## **Supplementary Material**

## Immune Checkpoint PD-1/PD-L1 CTLA-4/CD80 are Blocked by *Rhus verniciflua* Stokes and its Active Compounds

## Wei Li,<sup>1,#</sup> Tae In Kim,<sup>1,#</sup> Ji Hye Kim,<sup>1</sup> and Hwan-Suck Chung <sup>1</sup>,\*

<sup>#</sup> These authors contributed equally to this work.

\* Correspondence: hschung@kiom.re.kr (H.S.C); Tel: (+82) 53 940 3875; Fax: (+82) 53 940 3899

<sup>&</sup>lt;sup>1</sup> Korean medicine (KM)-Application Center, Korea Institute of Oriental Medicine (KIOM), Daegu 41062, Republic of Korea; hschung@kiom.re.kr (H.S.C); liwei1986@kiom.re.kr (W.L.); tikim@kiom.re.kr (T.I.K); jkim2903@kiom.re.kr (J.H.K)



S1. Structures of compounds 1-20 from RVS.



**S2.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **1** in methanol- $d_4$  (600 and 150 MHz)



**S3.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **2** in methanol- $d_4$  (600 and 150 MHz)



**S4.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **3** in methanol- $d_4$  (600 and 150 MHz)



**S5.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **4** in methanol- $d_4$  (600 and 150 MHz)



**S5.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **4** in methanol- $d_4$  (600 and 150 MHz)



**S6.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **5** in methanol- $d_4$  (600 and 150 MHz)



**S7.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **6** in methanol- $d_4$  (600 and 150 MHz)



**S8.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **7** in methanol- $d_4$  (600 and 150 MHz)



**S9.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **8** in methanol- $d_4$  (600 and 150 MHz)



**S10.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **9** in methanol- $d_4$  (600 and 150 MHz)



**S11.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **10** in methanol- $d_4$  (600 and 150 MHz)



**S12.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **11** in methanol- $d_4$  (600 and 150 MHz)



**S13.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **12** in methanol- $d_4$  (600 and 150 MHz)



**S14.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **13** in methanol- $d_4$  (600 and 150 MHz)



**S15.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **14** in methanol- $d_4$  (600 and 150 MHz)



**S16.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **15** in methanol- $d_4$  (600 and 150 MHz)



**S17.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **16** in methanol- $d_4$  (600 and 150 MHz)



**S18.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **17** in methanol- $d_4$  (600 and 150 MHz)



**S19.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **18** in methanol- $d_4$  (600 and 150 MHz)



**S20.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **19** in methanol- $d_4$  (600 and 150 MHz)



**S21.** <sup>1</sup>H and <sup>13</sup>C NMR spectrum of compound **20** in methanol- $d_4$  (600 and 150 MHz)



S22. Blocking effect of PD-1/PD-L1 interaction of isolated compound 1–20 at 10µM.



S23. Blocking effect of CTLA4/CD80 interaction of isolated compound 1-20 at  $10\mu$ M.