Supplementary Table 2: The preparation and chemical analysis of the oral treatments of herbs and their active constituents

Species/compound, source, concentration	Quality control reported? (Y/N)	Chemical analysis reported? (Y/N)	Purity (%) [For compound only]	References
Whole plant of <i>Angelica</i> gigas Nakai, [Ginbu GAP Farming Corp], 330 g No voucher specimen or batch number was mentioned	Y- Air-dried and extracted with 95% ethanol at 80°C for 4 h using the machine to obtain the extract, which was then concentrated. The extract was collected, concentrated again using a rotary vacuum evaporator.	N	N/A	(102)
Whole plant of <i>Artemisia</i> caruifolia BuchHam. ex Roxb., [Yeongcheon Oriental Herbal Market], 30 g No voucher specimen or batch number was mentioned	Y- Extracted with 70% ethanol in a 40°C shake-incubator for 24 h and filtered. The freezedried extract powder was dissolved in dimethyl sulfoxide and centrifuged at 14,000 rpm for 10 min. The resulting supernatant was filtered.	Y- HPLC-UV/VIS	N/A	(144)
Roots of <i>Ribes</i> fasciculatum var. chinense MAX., [Human herb (Gyeongbuk, Korea)], 100 g No voucher specimen or batch number was mentioned	Y- Chopped with 70% ethanol solution under room temperature for 24 h and then filtered on a water bath under vacuo, frozen and lyophilized to yield ethanol extracts.	Y- HPLC	N/A	(54)
Roots of Sanguisorba officinalis L. (Rosacease), [Yeongcheon Oriental Herbal Market], 50 g No voucher specimen or batch number was mentioned	Y- Extracted using 70% ethanol in a 40°C shaking incubator for 24 h.	N	N/A	(146)

Barks of <i>Poria</i> cocos F.A. Wolf, [Plant Extract Bank of Korea]	Y- Extracted using 95% ethanol.	N	N/A	(6)
No information on the concentration of the herb				
No voucher specimen or batch number was mentioned				
Whole plants of Salvia plebeia R. Br. (Lamiaceae), [herbal store in Jeongeup, Korea], 30 kg	Y- Ground and extracted with 95% ethanol for 2 weeks at room temperature. The ethanol extract was then concentrated with a rotary evaporator under reduced pressure to give a dried residue.	Y- HPLC	N/A	(20)
(number WSP-11-114) was deposited at the Herbarium of the College of Pharmacy, Woosuk University				
Tubers of <i>Helianthus</i> tuberosus L., [Yangwonfood (Cheonan, Korea)]	Y- Extracted with 30% ethanol then filtered, lyophilized, and stored at 4°C.	N	N/A	(58)
No information on the concentration of the herb				
No voucher specimen or batch number was mentioned				
Tabebuia avellanedae Lorentz ex Griseb., [Nutribiotech Co. Ltd. (Seoul, Korea)]	N	N	N/A	(108)
No information on the part of the herb and the concentration of the herb				
No voucher specimen or batch number was mentioned				

Artemisia argyi H. Lév. & Vaniot, [Omniherb Co., Ltd. (Daegu, South Korea)], 100 g	Y- Mixed with 75% ethanol at 60°C, incubated for 24 h and filtered.	N	N/A	(34)
No information on the part of the herb				
Voucher specimen (DKMP-201203-AAFE) was deposited at Korean Medical Physiology Laboratory, Dong-Eui University				
Whole plant of Persicaria tinctoria (Aiton) Spach, [cultivated traditionally in Naju, republic of Korea]	Y- Extracted with distilled water at 80°C for 3 h and filtered. The filtered extracts were lyophilized and reduced to powder.	Y- HPLC	N/A	(35)
No information on the concentration of the herb				
No voucher specimen or batch number was mentioned				
Angelica dahurica (Hoffm.) Benth. & Hook.f. ex Franch. & Sav., [Han-poong Pharm Co., Ltd (Jeonjoo, Republic of Korea)]	N	N	N/A	(69)
No information on the part of the herb and the concentration of the herb				
No voucher specimen or batch number was mentioned				
Ixeris dentata (Thunb.) Nakai (Compositae), [Wonkwang Institute of Biomedical Engineering Research]	Y- Decocting the herbs with distilled water for 3 h and filtered and kept at 4°C.	N	N/A	(49)

