

## Supporting Information for Publication

### The Novel Anti-Hepatitis B Virus Activity of *Euphorbia schimperi* and its Quercetin and Kaempferol Derivatives

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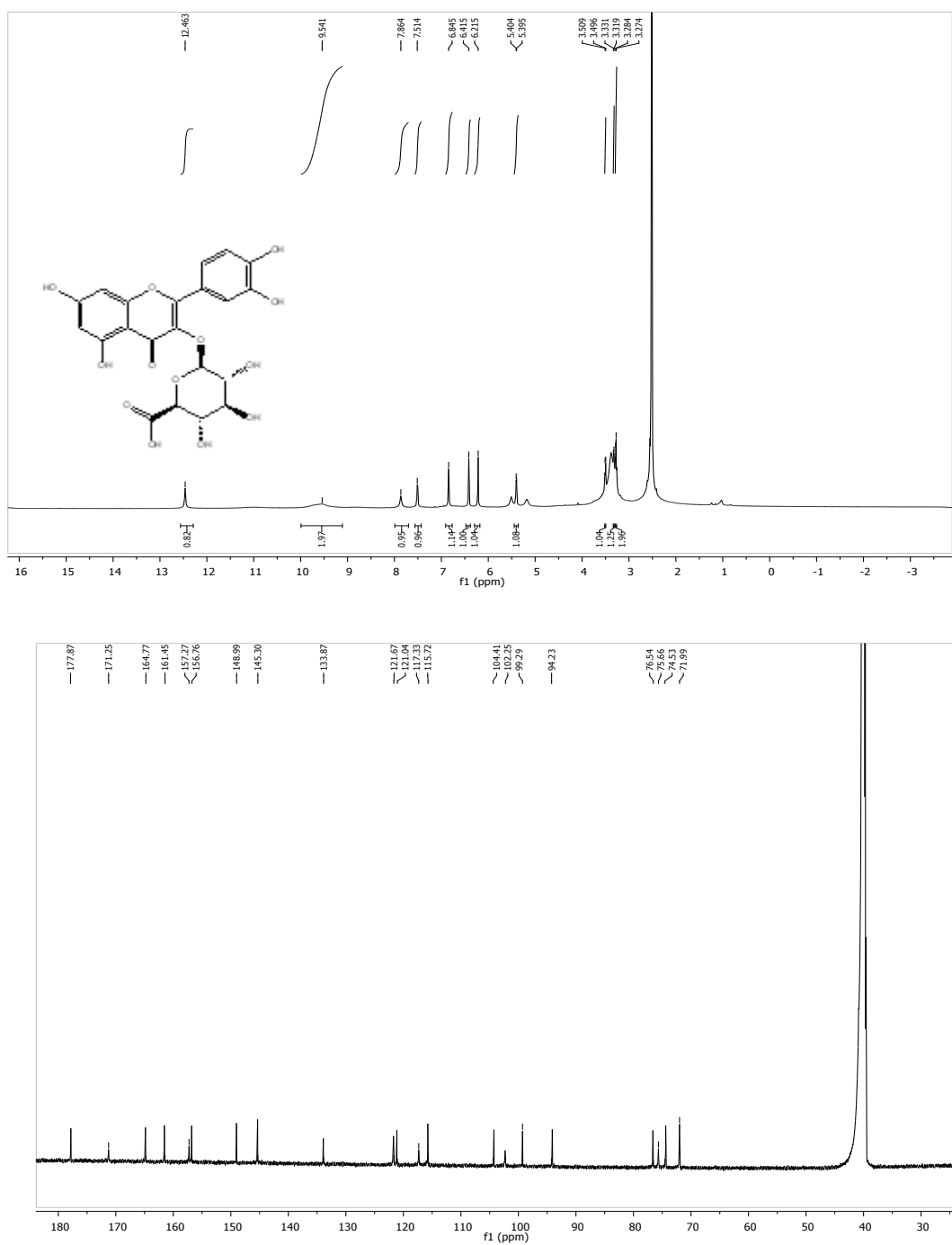
