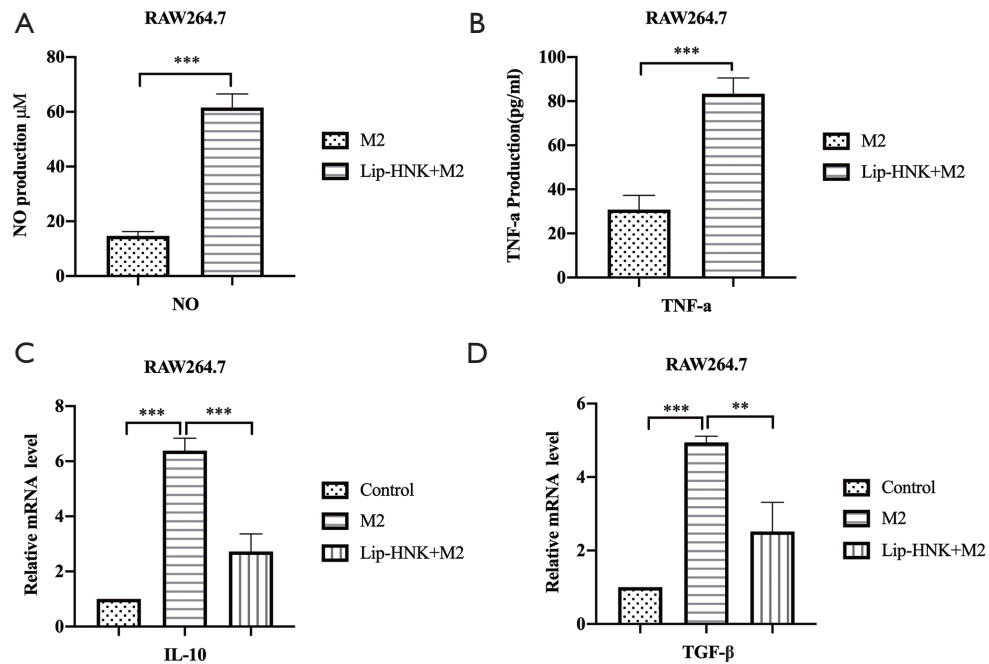
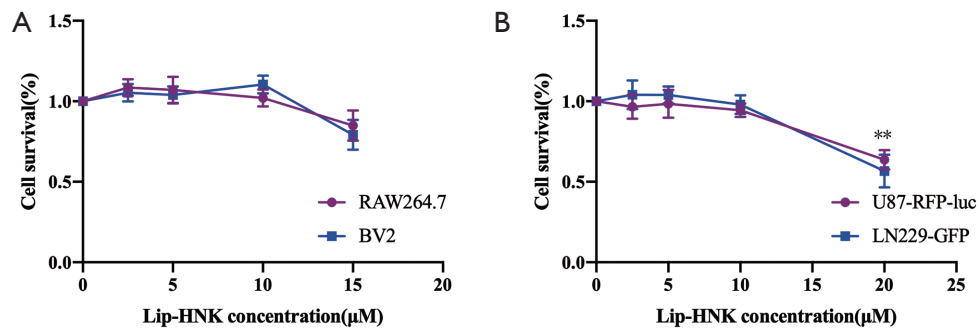


**Figure S1** The effect of liposomal honokiol (Lip-HNK) on lipopolysaccharide (LPS) + interferon  $\gamma$  (IFN- $\gamma$ )-mediated macrophage marker expression. The mRNA levels of tumor necrosis factor  $\alpha$  (TNF- $\alpha$ ) were detected by real-time reverse transcription polymerase chain reaction (RT-PCR) in RAW264.7 (A) and BV2 cells (B). \*,  $P < 0.05$ ; \*\*\*,  $P < 0.001$ .