No information on the part of the herb and the concentration of the herb				
Voucher specimen (No. 05-17-12) was deposited at the Herbarium of the College of Pharmacy, Wonkwang University				
Panax ginseng C.A. Mey., [Gansando agriculture cooperation, Jinan, Republic of Korea], 5 g	Y- Extracted with distilled water using a reflux condenser. The extract was then filtered and condensed under a vacuum.	N	N/A	(109)
No information on the part of the herb No voucher specimen or batch number was mentioned				
Roots of <i>Panax ginseng</i> Meyer, [Korea Ginseng Corporation (Daejeon, Korea)]	Y- Steaming fresh ginseng at 90-100°C for 3 h and then drying at 50-80°C, extracted with water for 8 h.	Y- HPLC	N/A	(60)
No information on the concentration of the herb				
No voucher specimen or batch number was mentioned				
Roots of <i>Polygala</i> tenuifolia Willd. [Oriental drug store (Dongwoodang Pharmacy Co., Ltd., Yeongcheon, Republic of Korea)], 120 g	Y- Immersed in distilled water, heated at 100°C, concentrated using a rotary evaporator and lyophilized using a freeze dryer.	Y- UPLC– ESI– MS	N/A	(126)
Voucher specimen (No. D0801130PTW) was deposited at the herbarium located in the college of Korean				

Medicine, Kyung Hee University				
Roots of <i>Liriope muscari</i> (Decne.) L.H. Bailey [plantations in the Miryang area (Korea)], 200 g Voucher specimens of LP (WPC-11-010) were deposited in the Functional Materials Bank of the PNU-Wellbeing RIS Center at Pusan National University	Y- Dry roots were steamed at 99°C for 3 h, airdried at 70°C for 24 h then reduced to powder, distilled water was added to the powder and the aqueous extracts were purified for 2 h at 100°C.	Y- HPLC	N/A	(72)
Dried whole plant of Saussurea lappa (Decne.) C.B. Clarke, [HMAX (Jecheon, Korea)], 100 g	Y- Extracted three times with 70% methanol (1 L) by refluxing for 90 min, then filtered, evaporated to dryness, and freeze-dried.	N	N/A	(87)
Voucher specimen (2009-KIOM62) has been deposited in the K- herb Research Center, Korea Institute of Oriental Medicine				
Dried whole plant of Saussurea lappa (Decne.) C.B. Clarke, [HMAX (Jecheon, Korea)], 100 g	Y- Extracted three times with 70% methanol (1 L) by refluxing for 90 min, then filtered, evaporated to dryness, and freeze-dried.	N	N/A	(86)
Voucher specimen (2009-KIOM62) has been deposited in the K- herb Research Center, Korea Institute of Oriental Medicine				
Stems of <i>Dendrobium</i> officinale Kimura & Migo, [green house of Chaoyang University ofTechnology]	Y- Dried stems of was grinded into fine powder, dissolved in methanol and sonicated at an amplitude of 15 for 10 min.	Y- LC/MS	N/A	(137)

No information on the concentration of the herb Voucher specimen number: (CMC DT 0303) has been deposited at the Institute of Chinese Pharmaceutical Sciences, China Medical				
University Dried whole plant of Patrinia scabiosifolia Link, [Korea Plant Extract Bank, Korea Research Institute of Bioscience and Biotechnology (Deajeon, Korea)], 33 g Voucher specimens (031 - 037) were deposited at the herbaria of the Department of the Herbal Pharmaceutical Development, Korea Institute of Oriental Medicine, Daejeon,	Y- Dried, powdered, and then underwent extraction with DMSO for 1 day at room temperature.	N	N/A	(13)
Byakkokakeishito • Gypsum (natural hydrous calcium sulfate), lot no. 7H16 M • Rhizome of Anemarrhena asphodeloides Bunge, 0E22 • Bark of the trunk of Cinnamomum verum J. Presl, 6K20 M • Seed of Oryza sativa L., 5G29	Y- Cutting or crushing of the whole crude drugs, boiled in 600 ml of distilled water for 60 min. The crude drugs were then removed from the paper bags and the decoctions were lyophilized.	Y- HPLC	N/A	(90)