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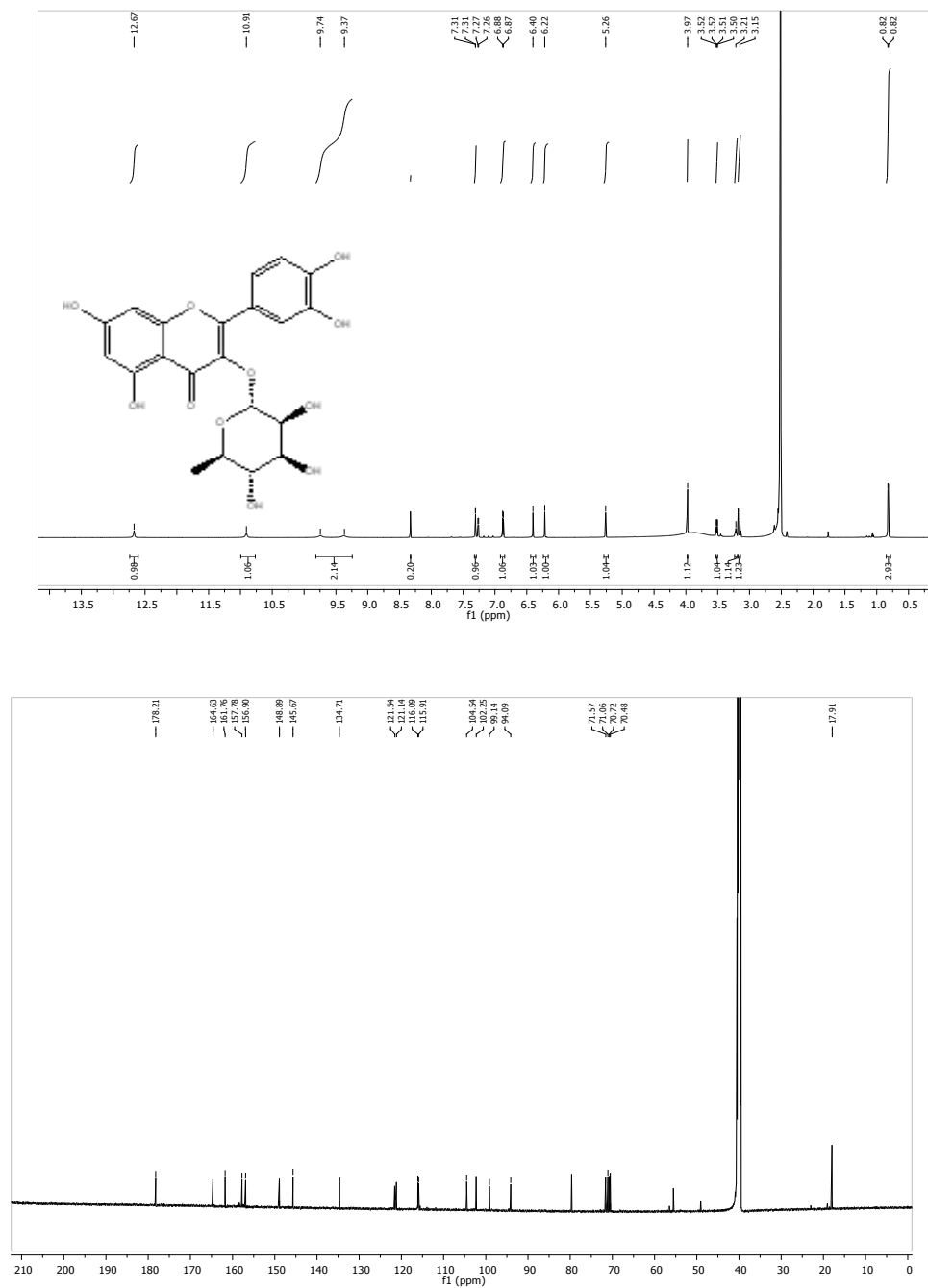
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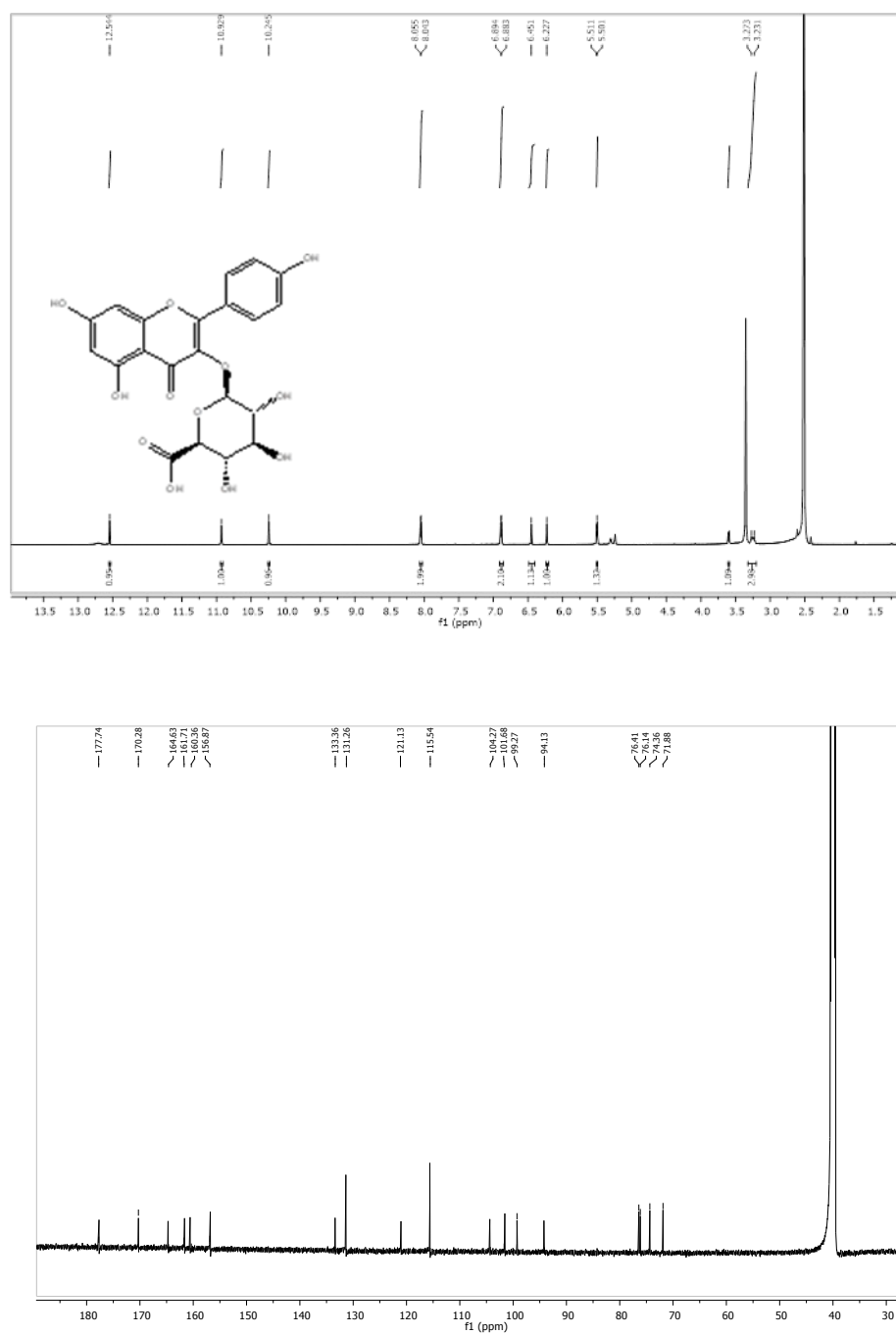
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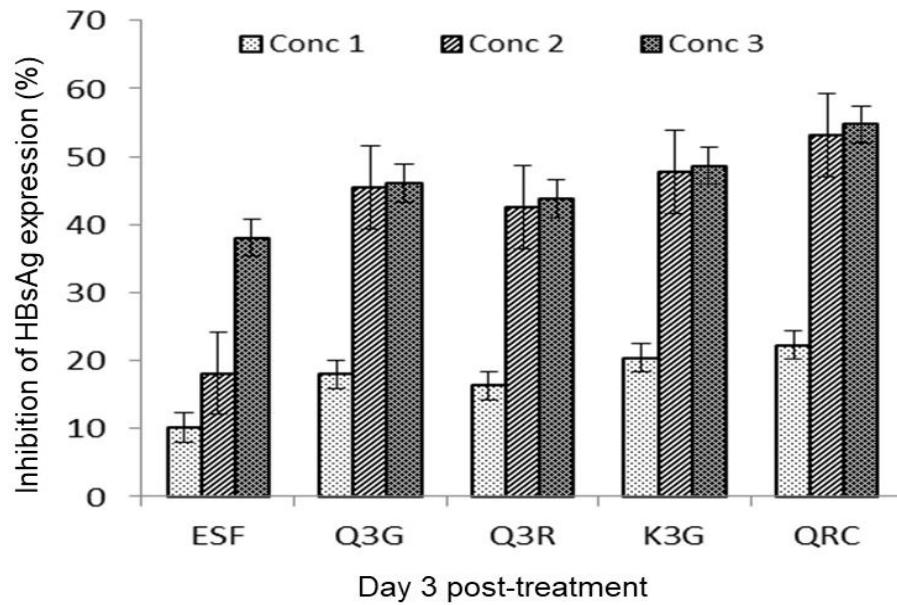
**Figure S1.** The <sup>1</sup>H NMR (700 MHz, DMSO-d<sub>6</sub>; upper panel) and <sup>13</sup>C NMR (175 MHz, DMSO-d<sub>6</sub>; lower panel) spectrum of *E. schimperi* derived quercetin 3-*O*-glucuronide.



