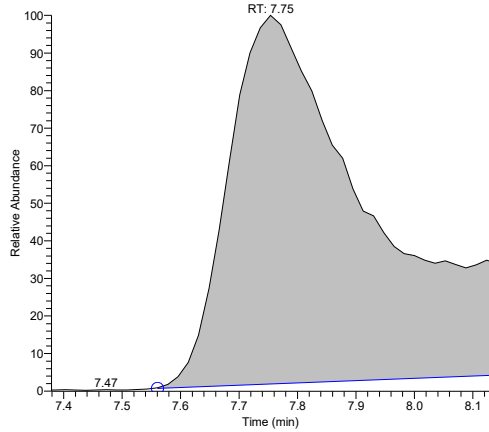
RT: 15.19 - 15.94 SM: 1G



NL: 2.93E3  
TIC F: + c ESI  
SRM ms2  
231.100  
[185.154-  
185.156] MS  
DGF1-1

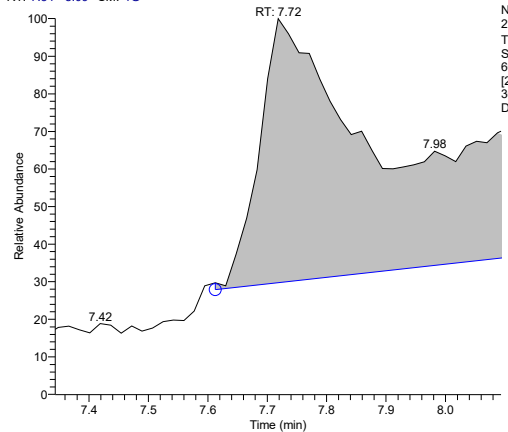
## 11. Rutin

RT: 7.38 - 8.13 SM: 1G



NL: 5.12E5  
TIC F: - c ESI  
SRM ms2  
609.100  
[299.999-  
300.001] MS  
ICIS 500ng

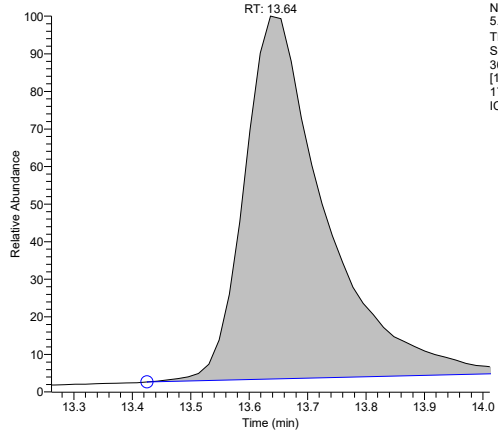
RT: 7.34 - 8.09 SM: 1G



NL: 2.70E4  
TIC F: - c ESI  
SRM ms2  
609.100  
[299.999-  
300.001] MS  
DGF1-1

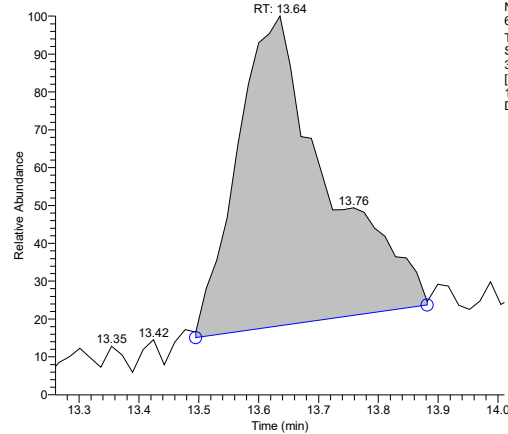
## 12. Curcumin

RT: 13.26 - 14.01 SM: 1G



NL: 5.38E6  
TIC F: + c ESI  
SRM ms2  
369.050  
[176.999-  
177.001] MS  
ICIS 500ng