[all purchased from Daiko Shoyaku (Nagoya, Japan)]				
• Root and stolon of <i>Glycyrrhiza uralensis</i> Fisch. ex DC., [Tsumura (Tokyo, Japan)], 23040711				
No information on the concentration and ratio of the herbs				
Qingre-Qushi • Hedyotis diffusa Willd. • Taraxacum (no specific species was mentioned) • Xanthium sibiricum Patrin ex Widder • Sophora flavescens Aiton	Y- The four herbs (4:4:1:4) were soaked in water and boiled at 100°C twice, 1 h each time then filtered through a Whatman number 2 filter paper, concentrated under vacuum conditions, and freeze- dried.	Y- HPLC	N/A	(14)
[Shuguang Hospital Affiliated with the Shanghai Chinese Medicine University]				
No information on the concentrations of each herb				
No voucher specimen or batch number was mentioned				
Yu-Ping-Feng-San • Astragalus mongholicus Bunge; [Zhangzhou Fushun Pharmaceutical	Y- Herbal mixture (RA: RAM: RS in a 3:1:1) were extracted twice with boiling water and further evaporated to crude extracts.	Y- HPLC	N/A	(148)

Co., Ltd. (Inner
Mongolia,
China)]

- Atractylodes macrocephala
 Koidz. [Shanxi
 Wanhui
 Pharmaceutical
 Co., Ltd.
 (Zhejiang,
 China)]
- Saposhnikovia divaricata (Turcz. ex Ledeb.)
 Schischk.,
 [Herbal Decoction Slices division of Nanjing Pharmaceutical Company (Heilongjiang, China)]

No information on the concentrations of each herb

No voucher specimen or batch number was mentioned

Yu-Ping-Feng-San	Y- Immersed (3:1:1) in ethanol	Y- HPLC	N/A	(138)
 Astragalus 	for 1 h and then refluxed for 2			
mongholicus	h. The extraction process was			
Bunge;	repeated twice and the extracts			
[Zhangzhou	were combined, filtered, and			
Fushun	evaporated to dryness using a			
Pharmaceutical	vacuum concentrator system.			
Co., Ltd. (Inner				
Mongolia,				
China)],				
20150601				

•	Atractylodes
	macrocephala
	Koidz., [Shanxi
	Wanhui
	Pharmaceutical
	Co., Ltd.
	(Zhejiang,
	China)], 150102

• Saposhnikovia divaricata (Turcz. ex Ledeb.)
Schischk.,
[Herbal Decoction Slices division of Nanjing Pharmaceutical Company (Heilongjiang, China)],
20150601

No information on the concentrations of each herb

Pentaherbs

- Lonicera japonica Thunb., 5 g
- Mentha canadensis L., 2.5 g
- Paeonia suffruticosa Andrews, 5 g
- Atractylodes lancea (Thunb.) DC., 5 g
- Phellodendron amurense Rupr., 5 g

No information of the supplier and the ratio of the herbs

Y- Extracted by refluxing in boiling water at 100°C for 1h. Extraction was repeated twice to obtain total water crude extract. The filtered extract was freeze-dried into powder.

Y- HPLC N/A (131)

batch number was mentioned				
Hataedock • Glycine max (L.) Merr., [Namyoung Pharm (Muju, Republic of Korea)], 100 g • Artemisia caruifolia Buch Ham. ex Roxb. • Morus alba L.	Y- Herbs (1:1) were boiled for 3 hours in distilled water and then filtered, concentrated by using a rotatory vacuum evaporator and then freezedried to obtain the extract.	Y- HPLC	N/A	(64)
No information of the supplier and the concentration of the herbs				
No voucher specimen or batch number was mentioned				
Hataedock • Coptis japonica (Thunb.) Makino, [Omniherb (Yeongcheon, Korea)], 100 g • Glycyrrhiza uralensis Fisch. ex DC., [Omniherb (Yeongcheon, Korea)], 100 g	Y- Herbs (1:1) were powdered and boiled for 3 h in distilled water then the resulting extract was filtrated, concentrated and freeze-dried.	Y- HPLC	N/A	(12)
Hataedock • Coptis chinensis Franch., 100 g • Glycyrrhiza uralensis Fisch. ex DC., 100 g No information of the	Y- Herbs (1:1) were decocted in distilled water for 3 hours and then filtered; after concentrating, the filtrate was freeze-dried.	Y- HPLC	N/A	(11)

