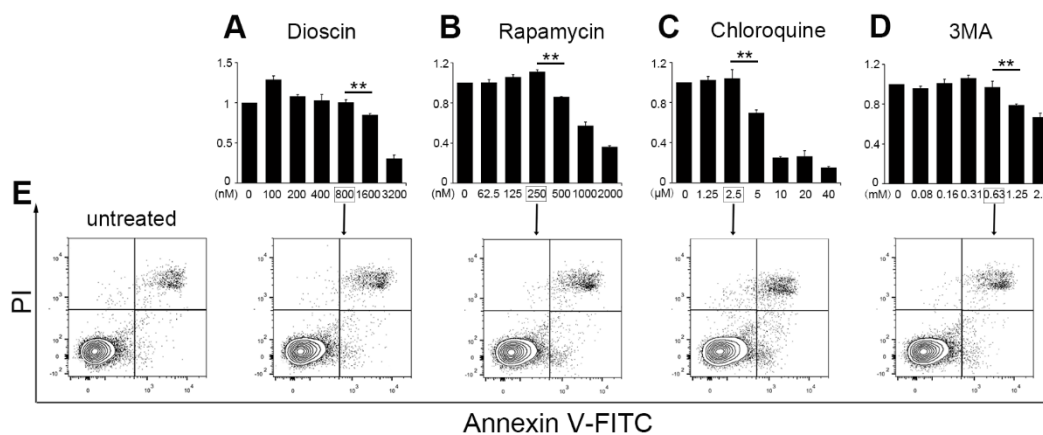
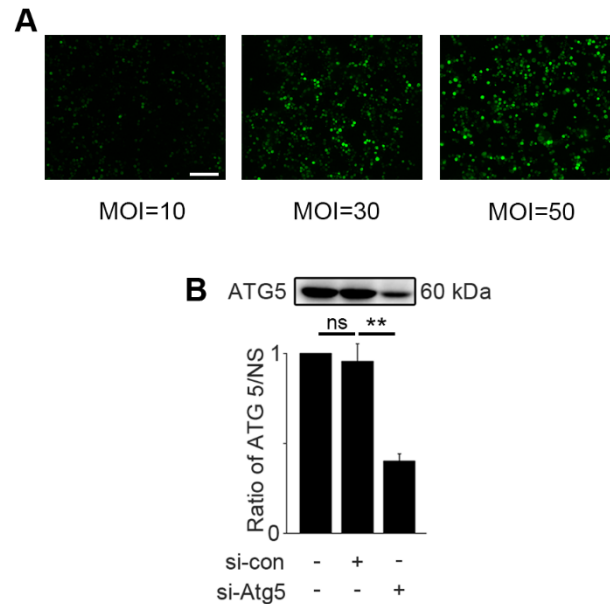


**Figure S1. Method of administration.** dioscin (80 mg/kg body weight) or vehicle (CMC-Na) was orally administered to C57BL/6 mice every day after saline or CS injection (n=10 per group).



