Supplementary Table 1.

Primer name	Sequence	Description
CD98-F	5'-GAGGACAGGCTTTTGATTGC-3'	CD98 gene RT-PCR forward primer
CD98-R	5'-ATTCAGTACGCTCCCCAGTG-3'	CD98 gene RT-PCR reverse primer
ΤΝFα-F	5'-AGGCTGCCCCGACTACGT-3'	Tumor necrosis factor gene RT-PCR forward primer
TNFα-R	5'-GACTTTCTCCTGGTATGAGATAGCAAA-3'	Tumor necrosis factor gene RT-PCR reverse primer
IL6-F	5'-ACAAGTCGGAGGCTTAATTACACAT-3'	Interleukin 6 gene RT-PCR forward primer
IL6-R	5'-TTGCCATTGCACAACTCTTTTC-3'	Interleukin 6 gene RT-PCR reverse primer
IL12-F	5'-GCCAGTACACCTGCCACAAA-3'	Interleukin 12 gene RT-PCR forward primer
IL12-R	5'-TGTGGAGCAGCAGATGTGAGT-3'	Interleukin 12 gene RT-PCR reverse primer
36B4-F	5'-TCCAGGCTTTGGGCATCA-3'	36B4 gene RT-PCR forward primer
36B4-R	5'-CTTTATCAGCTGCACATCACTCAGA-3'	36B4 gene RT-PCR reverse primer

CCP CP

Supplementary Figure 1.

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Supplementary Figure 2.







Supplementary Figure 4.

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Supplementary Figure 5.

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T (s)

CHR HANN



ALA ALA



Supplementary Figure 9.

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CER AN



Supplementary Figure 11.

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Supplementary Figure 12.

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Supplementary Figure 13.

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1 Nanoparticles with Surface Antibody Against CD98 and Carrying CD98 Small Interfering

- 2 RNA Reduce Colitis in Mice
- **3** Supplementary Materials and Methods

4 Preparation and Characterization of scCD98-PEG-UAC

The synthetic strategy of scCD98-PEG-UAC is outlined in Supplementary Figure 1. Chitosan 5 (Sigma) was obtained by further deacetylating and depolymerizing commercial chitosan using 6 alkali treatment and sodium nitrite, respectively. Degree of deacetylation of chitosan was 7 estimated as 95.6% by ¹H NMR. Molecular weight of the resultant chitosan was measured as 8 11.4 kDa by MOLDI-TOF. The UAC sample was synthesized described in previous report [1]. 9 10 UAC-PEG-MAL was prepared in phosphate-buffer solution (PBS, pH = 7.2) through specific reaction between UAC and the NHS groups of bifunctional PEG derivative (NHS-PEG-MAL, 11 MW 2000, Jenkem, Beijing, China) at the mole ratio of 1:4 overnight while stirring. The 12 resulting conjugate was purified by ultrafiltration with an Amicon[®] Ultra (regenerated cellulose 13 membrane, MWCO = 10 kDa, Millipore). Meanwhile, CD98 antibody (Biolegend) was reduced 14 to single-chain CD98 antibody (scCD98) using 2-mercapto ethylamine (Sigma). Then MAL-15 PEG-UAC was reacted with scCD98 at mole ratio of 1:2 in PBS (pH = 7.2) for 24 h at room 16 temperature. The terminal MAL groups of MAL-PEG-UAC were specifically reacted with the 17 thiol groups of scCD98, resulting in scCD98-PEG-UAC. Unreacted maleimide groups were 18 quenched with 5 eq. cysteine. The final conjugates were lyophilized and characterized by FT-IR 19 and ¹H NMR. SDS-PAGE was used to demonstrate the full reaction of thiolated scCD98. 20

Infrared spectra of chitosan and its derivatives were recorded with a Varian 600-UMA FT-IR
microscope equipped with a liquid N₂ cooled HgCdTe detector and coupled to a Varian 7000 FT IR spectrometer. NMR spectrum was recorded on a Bruker Avance spectrometer (INOVA-600

