

# Piezo1 integration of vascular architecture with physiological force

# Piezo1's role in endothelial cell regulation

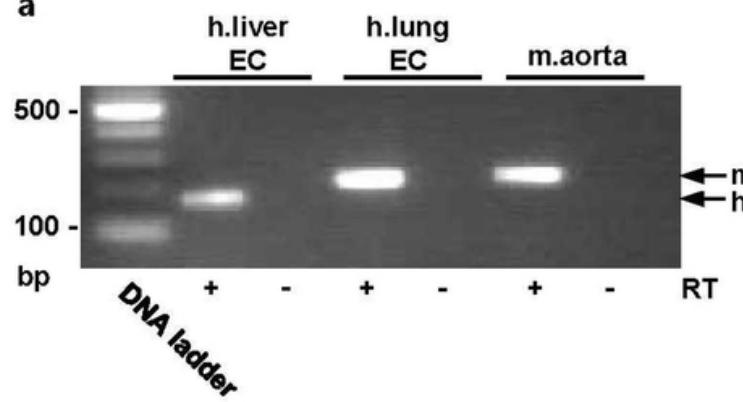
Piezo proteins as subunits of  $\text{Ca}^{2+}$  permeable cationic channels

Piezo1 channels as sensors of shear stress

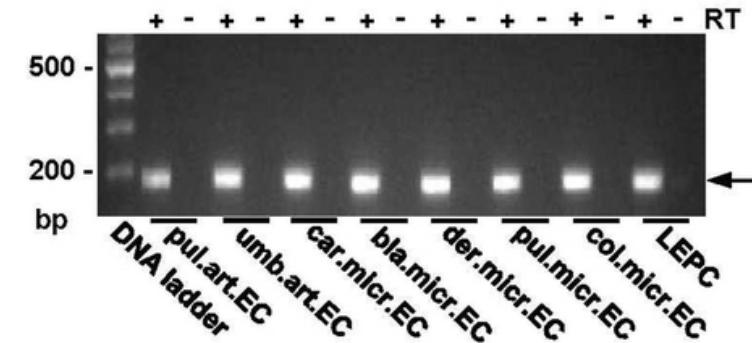
Piezo1 channels determinants of vascular structure

