## **Electronic Supplementary Information**

## Neuroprotective effect of *Cyperi rhizome* against corticosterone-induced PC12 cells via suppress $Ca^{2+}$ overloading

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identified metabolites					
Metabolic pathway	Total	Expected	Hits	Raw p	Impact
Sphingolipid metabolism	21	0.074893	2	0.0020808	0.14286
Glycerophospholipid metabolism	30	0.10699	1	0.10265	0.04444
Steroidhormone biosynthesis	70	0.24964	1	0.22622	0.01699

Tabe S1 Summary of metabolic pathway analysis with MetaboAnalyst 3.0.based on the identified metabolites



-0.10<sup>1</sup> 0.00 1.00 2.00 3.00 4.00 5.00 6.00 7.00 8.00 9.00 10.00 11.00 12.00 13.00 14.00 15.00 16.00 17.00 18.00 19.00 20.00 21.00 22.00 23.00 24.00 25.00 26.00 27.00 28.00 29.00 30.0 Time

Fig. S1 UPLC chromatograms of the CR-95E, CR-50E and CR-W extracts.



**Fig S2.** Effect of *CR-50E* on corticosterone-induced antioxidants enzymes (SOD, CAT, and MDA) activity; Cells were exposed to 200  $\mu$ M of corticosterone in the absence or presence of *CR-50E* for 24h. Results are presented as means  $\pm$  SD (n=6). \*\*p < 0.01, compared with corticosterone-treated group (Cort), #p < 0.05 or ##p < 0.01, compared with control group.



**Fig. S3** Base peak intensity (BPI) chromatograms of UPLC-Q-TOF/MS in positive ion mode from the PC12 cell samples in each group. A: control group; B: corticosterone-treated group; C: High dose CR-50E-treated group.



**Fig. S4** Summary of metabolic pathway analysis with MetaboAnalyst 3.0. Each point represents one metabolic pathway; the size of dot and shades of color is in positive correlation with the impaction of the metabolic pathway.