Supplementary figure 1: MI-2 treatment reduced the coltis-induced damage on colon tissues *in vovo*.



(a) Representative result of western blot analysis shows the expression level of ZO-1 protein from large intestine of healthy or DSS-induced colitis mice with/without treatment of MI-2 (30mg per Kg). Full-lenth blots are presented in Supplementary Figure 3. These results were from three independent experiments. The expression of β -actin was analyzed as a loading control. (b) The graph shows the expression levels of ZO-1 pooled from three independent experiments of large intestine from healthy or DSS-induced colitis mice with/without treatment of MI-2. (c) Representative result of western blot analysis shows the expression level of ZO-1 protein from Caco2 cell monolayers treated with DMSO, LPS (1 µg/mL), and LPS (1 µg/mL) + MI-2 (2 µM). Full-lenth blots are presented in Supplementary Figure 4. These results were from three independent experiments. The expression of β -actin was analyzed as a loading control. (d) The graph shows the expression levels of ZO-1 pooled from three independent experiments.

Supplementary figure 2 : MALT-1 Knockdown Downregulates the Production of Inflammatory Cytokines TNF α and IL-1 β .



(a) Representative western blots show the successful knockdown of MALT-1 on human macrophages by siRNA from three independent experiments at the protein expression level. Full-lenth blots are presented in Supplementary Figure 4. The mRNA levels of (b) MALT-1, (c) TNF α , and (d) IL-1 β were measured from untreated and LPS-treated macrophages with/without MALT-1 siRNA transfection using qPCR method. Pooled results are shown from three independent experiments. ** and *** reflect significant differences based on two-tailed unpaired Student's *t*-tests at *P* < 0.01 and *P* < 0.001, respectively. The error bars show the SEM values.

Supplementary figure 3: Immunoblot analysis of ZO-1 in human epithelial monolayer and mouse colon tissues.



(a) Murine colon tissues were used to assess the expression level of ZO-1 using immunoblot method. Colon tissues of healthy or DSS-induced colitis mice with/without treatment of MI-2 (30mg per Kg) were used for this experiment. (b) Human colon cell line, Caco-2 was used to assesse the expression of ZO-1 using immunoblot method with a use of β –actin as a loading control. Human Caco-2 cells were incubated treated with DMSO, LPS (1 µg/mL), and LPS (1 µg/mL) + MI-2 (2 µM). The level of β -actin was analysed as a loading control.

Supplementary figure 4: Immunoblot analysis of MALT-1 in mouse intestinal tissues and human macrophages.



(a) Mouse intestinal tissues of control, DSS and DSS + MI-2 mice group were used to assess the expression of MALT1 using immunoblot method with a use of β –actin as a loading control. N/S indicates non-specific band. (b) Human macrophages derived from peripheral blood monocytes in the absence or presence of LPS with or without knockdown of MALT-1 siRNA were used to assess the expression of MALT-1 using immunoblot method with a use of β –actin as a loading control.

Supplementary Table. 1 Disease activity index based on previously study

| score | Weight loss | Stool consistency | Bleeding stool |
|-------|--------------------------|------------------------|---------------------|
| 0 | No weight loss or weight | Normal and well formed | Normal color stool |
| | gain | | |
| 1 | 5-10 % weight loss | - | - |
| 2 | 11-15 % weight loss | Very soft and unformed | Reddish color stool |
| 3 | 16-20 % weight loss | - | - |
| 4 | >21 % weight loss | Watery stool | Bloody stool |