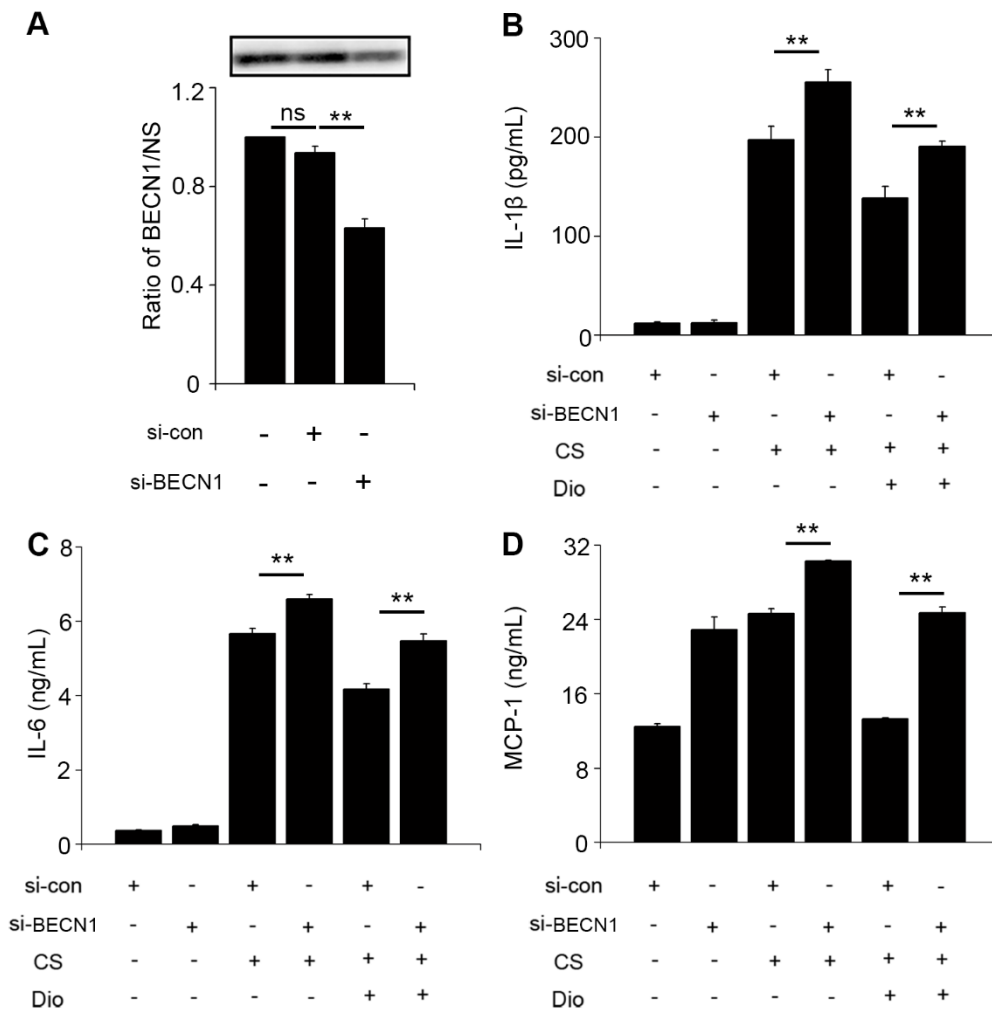
**Figure S2. Effects of chemicals on cell viability.** (A) MTT assay of dioscin-treated cells. (B) MTT assay of Rapamycin-treated cells. (C) MTT assay of Chloroquine-treated cells. (D) MTT assay of 3MA-treated cells. (E) Apoptosis assay of drugs used in experiment. \*,  $P < 0.05$ ; \*\*,  $P < 0.01$ . Error

bars indicate mean  $\pm$  SEM. All experiments were repeated three times.



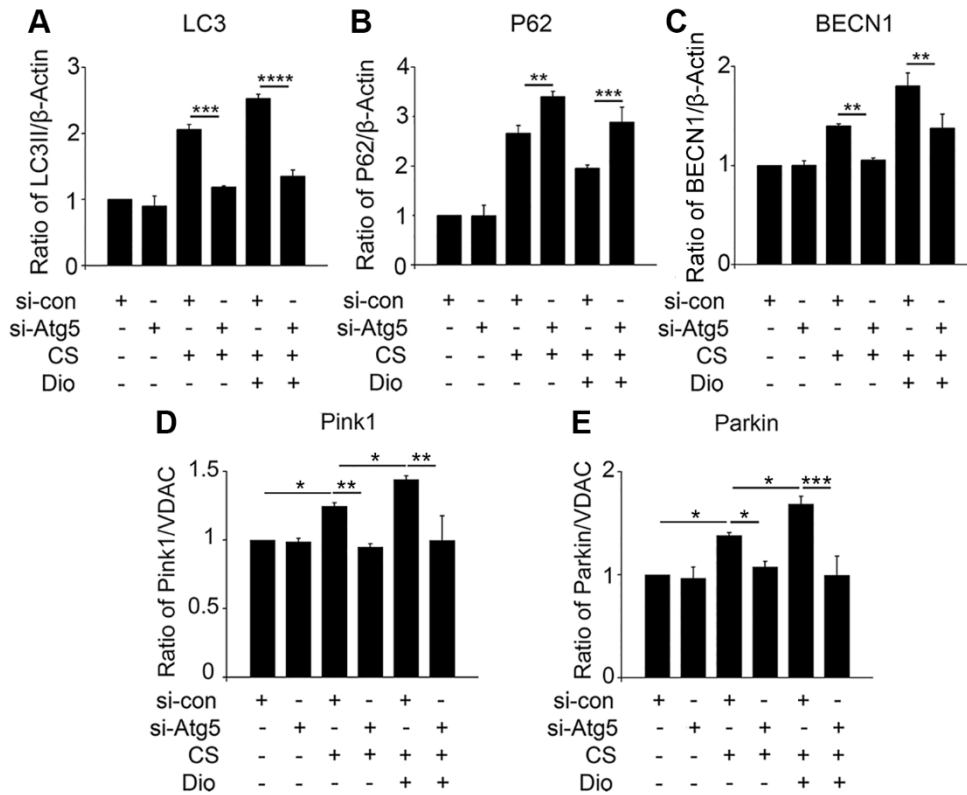
**Figure S3. Transfection efficiency of lentivirus with Atg5 siRNA sequence.**

(A) Multiplicity of Infection (MOI) gradient of mRFP-GFP-LC3 lentivirus infecting MH-S cells at 24 hours post infection, scale bar indicates 100  $\mu$ m. (B) Transfection efficiency was tested by measuring ATG5 by immunoblot assay (n=3). \*,  $P < 0.05$ ; \*\*,  $P < 0.01$ . Error bars indicate mean  $\pm$  SEM. All experiments were repeated three times.