# Piezo1 mRNA in aorta and endothelial cells

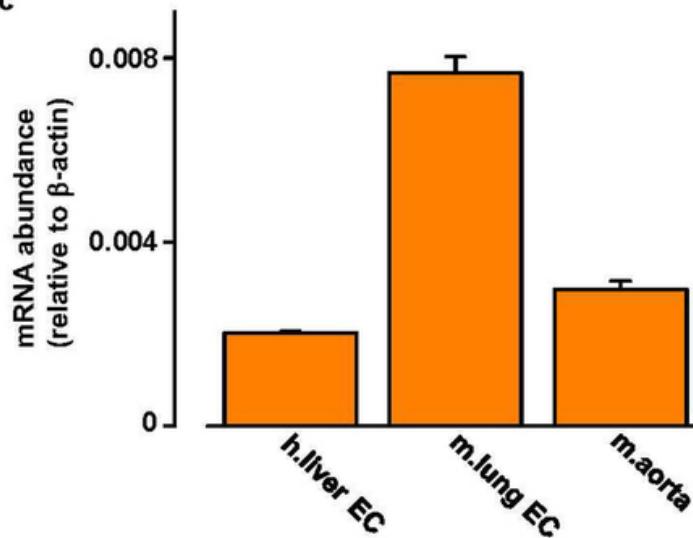
Ext. Fig. 1 a



b

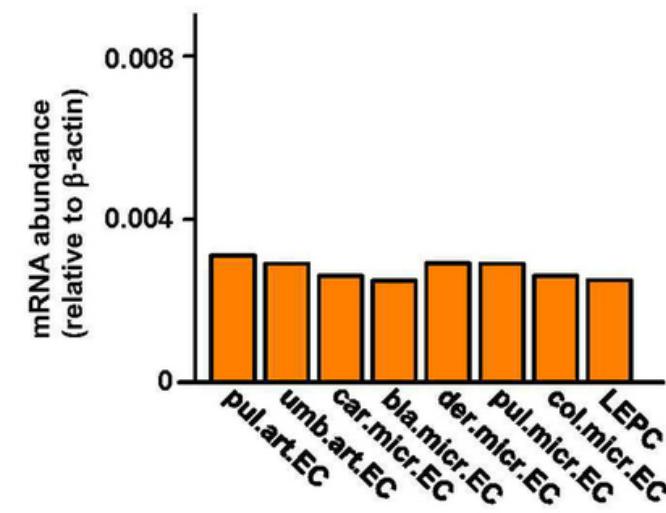


c



$n = 2$  each in duplicate

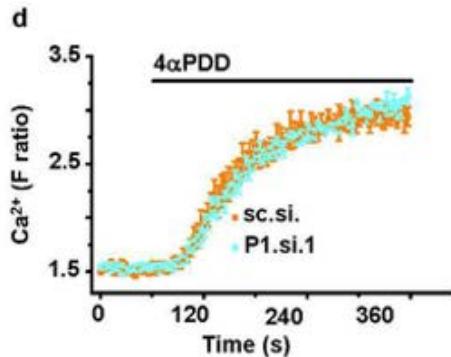
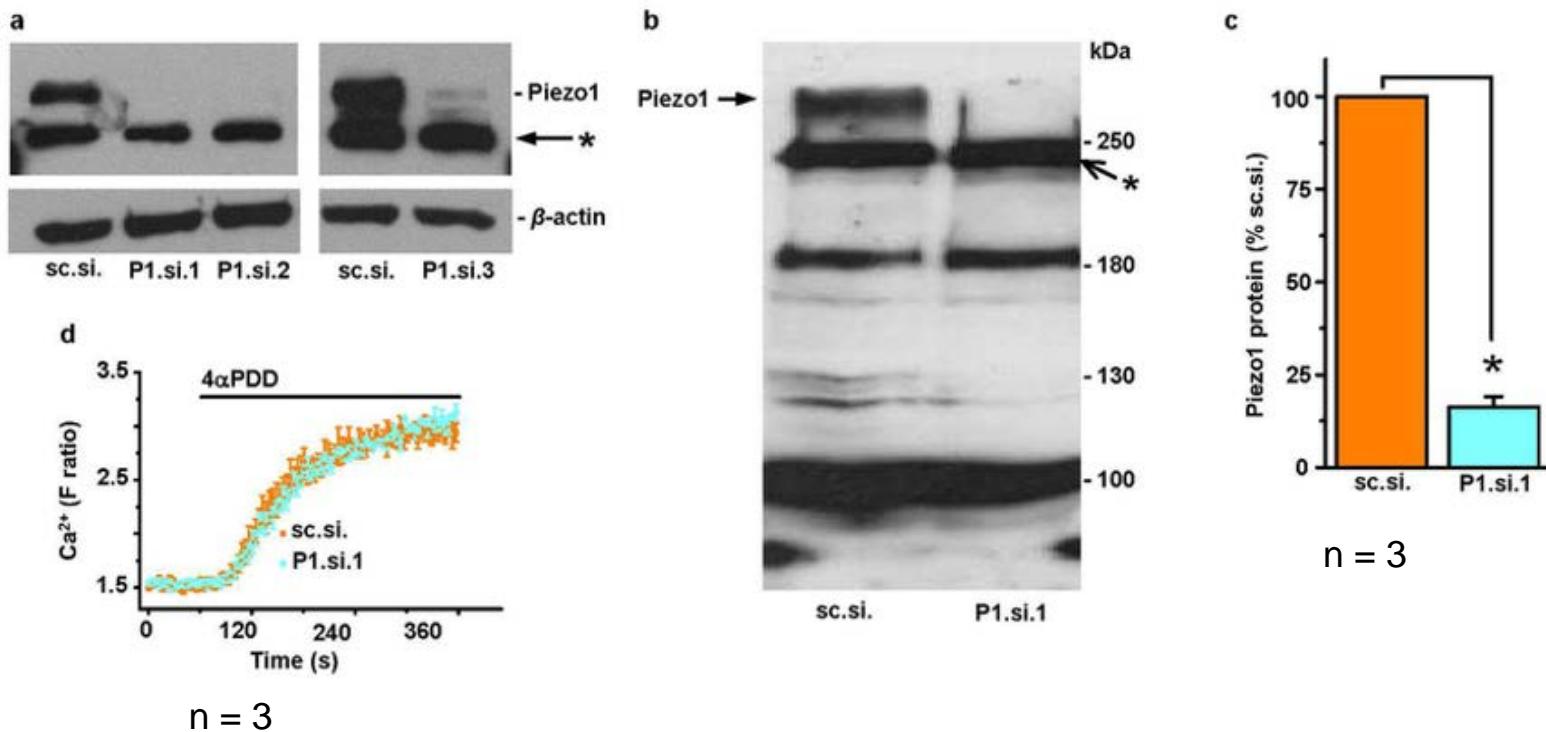
d