No voucher specimen or batch number was mentioned				
 Bupleurum falcatum Linne Pinellia ternate Breitenbach Zingiber officinale Roscoe Scutellaria baicalensis Georgi Panax ginseng C.A. Meyer Glycyrrhiza uralensis Fischer Zizyphus jujube Miller var. inermis Rehder [Omniherb (Korea) and 	Y- Extracted in distilled water at 100°C for 120 min, the extract was filtered through a standard sieve, evaporated and freeze-dried into powder.	Y- UPLC	N/A	(79)
HMAX (China)] Voucher specimens (2008–KE26–1 ~ KE26– 7) have been deposited at the K-herb Research Center, Korea Institute of Oriental Medicine No information on concentration and ratio of				
the herbs Gamisasangja-tang • Stemona sessilifolia (Miq.) Miq. • Spirodela polyrhiza (L.) Schleid. • Cnidium monnieri (L.) Cusson	Y- Boiling the herbs (1:1:1:1:0.5:0.5) in distilled water at 100°C for 2h. The boiled herbs were then filtered through a Whatman no.2 filter, concentrated under vacuum conditions, and freeze-dried.	Y- HPLC	N/A	(106)

Schleid. • Cnidium monnieri (L.) Cusson	concentrated under vacuum conditions and freeze-dried.			
• Spirodela polyrhiza (L.)	were then filtered through a Whatman no. 2 filter,			
• Stemona japonica (Blume) Miq.	(1:1:1) in distilled water at 100°C for 2 h. The boiled herbs			
herbs SSC201	Y- Boiling the three herbs	Y- HPLC	N/A	(105)
No information on concentration of the				
Daejeon, Korea				
of Oriental Medicine, Daejeon University,				
deposited at the College				
Voucher specimens (No. 2014-004 to 009) were				
Co., Ltd. (Seoul, Korea)]				
[Dong Kyung Pharm				
mandshurica (Rupr.) Ohwi				
terniflora var.				
• Clematis				
 Angelica gigas Nakai 				
flavescens Aiton				

- Rehmannia glutinosa (Gaertn.) DC.
- Betula platyphylla var. japonica
- Rubus coreanus Mig.

[Hanpoong Pharmaceutical (Jeon-ju, Korea)]

No information on concentration and ratio of the herbs

No voucher specimen or batch number was mentioned

Huang-Lian-Jie-Du

- Coptis chinensis Franch.
- Scutellaria baicalensis Georgi
- Phellodendron chinense C.K. Schneid.
- *Gardenia* jasminoides J. Ellis

[Zhixin Herbal Pharmaceutical Company Ltd]

Voucher specimen (no. AD01–04) were deposited in the Herbarium of the School of Chinese Medicine, CUHK

Y- Herbs (3:2:2:3) were ground to powder or pieces and mixed. The mixture was extracted with 80% aqueous ethanol in an ultrasonic bath for 30 min and then was filtered.

Y- HPLC N/A (15)

No information on concentration of the herbs				
Sanpaocao • Cardiospermum halicacabum L., 200 g • Physalis angulata L., 200 g • Ludwigia adscendens (L.) H. Hara, 200 g [Haikou, Hainan Province, China] No voucher specimen or batch number was	Y- Herbs (1:1:1) were extracted with 70% ethanol using maceration for 3 days with 90 rmp shaking at 40°C, repeated three times. The extract was filtered through a cotton bed followed by Whatman No. 1 filter paper. The filtrate was evaporated under reduced pressure at 45°C using a Buchii Rotaroy Evaporator to leave a gummy concentrate and through vacuum drying.	N	N/A	(121)
mentioned BuShenYiQi • Astragalus mongholicus Bunge, 1204811 • Epimedium sagittatum (Siebold & Zucc.) Maxim, 1111018 • Rehmannia glutinosa (Gaertn.) DC., 1203813	Y- Soaked in water for 2h and extracted twice with H ₂ O at 100°C for 1h, the filtered liquid was concentrated to a clear cream. After spray drying, the spray powder was mixed homogeneously and made into mesh particles.	Y- LC-ESI-Q- TOF-MS	N/A	(68)
[Anhui Bencao ChineseMedicine Herb Pieces Co., Ltd (Bozhou, China)]				
concentration and the ratio of the herbs				
Yupingfeng	N	N	N/A	(147)

Astragalus mongholicus Bunge Atractylodes macrocephala Koidz. Saposhnikovia divaricata (Turcz. ex Ledeb.) Schischk. [Guangdong HuanQiu Pharmaceutical Company] No information on the concentration and ratio of the herbs No voucher specimen or batch number was mentioned PTQX Y- UHPLC-LTQ-N N/A (139)Atractylodes Orbitrap-MS macrocephala Koidz., 10 g Pseudostellaria heterophylla (Miq.) Pax, 10 g Dioscorea oppositifolia L., 15 g Coix lacrymajobi var. ma-yuen (Rom.Caill.) Stapf, 20 g *Imperata* cylindrica (L.) P. Beauv., 15 g Forsythia suspensa (Thunb.) Vahl, 10

