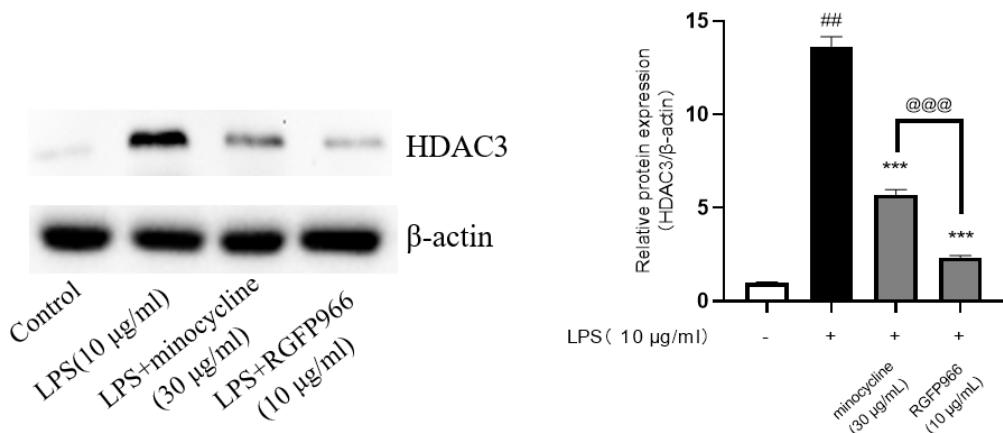


## 1. Positive Control

To further verify the inhibitory effect of minocycline on HDAC3, we employed the HDAC3-selective inhibitor RGFP966 as a positive control to perform a comparison experiment. The western blot results are shown in Figure S1. RGFP966 (10 µg/mL) can significantly inhibit the expression of HDAC3 in LPS-induced A549 cells ( $p=0.000$ ). The western blot results of minocycline have the same trend as those of RGFP966, which can significantly inhibit the expression of HDAC3 ( $p=0.000$ ). However, the HDAC3-selective inhibitors exhibited stronger inhibition of HDAC3 than minocycline ( $p=0.000$ ). According to the comparison results of positive drugs, minocycline has a definite inhibitory effect on HDAC3, and its drug potency is weaker than that of RGFP966.



**Appendix 1.** Western blot analysis of the inhibitory effects of minocycline and RGFP966 on HDAC3. (n=6) Values are mean  $\pm$  SD. ###P < 0.01 vs. the control group, \*\*\*p < 0.001 vs. LPS-treated group, @@@@ p < 0.001 vs. minocycline (30 µg/mL) group.

## 2. Statistical Details

The exact  $p$ -values of all the experimental result statistics are shown as follows. The statistical details presented in the subsequent tables (**Table S1-S4**) accurately correspond with each image depicted in the article.

## Appendix 2. The exact $p$ -values of all the results in Figure 1

### **Appendix 3. The exact $p$ -values of all the results in Figure 2**

#### **Appendix 4. The exact $p$ -values of all the results in Figure 3**

LPS(40µg/mL)	Minocycline (mg/kg)	p-values				
		apoptotic rate	Relative intensity of fluorescence	Bcl-2	Bax	Cleaved-caspase 3
-	0	-	-	-	-	-
+	0	0.000	0.000	0.000	0.000	0.000
+	10	0.000	0.000	0.027	0.000	0.000
+	30	0.000	0.000	0.000	0.000	0.000

## **Appendix 5.The exact $p$ -values of all the results in Figure 4**