$n = 1$  each in duplicate

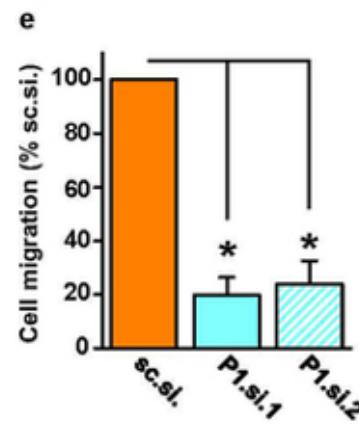
# HUVEC: cultured human umbilical vein endothelial cells

Ext. Fig. 2

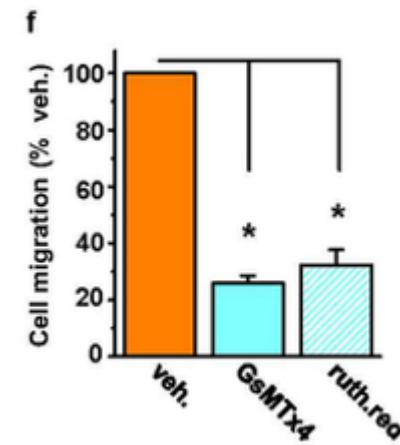


$n = 3$

Ext. Fig. 2

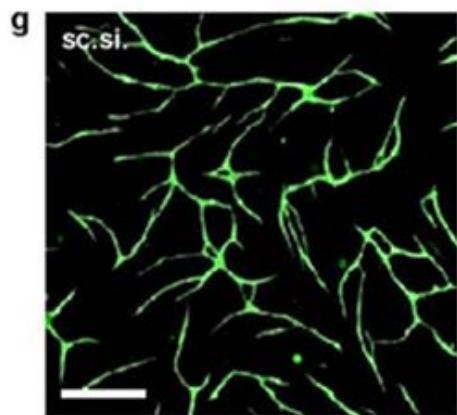


n = 4

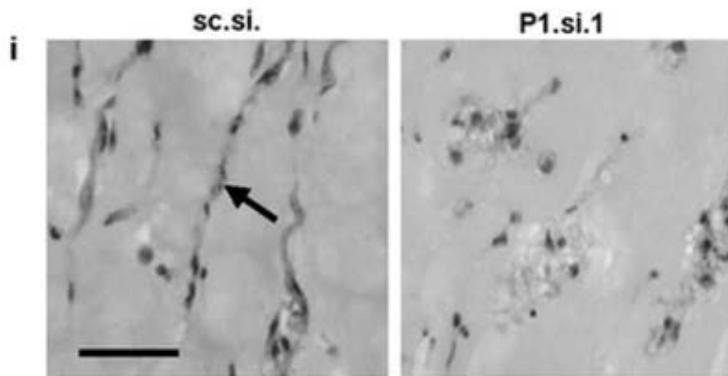
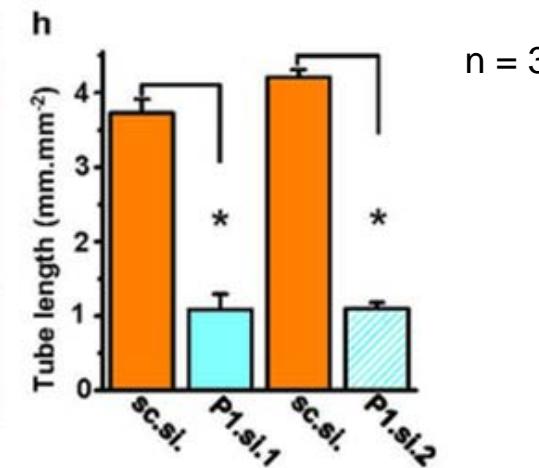
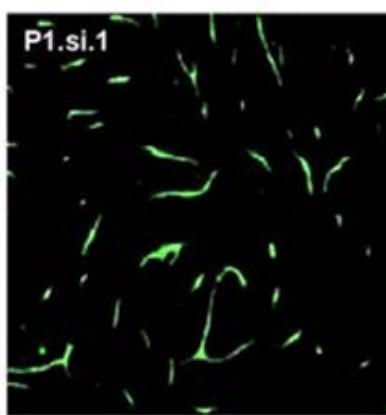


