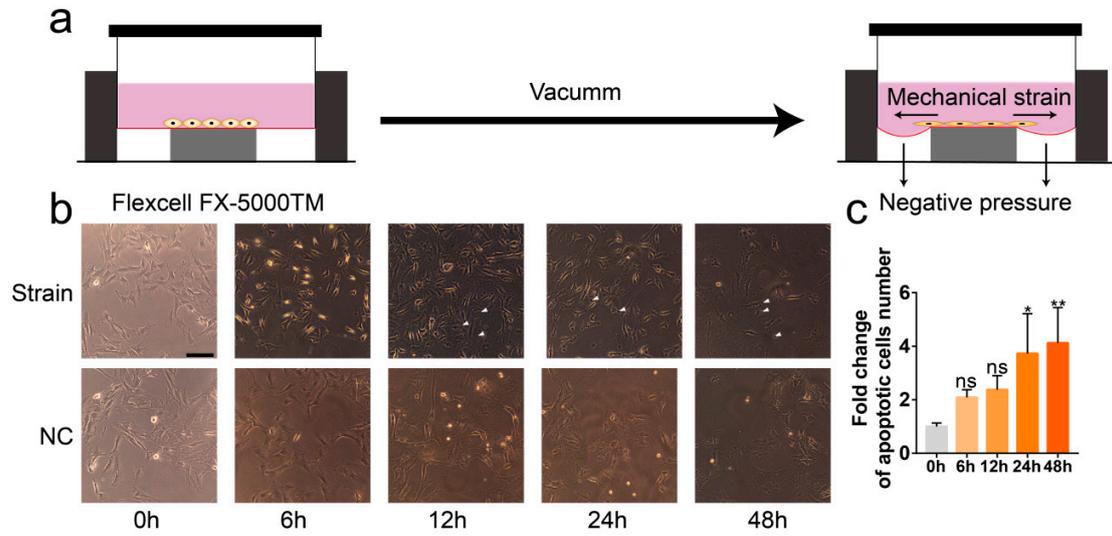
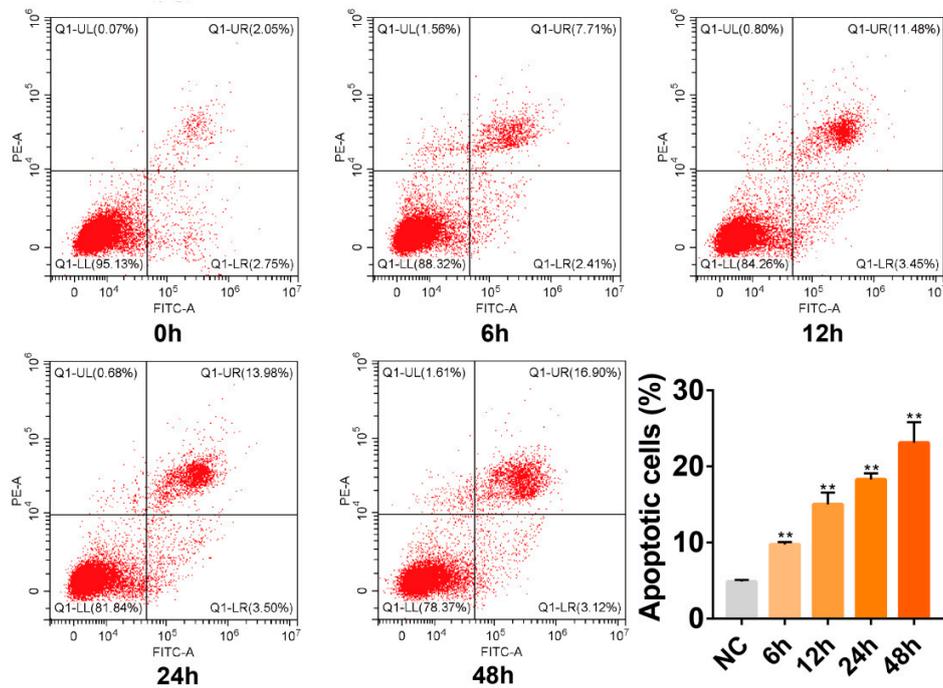