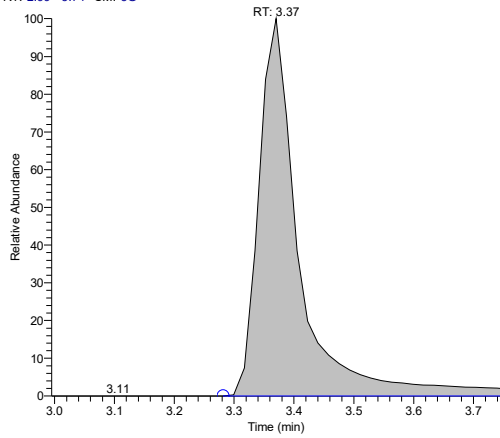
RT: 13.26 - 14.01 SM: 1G



NL: 6.26E3  
TIC F: + c ESI  
SRM ms2  
369.050  
[176.999-  
177.001] MS  
DGF1-1

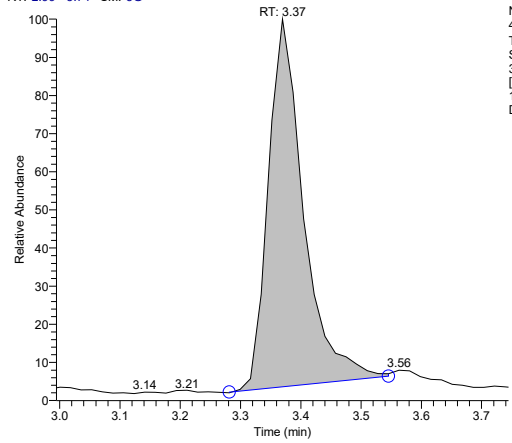
### 13. Rosmarinic acid

RT: 2.99 - 3.74 SM: 3G



NL: 1.70E6  
TIC F: - c ESI  
SRM ms2  
359.000  
[161.070-161.072] MS  
500ng

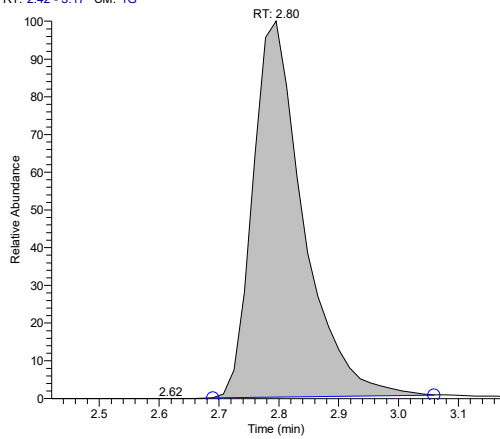
RT: 2.99 - 3.74 SM: 3G



NL: 4.43E4  
TIC F: - c ESI  
SRM ms2  
359.000  
[161.070-161.072] MS  
DGF1-1

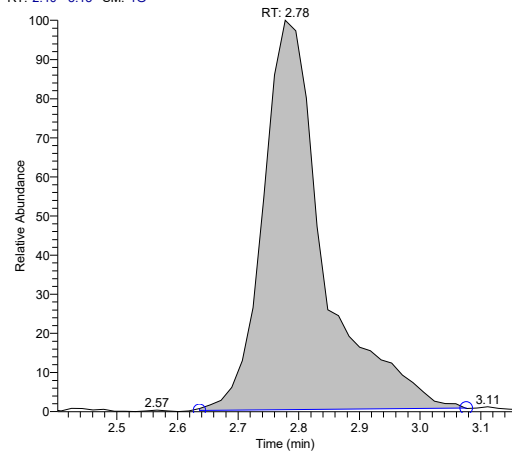
### 14. Salvianolic acid B

RT: 2.42 - 3.17 SM: 1G



NL: 2.47E5  
TIC F: - c ESI  
SRM ms2  
717.113  
[519.064-519.066] MS  
500ng