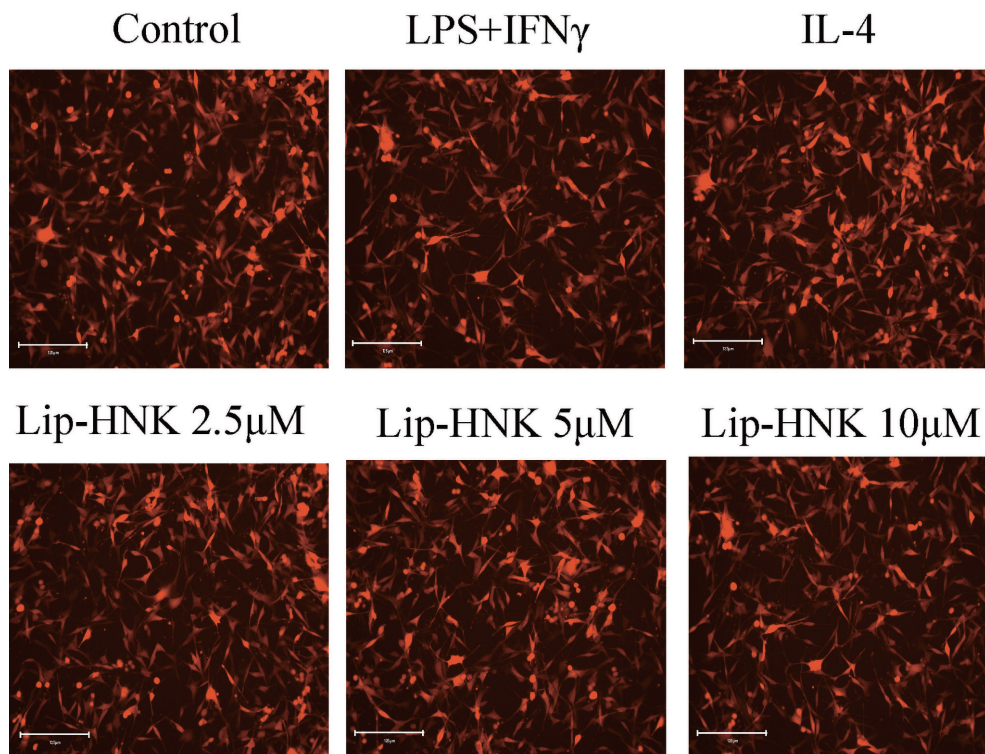


**Figure S2** The levels of M1 and M2 related markers in RAW264.7 cells after induced with interleukin (IL)-4 and liposomal honokiol (Lip-HNK) treatment. The produced levels of nitric oxide (NO) (A), tumor necrosis factor  $\alpha$  (TNF- $\alpha$ ) (B), and the relative mRNA expression of IL-10 (C) and transforming growth factor  $\beta$  (TGF- $\beta$ ) (D) were detected by real-time reverse transcription polymerase chain reaction (RT-PCR) in RAW264.7 cells. \*\*,  $P < 0.01$ ; \*\*\*,  $P < 0.001$ .

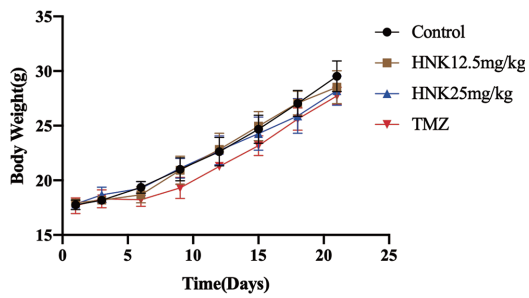


**Figure S3** The effect of liposomal honokiol (Lip-HNK) on the proliferation of tumor, and macrophage cells. (A) Macrophage cells were treated with different concentration of Lip-HNK for 48 h; (B) U87 and LN229 tumor cell lines were treated with different concentration of Lip-HNK for 48 h. \*\*,  $P < 0.01$ .