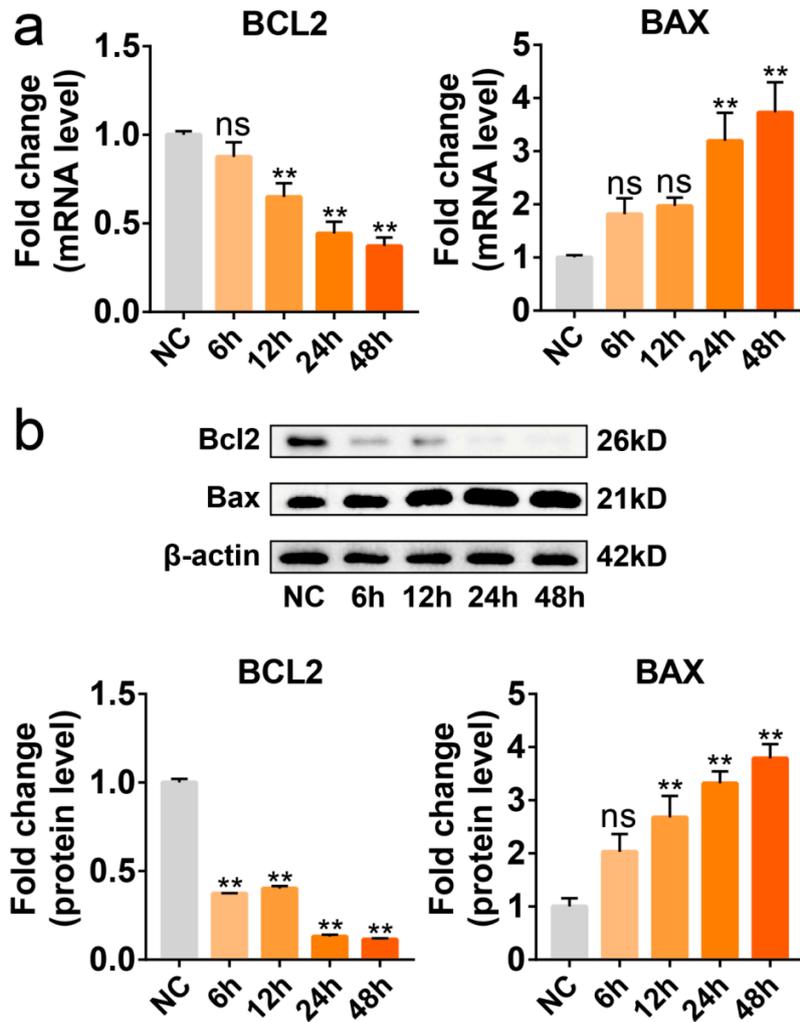
## Supplementary Materials



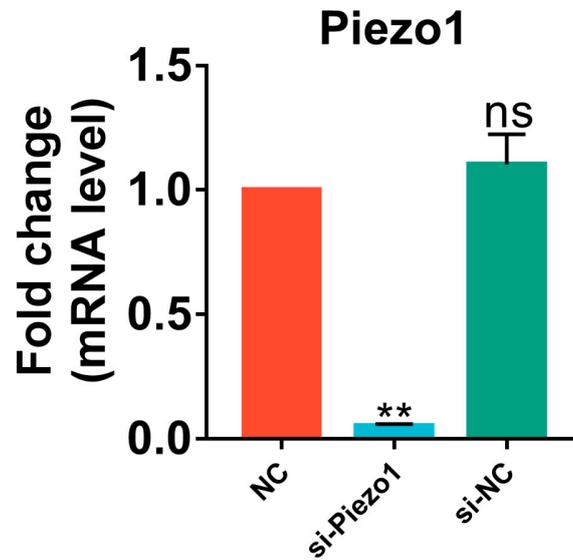
**Figure S1.** Morphological changes of chondrocytes under mechanical strain. **(a)** A schematic diagram of the mechanical strain treatment on chondrocytes. **(b)** The morphological characteristics of chondrocytes undergoing mechanical strain. **(c)** Quantified analysis of apoptotic chondrocytes ( $n=3$ ). \*,  $p < 0.05$  vs NC; \*\*,  $p < 0.01$  vs NC; ns, no significant differences. one-way ANOVA with Bonferroni's test for **(c)**. Scale bar: 100µm.



**Figure S2.** Flow cytometry analysis of chondrocyte stained with Annexin V-FITC and PI ( $n=3$ ). \*\*,  $p < 0.01$  vs NC. one-way ANOVA with Bonferroni's test was performed the statistical analysis.



**Figure S3.** Mechanical strain led to apoptosis and anabolic/catabolic imbalance in chondrocytes in a time-dependent manner. **(a)** RT-qPCR analysis of BAX and BCL2 in chondrocytes exposed to mechanical strain ( $n=3$ ). **(b)** Western blots analysis of BAX and BCL2 in chondrocytes exposed to mechanical strain ( $n=3$ ). \*\*,  $p < 0.01$  vs NC; ns, no significant differences. one-way ANOVA with Bonferroni's test for **(a, b)**.



**Figure S4.** The efficiency of si-Piezo1 investigated by qRT-PCR ( $n=3$ ). \*\*,  $p < 0.01$  vs NC; ns, no significant differences. one-way ANOVA with Bonferroni's test for the statistical analysis.

Table S1. Clinical information of human samples

I D	Age	Gender	Surgical site	ICRS
1	70	F	Right knee and Left knee	Intact: I Damaged: IV
2	68	F	Right knee	Intact: I Damaged: IV
3	82	F	Left knee	Intact: I Damaged: IV
4	73	F	Left knee	Intact: I Damaged: IV
5	59	R	Right knee	Intact: I Damaged: IV

Table S2. Primer sequences used in this study

Gene name	F/R	Sequences 5'-3'
$\beta$ -actin	F	TGCTATGTTGCCCTAGACTTCG
	R	GTTGGCATAGAGGTCTTTACGG
Piezo1	F	AGCAAGCAGGCACAAAGGC
	R	CGCACAAACTTGCCAACGAC

Bax	F	TTTTTGCTACAGGGTTTC
	R	TTGTTGTCCAGTTCATCG
Bcl2	F	GAGAGCGTCAACAGGGAG
	R	GCCAGGAGAAATCAAACA
Mmp3	F	CATGAACTTGGCCACTCCCT
	R	TGGGTACCACGAGGACATCA
Mmp13	F	GCCACCTTCTTCTTGTTGAGTTG
	R	GACTTCTTCAGGATCCCCGA
Aggreca	F	AGTGACCCATCTGCTTACCCTG
	R	CTGCATCTATGTCGGAGGTAGTG
Col2a1	F	GTGTCAAGGGTCACAGAGGTTAC
	R	CGCTCTCACCCCTCACACCT

---