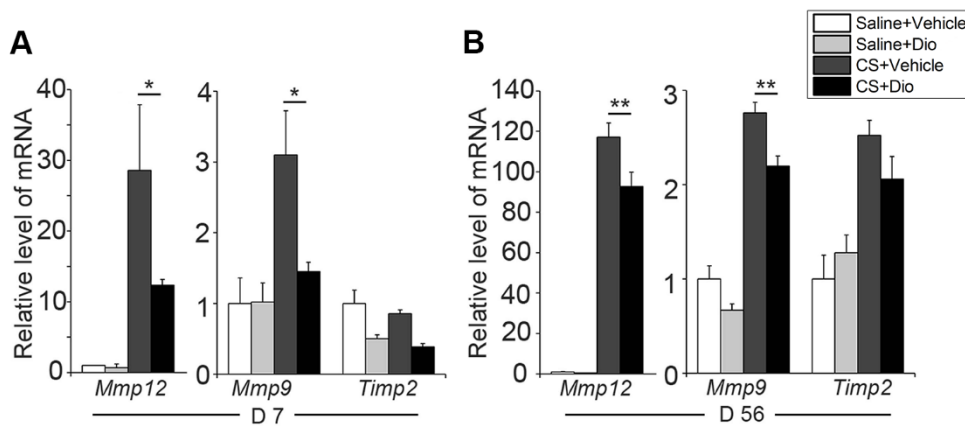


**Figure S4. BECN1 knockdown could affect function of dioscin.**

(A) Transfection efficiency was tested by measuring Becn1 by immunoblot assay (n=3) (B-D) ELISA analysis of IL-1 $\beta$ , IL-6, and MCP-1 in MH-S cell supernatant (n=3). (B-G). \*,  $P < 0.05$ ; \*\*,  $P < 0.01$ . Error bars indicate mean  $\pm$  SEM. All experiments were repeated three times with similar results.



**Figure S5. Quantification of autophagy and mitophagy protein. (A)** Quantification of LC3II. **(B)** Quantification of P62. **(C)** Quantification of BECN1. **(D)** Quantification of Pink1. **(E)** Quantification of Parkin. \*,  $P < 0.05$ ; \*\*,  $P < 0.01$ ; \*\*\*,  $P < 0.001$ , \*\*\*\*,  $P < 0.0001$ . Error bars indicate mean  $\pm$  SEM. All experiments were repeated three times.



**Figure S6. Dioscin treatment reduced the production of MMPs and TIMP**

**in gene level. (A)** mRNA level of *Mmp12*, *Mmp9*, and *Timp2* in lung tissue at day 7 (n=5). **(B)** mRNA level of *Mmp12*, *Mmp9*, and *Timp2* in lung tissue at day 56 (n=5). \*,  $P < 0.05$ ; \*\*,  $P < 0.01$ . Error bars indicate mean  $\pm$  SEM. All experiments were repeated three times.

**Table S1. Primer sequences for qPCR in this study.**

Gene	Forward (5'-3')	Reverse (5'-3')
<i>BECN1</i>	ATGGAGGGGTCTAAGGCGTC	TCCTCTCCTGAGTTAGCCTCT
<i>P62</i>	AGGATGGGGACTTGGTTGC	TCACAGATCACATTGGGGTGC
<i>Atg5</i>	TGTGCTTCGAGATGTGTGGTT	GTCAAATAGCTGACTCTTGGCAA
<i>Atg7</i>	GTTTCGCCCCCTTTAATAGTGC	TGAACTCCAACGTCAAGCGG
<i>ULK1</i>	TGGAGGTGGCCGTCAAATG	CGCATAGTGTGCAGGTAGTC
<i>Mmp9</i>	CTGGACAGCCAGACACTAAAG	CTCGCGGCAAGTCTTCAGAG
<i>Mmp12</i>	CTGCTCCCATGAATGACAGTG	AGTTGCTTCTAGCCCAAAGAAC
<i>Il-1b</i>	GCAACTGTTCTGAACTCAACT	ATCTTTTGGGGTCCGTCAACT
<i>Mcp-1</i>	TTAAAAACCTGGATCGGAACCAA	GCATTAGCTTCAGATTTACGGGT
<i>Il-6</i>	TAGTCCTTCTACCCCAATTTCC	TTGGTCCTTAGGCCACTCCCTTC
<i>Col1a1</i>	GCTCCTCTTAGGGGCCACT	CCACGTCTCACCATTGGGG
<i>Col3a1</i>	CTGTAACATGGAACTGGGGAAA	CCATAGCTGAACTGAAAACCACC