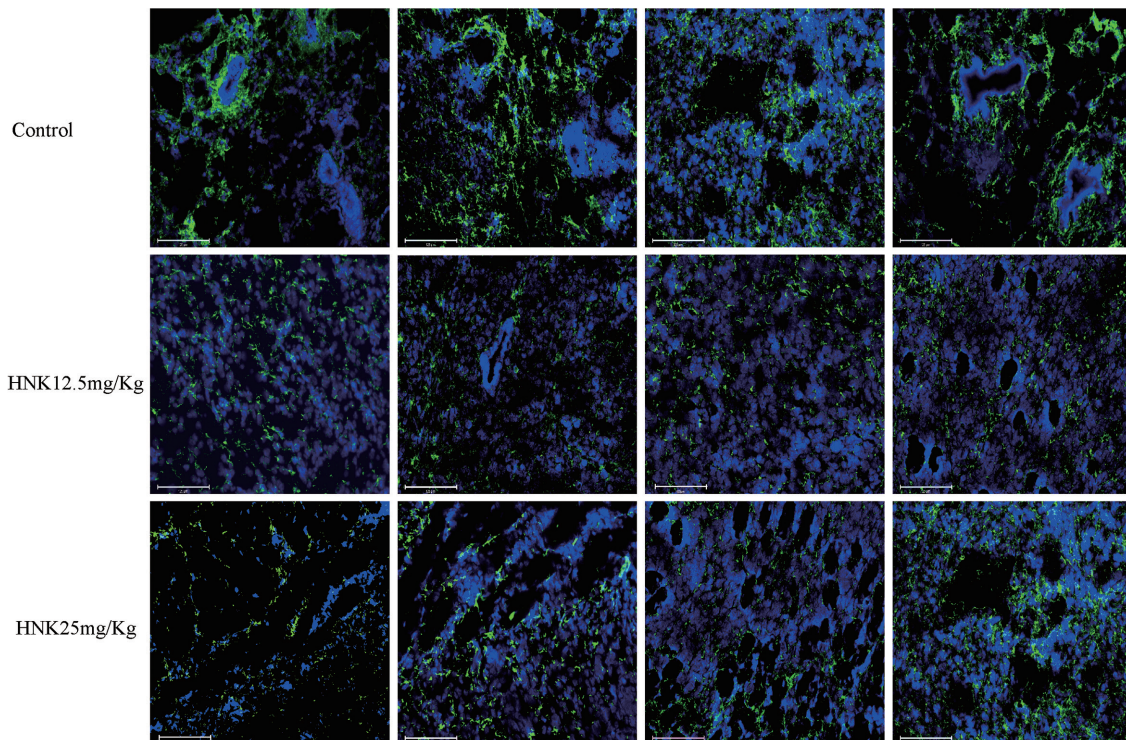
### U87-RFP-Luc cells



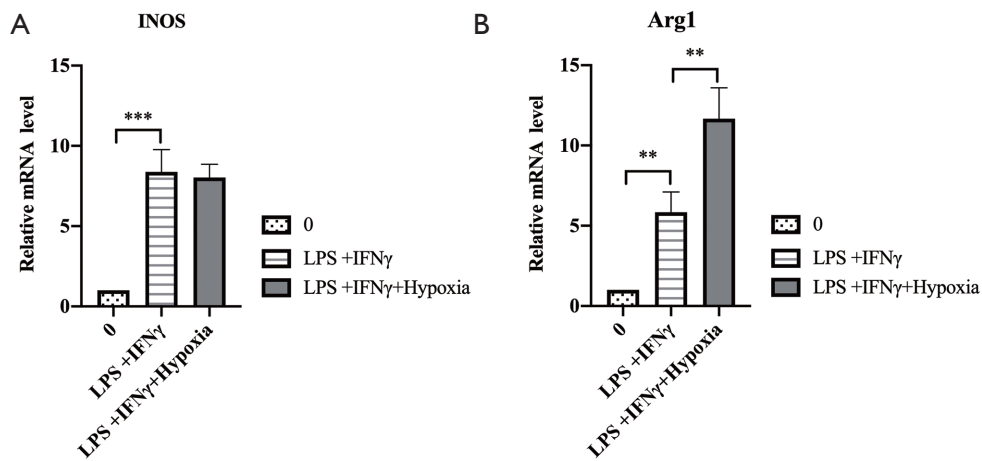
**Figure S4** The effect of liposomal honokiol (Lip-HNK) and M1/M2 inducer on the proliferation of U87-RFP-Luc tumor cells. U87-RFP-Luc glioma cells were treated with different concentration of Lip-HNK or interferon  $\gamma$  (IFN- $\gamma$ )/lipopolysaccharide (LPS) and interleukin 4 (IL-4). The morphological feature of tumor cells was examined under fluorescence microscopy.



**Figure S5** Effect of liposomal honokiol (Lip-HNK) on glioblastoma progression in G422 xenograft model. Body weight in individual mice were detected. TMZ, temozolomide.



**Figure S6** The expression of CD31 in liposomal honokiol (Lip-HNK) group and control group. Green fluorescence was used to detect the expression of CD31 in tumor tissue of G422 mice, blue fluorescence represents 4',6-diamidino-2-phenylindole (DAPI).



**Figure S7** The expression of macrophage related markers in lipopolysaccharide (LPS) + interferon  $\gamma$  (IFN- $\gamma$ ) (A) and interleukin 4 (IL-4) (B) group under hypoxia. \*\*,  $P < 0.01$ ; \*\*\*,  $P < 0.001$ .

**Table S1** The primer sequences used in the real time reverse transcription polymerase chain reaction (RT-PCR) analysis

| Gene  | Strands | Sequence                          |
|---|---------|-----------------------------------|
| Inducible nitric oxide synthase ( <i>iNOS</i> )                   | Forward | 5'-CTC AGC CCA ACA ATA CAA G-3'   |
|   | Reverse | 5'-CTA CAG TTC CGA GCG TCA-3'     |
| Arginase 1 ( <i>Arg1</i> )  | Forward | 5'- ATCAACTCTCCCCTGACAACC-3'      |
|   | Reverse | 5'- CGCAAGCCAATGTACACGAT-3'       |
| <i>GAPDH</i>  | Forward | 5'-GGT TGT CTC CTG CGA CTT CA-3'  |
|   | Reverse | 5'-TGG TCC AGG GTT TCT TAC TCC-3' |
| Tumor necrosis factor alpha ( <i>TNF-<math>\alpha</math></i> )    | Forward | 5'-ATGAGCACAGAAAGCATGATC-3'       |
|   | Reverse | 5'-TACAGGCTTGCTCACTCGAATT-3'      |
| Interleukin-10 ( <i>IL-10</i> )                                   | Forward | 5'-CAGAGCCACATGCTCCTAGA-3'        |
|   | Reverse | 5'-TGTCCAGCTGGTCCTTTGTT-3'        |
| <i>TGF-<math>\beta</math>1</i> (Transforming growth factor beta1) | Forward | 5'-GCCACTGCCCATCGTCTACT-3'        |
|   | Reverse | 5'-CACTTGCAAGGAGCGCACAAAT-3'      |