Supporting information

Metabolomics of artichoke bud extract in spontaneously hypertensive rats

Zhi-Bin Wang^{1,2}, Shi-Long Jiang^{1,2}, Shao-Bo Liu³, Jing-Bo Peng^{1,2}, Shuo Hu⁴, Xu Wang^{1,2}, Wei Zhuo^{1,2}, Tong Liu^{1,2}, Ji-Wei Guo^{1,2}, Hong-Hao Zhou^{1,2}, Zhi-Quan Yang^{5*}, Xiao-Yuan Mao^{1,2*} and Zhao-Qian Liu^{1,2*}

¹Department of Clinical Pharmacology, Hunan Key Laboratory of Pharmacogenetics, and National Clinical Research Center for Geriatric Disorders, Xiangya Hospital, Central South University, Changsha 410008, P. R. China; ²Institute of Clinical Pharmacology, Engineering Research Center for applied Technology of Pharmacogenomics of Ministry of Education, Central South University, Changsha 410078, P. R. China; ³Department of Pharmacy, Xiangya Hospital, Central South University, Changsha, 410008, P. R. China; ⁴Department of Nuclear Medicine and Key Laboratory of Biological Nanotechnology of National Health Commission, Xiangya Hospital, Central South University, Changsha 410008, P. R. China; ⁵Department of Neurosurgery, Xiangya Hospital, Central South University, Changsha 410008, P. R. China.

To whom correspondence should be addressed: Professor Zhao-Qian Liu, Department of Clinical Pharmacology, Hunan Key Laboratory of Pharmacogenetics, Xiangya Hospital, Central South University, Changsha 410008; P. R. China. Tel: +86 731 89753845, Fax: +86 731 82354476, E-mail: zqliu@csu.edu.cn.

Or Professor Xiao-Yuan Mao, Department of Clinical Pharmacology, Hunan Key Laboratory of Pharmacogenetics, Xiangya Hospital, Central South University, Changsha 410008; P. R. China. E-mail: xiaoyuanm@csu.edu.cn.

Or Professor Zhi-Quan Yang, Department of Neurosurgery, Xiangya Hospital, Central South University, Changsha 410008, P. R. China. E-mail: y66406914@163.com.

Supporting Information

The toxicity assessment of ABE; log2 fold changes of metabolites level mapped onto the KEGG pathway module; the chromatograms of artichoke bud extract that used in this study and the standards (chlorogenic acid, cynarin); chemical structures of active components contained in ABE; sequences of primers used for qRT-PCR.



Figure S1 Toxicity assessment of ABE. SP change (A) and DP change (B) compared to 0 week of each rat in different groups. SP (C) and DP (D) trend along with time in different groups. n(BLK)=6; n(TC)=5. ns, $P \ge 0.05$; *P < 0.05; **P < 0.01; ***P < 0.001; ***P < 0.001 (ANOVA with repeated measurments). Effect of ABE on the serum biochemical parameters of WKY rats liver function (E), kidney function (F) and serum lipid profile (G). The bars at each data point in C and D indicate standard deviation. SP, Systolic blood pressure; DP, diastolic blood pressure; BLK, blank; TC, toxicity control, 50 mg/kg/d artichoke bud extract; TP, total protein; ALB, albumin; GLB, globulin; TBIL, total bilirubin; DBIL, direct bilirubin; TBA, total bile acids; ALT, alanine aminotransferase; AST, aspartate aminotransferase; UA, uric acid; CRE, creatinine; TG, triglyceride; TC, cholesterol; HDL-C, high-density lipoprotein cholesterol c; LDL-C, low-density lipoprotein cholesterol c.



B



Figure S2 Log2 fold changes of metabolites level mapped onto the KEGG pathway module (A) "Purine metabolism" and (B) "Nicotinate and nicotinamide metabolism" by R package "Pathview". Left half of circle, Group HPM vs. Group BLK; right half of circle, Group ABE_a vs. Group HPM.



Figure S3 The chromatograms (UV 330 nm) of artichoke bud extract that used in this study (A) and the standards (B for chlorogenic acid, C for cynarin).



Figure S4 Chemical structures of active components contained in ABE. A for chlorogenic acid, B for cynarin.

Name	Direction	Sequence (5'-3')
Actb-rat	Forward	GGGAAATCGTGCGTGACATT
	Reverse	GCGGCAGTGGCCATCTC
Prdx2-rat	Forward	CTTCGCCAGATCACAGTCAA
	Reverse	ATACTGAAAGGCCTGGACGA
Sod2-rat	Forward	ATTAACGCGCAGATCATGCA
	Reverse	CCTCGGTGACGTTCAGATTGT
Gpx4-rat	Forward	GCCGTCTGAGCCGCTTATT
	Reverse	ACGCAACCCCTGTACTTATCCA
Gsr-rat	Forward	ACTTCTCACCCCAGTTGCG
	Reverse	CCACGGTAGGGATGTTGTCA

Table S1 Sequences of primers used for qRT-PCR.