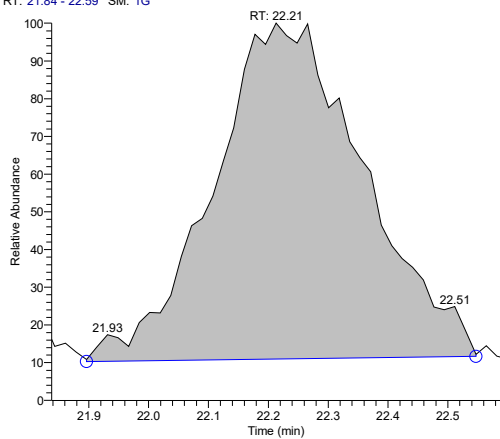
RT: 2.40 - 3.15 SM: 1G



NL: 2.05E4  
TIC F: - c ESI  
SRM ms2  
717.113  
[519.064-519.066] MS  
ICIS DGF1-1

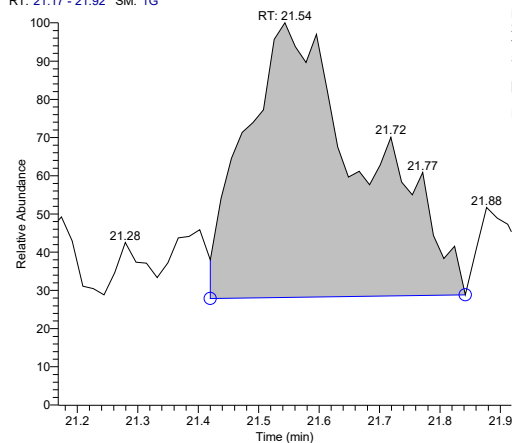
### 15. p-Coumaric acid

RT: 21.84 - 22.59 SM: 1G



NL: 7.18E3  
TIC F: + c ESI  
SRM ms2  
163.000  
[107.053-107.055] MS  
500ng

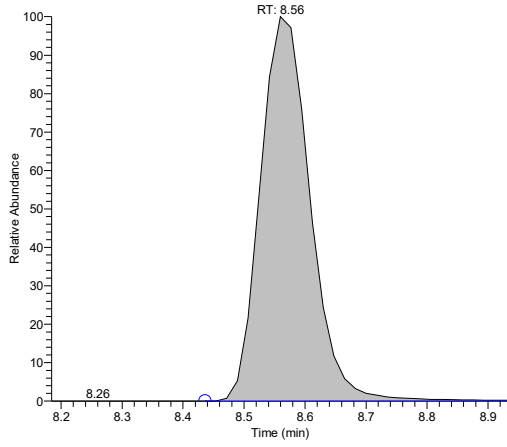
RT: 21.17 - 21.92 SM: 1G



NL: 2.12E3  
TIC F: + c ESI  
SRM ms2  
163.000  
[107.053-107.055] MS  
DGF1-1

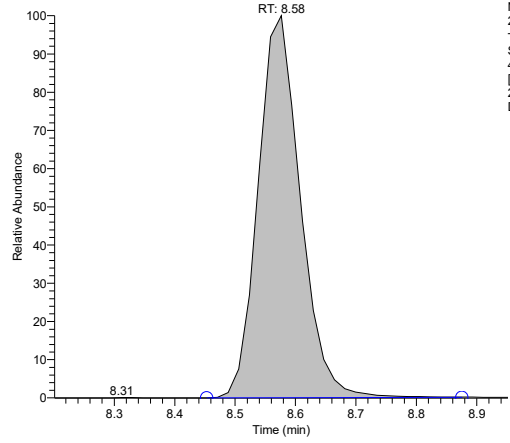
# 16. Liquiritin

RT: 8.18 - 8.93 SM: 1G



NL:  
1.54E7  
TIC F: - c ESI  
SRM ms2  
417.050  
[254.999-  
255.001] MS  
500ng