**Figure S2.** The <sup>1</sup>H NMR (700 MHz, DMSO-d<sub>6</sub>; upper panel) and <sup>13</sup>C NMR (175 MHz, DMSO-d<sub>6</sub>; lower panel) spectrum of *E. schimperi* derived quercetin 3-O-rhamnoside.



**Figure S3.** The <sup>1</sup>H NMR (700 MHz, DMSO-d<sub>6</sub>; upper panel) and <sup>13</sup>C NMR (175 MHz, DMSO-d<sub>6</sub>; lower panel) spectrum of *E. schimperi* derived kaempferol-3-*O*-glucuronide.



**Figure S4.** Dose-dependent inhibitory effect of *E. schemperi* fraction (ESF), quercetin-3-*O*-glucuronide (Q3G), quercetin-3-*O*-rhamnoside (Q3R) and kaempferol-3-*O*-glucuronide (K3G) on HBsAg expression at 48 h post-treatment. ESF (Conc 1: 25  $\mu$ g/ml; Conc 2: 50  $\mu$ g/ml; Conc 3: 100  $\mu$ g/ml). Q3G, Q3R and K3G (Conc 1: 6.25  $\mu$ g/ml; Conc 2: 12.5  $\mu$ g/ml; Conc 3: 25.0  $\mu$ g/ml). Quercetin (QRC; Conc 1: 6.25  $\mu$ g/ml; Conc 2: 12.5  $\mu$ g/ml; Conc 3: 25.0  $\mu$ g/ml) was used as positive control. Values on Y-axis are means of three determinations.

**Table S1.** Docking energies (kcal/mol) of the tested flavonols and standard drugs.

<i>Ligands</i>	<i>Target</i>	
	HBV-Pol/RT	HBV-Core
Quercetin	-8.309	-8.736
Quercetin-3- <i>O</i> -glucuronide*	-9.313	-7.443
Quercetin-3- <i>O</i> -rhamnoside*	-6.493	-9.001
Kaempferol	-8.976	-9.128
Kaempferol-3- <i>O</i> -glucuronide*	-9.197	-8.178
Kaempferol-3- <i>O</i> -rhamnoside	-9.382	-8.212
Lamudivine triphosphate	-9.153	-
Heteroaryldihydropyrimidine	-	-8.876

\**E. schimperi* derived