**Electronic Supplementary Information for** 

## Superior Solubility of Anhydrous Quercetin and Polymer Physical Mixtures Compared to Amorphous Solid Dispersions

Xu Ma, <sup>a,b</sup> Hong Su, <sup>b</sup> Yongming Liu,<sup>c</sup> Fenghua Chen,<sup>b,\*</sup> and Rongrong Xue <sup>b,\*</sup>

<sup>a</sup> College of Chemistry and Materials Science, Fujian Normal University, Fuzhou 350117, Fujian,

China.

<sup>b</sup> School of Resources and Chemical Engineering, Sanming University, Sanming 365004, Fujian,

China.

<sup>c</sup> School of Education and Music, Sanming University, Sanming 365004, Fujian, China.

Corresponding Authors \*(F. C.) E-mail: fenghuachen@fjsmu.edu.cn. \*(R. X.) E-mail: rongrongxue@fjsmu.edu.cn.



**Figure S1.** Macro-thermogravimetric curve of QUE-DH at 150 °C in air. The weight loss reaches 9.7% in 15 minutes, which is the same as that of QUE-DH heated at 100 °C for 1 hour. The dehydration product is QUE-AH. The weight loss is up to 10.0% in 50 minutes, which indicates that a small amount of QUE molecules undergo degradation. Because the weight loss in the process was measured by macroscopic thermogravimetric, a 0.3 % difference is enough to judge the occurrence of degradation.



Figure S2. Hydration process of QUE-AH at room temperature and  $\sim 60\%$  humidity.



Figure S3. UV-vis spectra of QUE aqueous solution with different storage time.



**Figure S4.** UV-vis spectra of the QUE aqueous solutions in the concentration range of (a)  $1.8 - 12.5 \ \mu g \cdot ml^{-1}$  and (b)  $13.8 - 17.8 \ \mu g \cdot ml^{-1}$ . The UV-vis spectra exhibit a good gradient increase in the concentration range of  $1.8 - 12.5 \ \mu g \cdot ml^{-1}$ . The absorption of QUE aqueous solutions with concentration of 13.8 and 17.8  $\ \mu g \cdot ml^{-1}$  decreased significantly at 367 nm, while the baseline absorption increased significantly at 400 – 500 nm, indicating the occurrence of recrystallization.



Figure S5. Working curve establishment of QUE 50 vol% ethanol solutions in the concentration

range of  $0 - 49.4 \ \mu g \cdot ml^{-1}$ . (a) UV-vis spectra, (b) the fitted working curve.



**Figure S6.** PXRD pattern of commercial QUE-AH stored for years. The most PXRD peaks are related to QUE-DH, and the peak of QUE-AH is marked.



Figure S7. IR spectra of QUE polymorphs in the range of 2600 - 3600 cm<sup>-1</sup>.



Figure S8. Low-frequency Raman spectra (in the range of 50 - 300 cm<sup>-1</sup>) of QUE polymorphs.

QUE polymorphs			
QUE-AM	QUE-AH	QUE-DH	assignments
519	521	522	$\gamma(A) + \gamma(C) + \delta(COH)_A$
598	597	603	$\gamma(OH)_{C3} + \gamma(A) + \gamma(C) + \gamma(OH)_{A5}$
1319	1321	1327	$\nu(A) + \nu(C) + \delta(CH)_{A6} + \nu(C-O)_{A5}$
1558	1553	1547	$\delta(CH)_{B2'} + \nu(B) + \nu(C-O)B_{4'}$
1612	1613	1606	$\nu(B) + \delta(CH)_{B5'} + \nu(C=O)$

Table S1. Raman bands (cm<sup>-1</sup>) and assignments of quercetin (QUE).<sup>1</sup>

1. Hanuza, J.; Godlewska, P.; Kucharska, E.; Ptak, M.; Kopacz, M.; Mączka, M.; Hermanowicz, K.; Macalik, L., Molecular structure and vibrational spectra of quercetin and quercetin-5'-sulfonic acid. *Vibrational Spectroscopy* **2017**, *88*, 94-105.



Figure S9. UV-vis spectra of 1 mg·ml<sup>-1</sup> PVP and 1 mg·ml<sup>-1</sup> Soluplus solutions.



Figure S10. (a) IR and (b) Raman spectra of QUE ASDs with PVP (BM-PVP- $\omega_{QUE}$ ) and PVP.



Figure S11. Fluorescence spectra of QUE-DH and QUE-AH. ( $\lambda_{ex}$ =350 nm)