RT: 8.20 - 8.95 SM: 1G

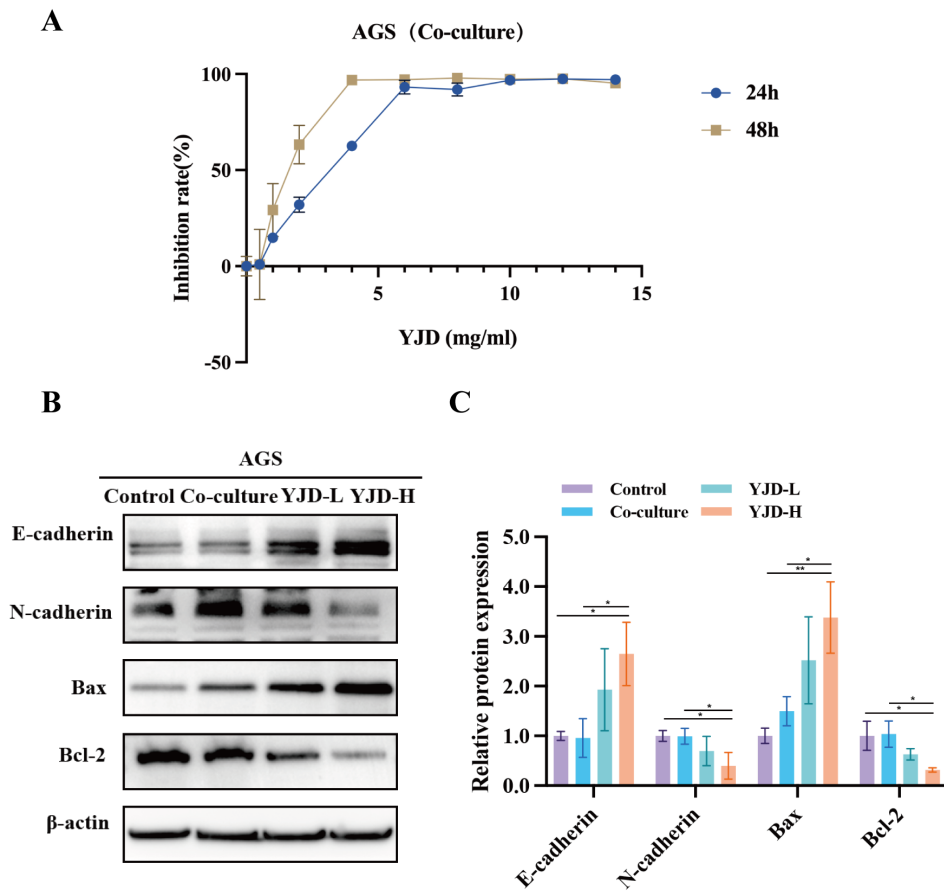


NL:  
2.17E6  
TIC F: - c ESI  
SRM ms2  
417.050  
[254.999-  
255.001] MS  
DGF1-1

**Table S3. The active ingredients of Yi-qi-hua-yu-jie-du decoction (YJD)**

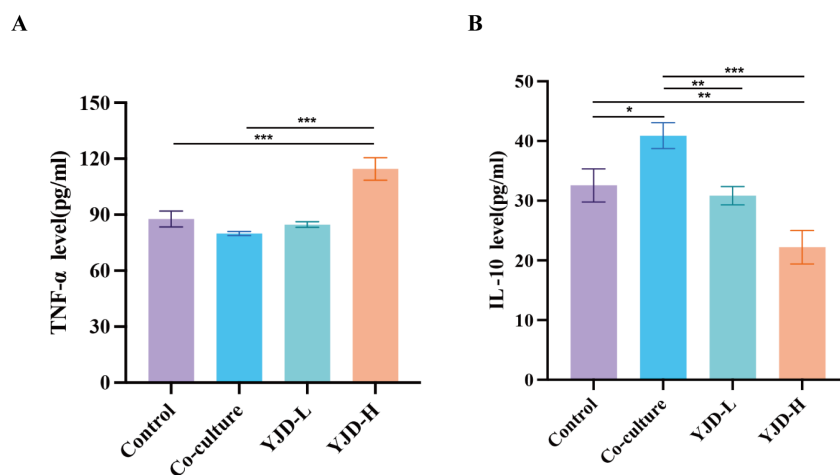
NO	Proposed compound	CAS	PubChem CID	Canonical SMILES
YJD1	Quinic acid	77-95-2	6508	<chem>C1C(C(C(CC1(C(=O)O)O)O)O)O</chem>
YJD2	Tyrosine	60-18-4	6057	<chem>C1=CC(=CC=C1CC(C(=O)O)N)O</chem>
YJD3	Danshensu	76822-21-4	11600642	<chem>C1=CC(=C(C=C1CC(C(=O)O)O)O)O</chem>
YJD4	trans-caffeic acid	501-16-6	689043	<chem>C1=CC(=C(C=C1C=CC(=O)O)O)O</chem>
YJD5	Tryptophan	73-22-3	6305	<chem>C1=CC=C2C(=C1)C(=CN2)CC(C(=O)O)N</chem>
YJD6	Dihydrocaffeic acid	1078-61-1	348154	<chem>C1=CC(=C(C=C1CCC(=O)O)O)O</chem>
YJD7	Codonopsine	26989-20-8	442631	<chem>CC1C(C(C(N1C)C2=CC(=C(C=C2)OC)OC)O)O</chem>
YJD8	cis-caffeic acid	4361-87-9	1549111	<chem>C1=CC(=C(C=C1C=CC(=O)O)O)O</chem>
YJD9	Ferulic acid	537-98-4	445858	<chem>COC1=C(C=CC(=C1)C=CC(=O)O)O</chem>
YJD10	Trans-Melilotoside	618-67-7	5280759	<chem>C1=CC=C(C(=C1)C=CC(=O)O)OC2C(C(C(O2)CO)O)O</chem>
YJD11	Isofraxidin	486-21-5	5318565	<chem>COC1=C(C(=C2C(=C1)C=CC(=O)O2)OC)O</chem>