Dictamnus dasycarpus Turcz., 10 g

- Margarita, 0.3g (no specific species)
- Glycyrrhiza glabra L., 5 g

[Jiangyin Tian Jiang Pharmaceutical Co., Ltd. (Jiangsu, China)]

The formula has been patented for the treatment of AD (patent no.: ZL2013 1 0328668.4, China

Taeumjowi-tang

(107) Y- UPLC N/A N

- Coix lacrymajobi var. ma-yuen (Rom.Caill.) Stapf
- Castanea crenata Siebold & Zucc.
- Raphanus raphanistrum subsp. sativus (L.) Domin
- Schisandra chinensis (Turcz.) Baill.
- Platycodon grandiflorus (Jacq.) A. DC.
- Acorus gramineus Aiton
- Ephedra sinica Stapf
- Liriope muscari (Decne.) L.H. Bailey

[I-World Pharm. Co. (Incheon, Republic of Korea)]

No information on the concentration and the ratio of the herbs No voucher specimen or batch number was mentioned Jianpi Chushi Y-Boiled for 30 min, filtrated N/A N (133)Poria cocos F.A. and concentrated. Wolf, 20 g Dioscorea oppositifolia L., 20 g Dictamnus dasycarpus Turcz., 20 g Zaocys dhumnade, 15 g (no information in MPNS) Atractylodes lancea (Thunb.) DC., 15 g Citrus aurantium L., 15 g • Scutellaria baicalensis Georgi, 10 g Sophora flavescens Aiton, 10 g Paeonia lactiflora Pall., 10 Glycyrrhiza glabra L., 5 g [Tibet Qizheng Tibetan Medicine Ltd.] No information on the ratio of the herbs

No voucher specimen or

batch number was

mentioned

Calycosin (pure compound)	N/A	N/A	≥ 98.75%	(129)
[Tianjin Marker Bio- Tech Co. Ltd. (Tianjin, China)]				
No information on the concentration of the compound				
Physcion (pure compound)	N/A	N/A	≥ 98%	(96)
[Cayman Chemical Company (Ann Arbor, MI, USA)]				
No information on the concentration of the compound				
Chrysophanol	N/A	N/A	N	(37)
[Sigma Chemical Co.]				
No information on the concentration of the compound				
Cimifugin (pure compound)	N/A	N/A	≥ 99%	(135)
[National Institutes for Food and Drug Control (Shanghai, China)]				
No information on the concentration of the compound				
Esculetin, 20 mg (112.27μMol)	Y- Refluxed twice with 95% ethanol for 2 h. The extracted solution was filtered through a	Y- NMR	N	(50)
[local market (Daegu, Republic of Korea)]	filter paper. The filtrate was concentrated to dryness by a rotary evaporator under reduced pressure below 40°C. The extract was suspended in			

distilled water and successively partitioned. 5 fractions were rechromatographed by using silica gel to yield six subfractions.			(2.1)
N	N	N	(94)
Y- 4-year-old ginseng was ground into small pieces and refluxed with 70% fermented ethanol eight times for 8 h each at 80°C. The extracts were concentrated, dissolved in distilled, cold water and stored at 4°C for 24–96 h. The supernatant and precipitate of water fractionation after ethanol extraction of ginseng was separated by centrifugation. The precipitate after centrifugation was lyophilized.	N	N	(75)
N	N	N	(36)
N	N	≥98%	(5)
	y- 4-year-old ginseng was ground into small pieces and refluxed with 70% fermented ethanol eight times for 8 h each at 80°C. The extracts were concentrated, dissolved in distilled, cold water and stored at 4°C for 24–96 h. The supernatant and precipitate of water fractionation after ethanol extraction of ginseng was separated by centrifugation. The precipitate after centrifugation was lyophilized.	partitioned. 5 fractions were rechromatographed by using silica gel to yield six subfractions. N N N N N N N N N N N N N	partitioned. 5 fractions were rechromatographed by using silica gel to yield six subfractions. N N N N N N N N N N N N N N N

N/A: Not applicable or the articles did not mention the parts of the plant used nor the concentration and purity, ESI: electrospray ionization, MS: mass spectrometry, UPLC: ultra-performance liquid

chromatography, HPLC: high-performance liquid chromatography, QTOF: Quadrupole Time-of-Flight, NMR: nuclear magnetic resonance, MPNS: Medicinal Plant Names Services