

Supplementary Information for

Tauroursodeoxycholic acid (TUDCA) counters osteoarthritis by regulating intracellular cholesterol levels and membrane fluidity of degenerated chondrocytes

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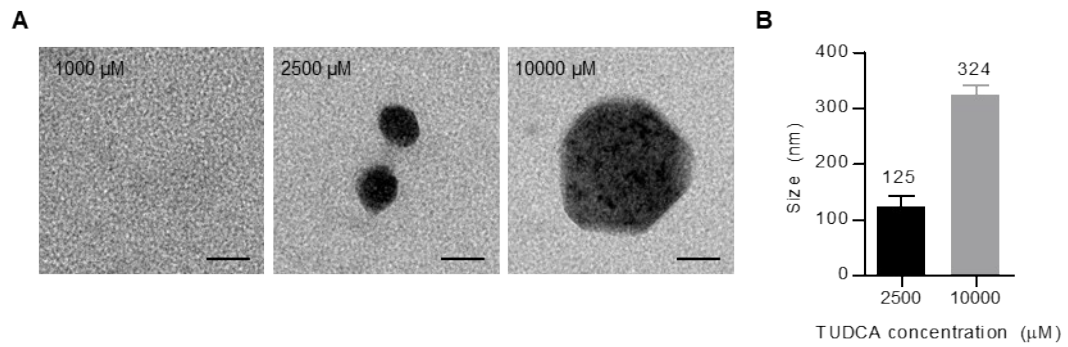
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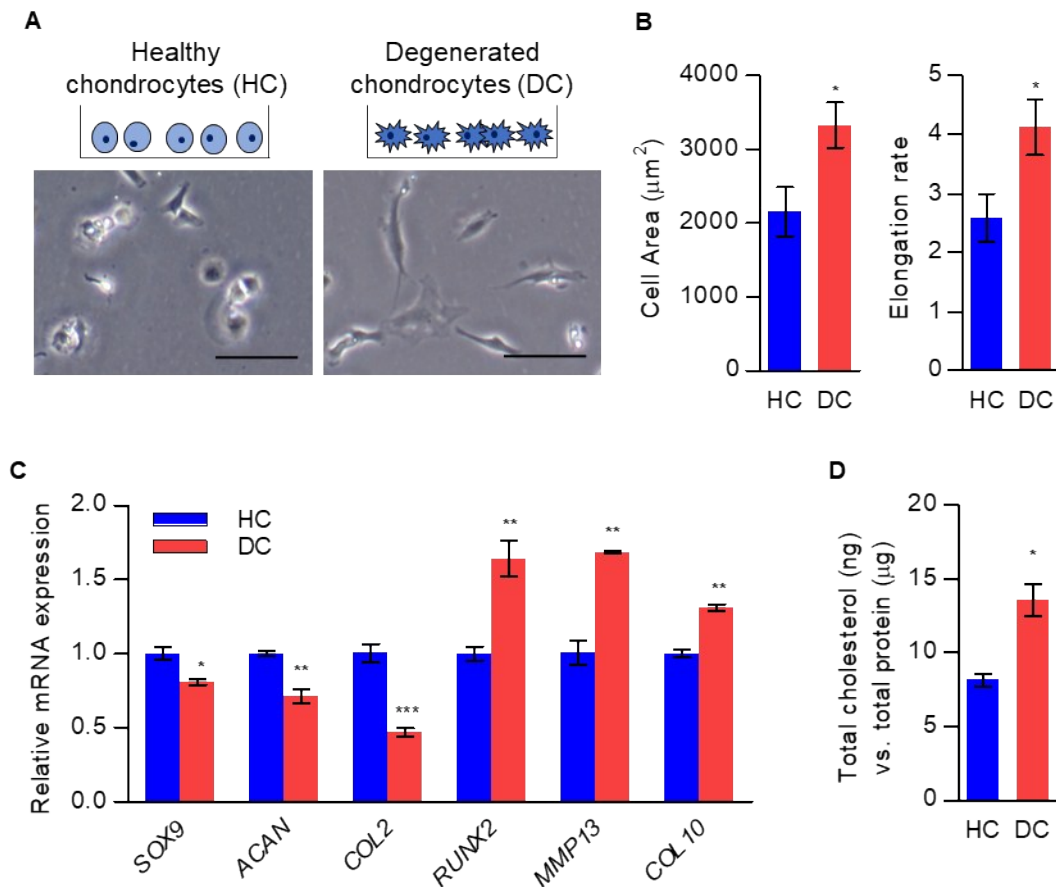
Keywords: Tauroursodeoxycholic acid (TUDCA), Cholesterol, Osteoarthritis, Membrane fluidity, Chondrocyte, Bile acid

Supplementary Figure 1



Supplementary Figure 1. Micelle formation of TUDCA in DPBS. (A) TEM images of TUDCA micelles in DPBS at 1000, 2500, and 10000 μM . Scale bar, 100 nm. (B) Average diameter of a TUDCA micelle was measured using ImageJ software. Data are mean \pm s.e.m. (n=4)

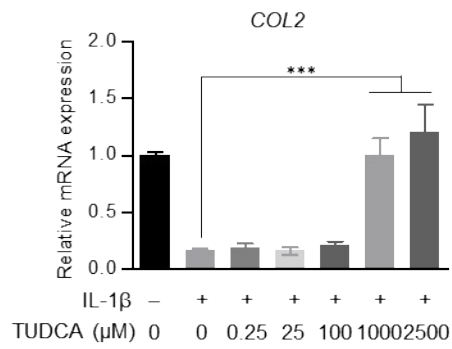
Supplementary Figure 2



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plementary Figure 2. Isolation of healthy or degenerated chondrocytes from patients with OA. (A) Isolation of healthy (HCs) and degenerated chondrocytes (DCs) from human knee cartilage. Cell morphology of HCs and DCs at passage 2. Scale bar, 100 μm . **(B)** Cell area and elongation rate of HCs and DCs were measured and quantified at passage 2 using ImageJ software. Data are mean \pm s.e.m. ($n=15$, Student's t-test, $*P<0.05$). **(C)** mRNA expression of chondrogenic marker genes (SOX9, COL2, and ACAN) and hypertrophic marker genes (COL10, MMP13, and RUNX2) in HCs and DCs were assessed by qPCR at 7 days of treatment. Data are mean \pm s.e.m. ($n=3$, Student's t-test, $*P<0.05$, $**P<0.01$). **(D)** Total cholesterol of HCs and DCs at passage 2 was normalized to the total protein content of the cells. Data are mean \pm s.e.m. ($n=3$, Student's t-test, $*P<0.05$).

Supplementary Figure 3



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ntary Figure 3. The effect of TUDCA on hypertrophic-induced chondrocytes using IL-1 β . mRNA expression of *COL2* in hypertrophy induced HCs following IL-1 β treatment was assessed by qPCR at 7 days after treatment with or without TUDCA. Data are mean \pm s.e.m. (n=4, One-way ANOVA, *** P <0.001).