YJD12	7-Hydroxycoumarin	93-35-6	5281426	<chem>C1=CC(=CC2=C1C=CC(=O)O2)O</chem>
YJD13	loliolide	5989/2/6	100332	<chem>CC1(CC(CC2(C1=CC(=O)O2)C)O)C</chem>
YJD14	Syringaldehyde	134-96-3	8655	<chem>COC1=CC(=CC(=C1O)OC)C=O</chem>
YJD15	paeoniflorin	23180-57-6	442534	<chem>CC12CC3(C4CC1(C4(C(O2)O3)COC(=O)C5=CC=CC=C5)OC6C(C(C(O6)CO)O)O)O</chem>
YJD16	Scopoletin	92-61-5	5280460	<chem>COC1=C(C=C2C(=C1)C=CC(=O)O2)O</chem>
YJD17	Germacrone-4,5-epoxide	92691-35-5	11637239	<chem>CC1=CCCC2(C(O2)CC(=C(C)C)C(=O)C1)C</chem>
YJD18	Salvianolic acid J	-	24177556	<chem>C1=CC(=C(C=C1CC(C(=O)O)OC(=O)C=CC2=CC3=C(C=C2)OC(C(O3)C(=O)O)C4=CC(=C(C=C4)O)O)O)O</chem>
YJD19	Zedoalactone A	-	15226639	<chem>CC1=C2CC3C(CCC3(C)O)C(CC2OC1=O)(C)O</chem>
YJD20	Liquiritigenin	578-86-9	114829	<chem>C1C(OC2=C(C1=O)C=CC(=C2)O)C3=CC=C(C=C3)O</chem>
YJD21	camphoric acid	560-05-4	21491	<chem>CC1(C(CCC1(C)C(=O)O)C(=O)O)C</chem>
YJD22	Luteolin	491-70-3	5280445	<chem>C1=CC(=C(C=C1C2=CC(=O)C3=C(C=C(C=C3O2)O)O)O)O</chem>
YJD23	Perlolyrine	29700-20-7	160179	<chem>C1=CC=C2C(=C1)C3=C(N2)C(=NC=C3)C4=CC=C(O4)CO</chem>
YJD24	Acacetin	480-44-4	5280442	<chem>COC1=CC=C(C=C1)C2=CC(=O)C3=C(C=C(C=C3O2)O)O</chem>

YJD25	kaempferol	520-18-3	5280863	<chem>C1=CC(=CC=C1C2=C(C(=O)C3=C(C=C(C=C3O2)O)O)O</chem>
YJD26	Liquiritin	551-15-5	503737	<chem>C1C(OC2=C(C1=O)C=CC(=C2)O)C3=CC=C(C=C3)OC4C(C(C(C(O4)CO)O)O)O</chem>
YJD27	Hesperetin	520-33-2	72281	<chem>COC1=C(C=C(C=C1)C2CC(=O)C3=C(C=C(C=C3O2)O)O)O</chem>
YJD28	Methylnissolin-3-O-glucoside	94367-42-7	101679160	<chem>COC1=C(C2=C(C=C1)C3COC4=C(C3O2)C=CC(=C4)OC5C(C(C(C(O5)CO)O)O)O)OC</chem>
YJD29	pratensein	2284-31-3	5281803	<chem>COC1=C(C=C(C=C1)C2=COC3=CC(=CC(=C3C2=O)O)O)O</chem>
YJD30	Isoliquiritigenin	961-29-5	638278	<chem>C1=CC(=CC=C1C=CC(=O)C2=C(C=C(C=C2)O)O)O</chem>
YJD31	Calycosin	20575-57-9	5280448	<chem>COC1=C(C=C(C=C1)C2=COC3=C(C2=O)C=CC(=C3)O)O</chem>
YJD32	Poncirin	14941-08-3	442456	<chem>CC1C(C(C(C(O1)OC2C(C(C(OC2OC3=CC(=C4C(=O)CC(OC4=C3)C5=CC=C(C=C5)OC)O)CO)O)O)O)O)O</chem>
YJD33	Biochanin A	491-80-5	5280373	<chem>COC1=CC=C(C=C1)C2=COC3=CC(=CC(=C3C2=O)O)O</chem>
YJD34	Sinensetin	2306-27-6	145659	<chem>COC1=C(C=C(C=C1)C2=CC(=O)C3=C(C(=C(C=C3O2)OC)OC)OC)OC</chem>
YJD35	Alizarin	72-48-0	6293	<chem>C1=CC=C2C(=C1)C(=O)C3=C(C2=O)C(=C(C=C3)O)O</chem>
YJD36	Curcumenone	100347-96-4	153845	<chem>CC(=C1CC2C(C2(CC1=O)C)CCC(=O)C)C</chem>
YJD37	asiatic acid	464-92-6	119034	<chem>CC1CCC2(CCC3(C(=CCC4C3(CCC5C4(CC(C(C5(C)C(O)O)O)C)C)C2C1C)C)C(=O)O</chem>
YJD38	formononetin	485-72-3	5280378	<chem>COC1=CC=C(C=C1)C2=COC3=C(C2=O)C=CC(=C3)O</chem>
YJD39	Nobiletin	478-01-3	72344	<chem>COC1=C(C=C(C=C1)C2=CC(=O)C3=C(O2)C(=C(C(=C3OC)OC)OC)OC)OC</chem>
YJD40	Astragaloside III	84687-42-3	441905	<chem>CC1(C(CCC23C1C(CC4C2(C3)CCC5(C4(CC(C5C6(CC(C(O6)C(C)C)O)C)O)C)O)OC7C(C(C(CO7)O)O)OC8C(C(C(C(O8)CO)O)O)O)C</chem>
YJD41	Atractylenolide II	73069-14-4	14448070	<chem>CC1=C2CC3C(=C)CCCC3(CC2OC1=O)C</chem>



**Fig. S2** YJD suppresses AGS cell proliferation and induces apoptosis via TAM modulation in vitro.

(A) Viability of AGS cells exposed to conditioned medium from YJD-treated macrophages. (B) Western blotting analysis of E-cadherin, N-cadherin, Bax, and Bcl-2 in AGS cells. Data are presented as means  $\pm$  SD ( $n = 3$  independent experiments). \* $P < 0.05$ , \*\* $P < 0.01$ , \*\*\* $P < 0.001$ .



**Fig. S3** Effect of YJD on TNF- $\alpha$  and IL-10 secretion by TAMs in MKN-45 co-culture systems. (A) TNF- $\alpha$  levels in the supernatant of TAMs. (B) IL-10 levels in the supernatant of TAMs. Data are expressed as mean  $\pm$  SD ( $n = 3$  independent experiments). \* $P < 0.05$ , \*\* $P < 0.01$ , \*\*\* $P < 0.001$ .