n = 3

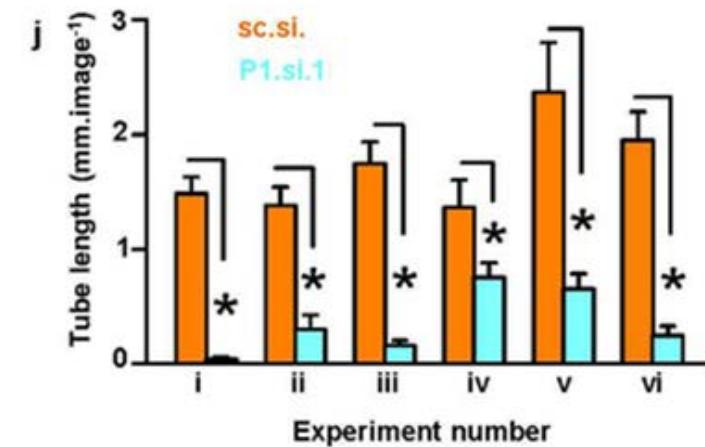
Ext. Fig. 2



Scale bar, 400  $\mu\text{m}$ .

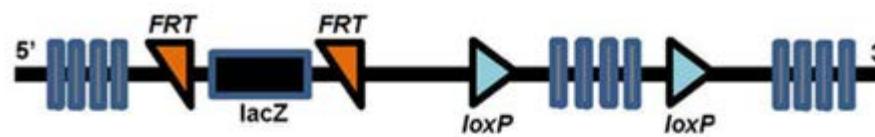


Scale bar, 50  $\mu\text{m}$ .

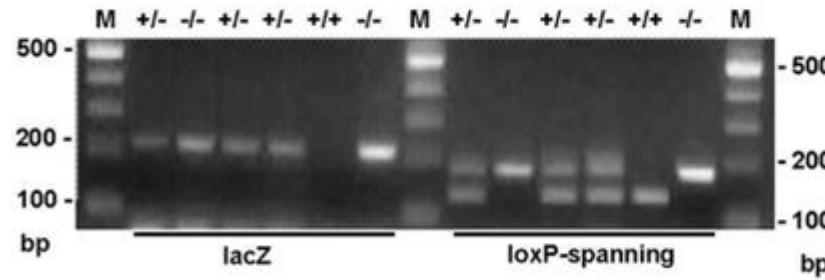


## Global and endothelial-specific Piezo1 modification

Ext. Fig. 3 a



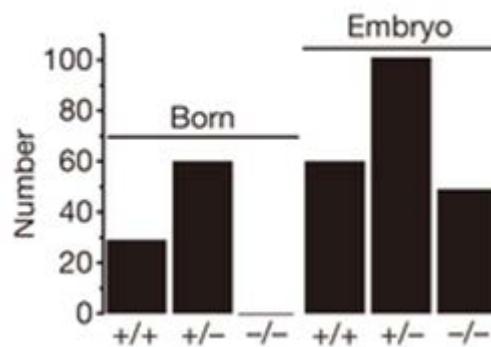
b



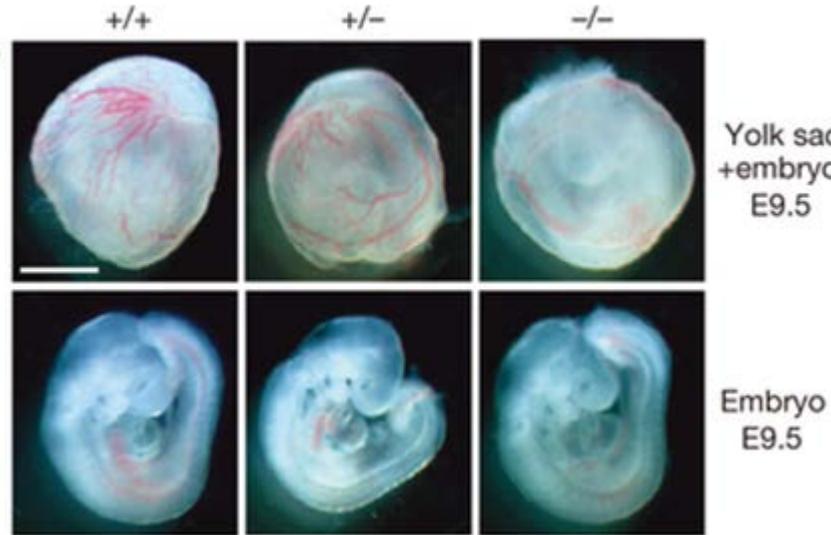
# Piezo1 function in murine embryos

**a**

Fig. 1



**b**



Scale bar, 1 mm.

Ext. Fig. 3

**c**

E10.5 +/+



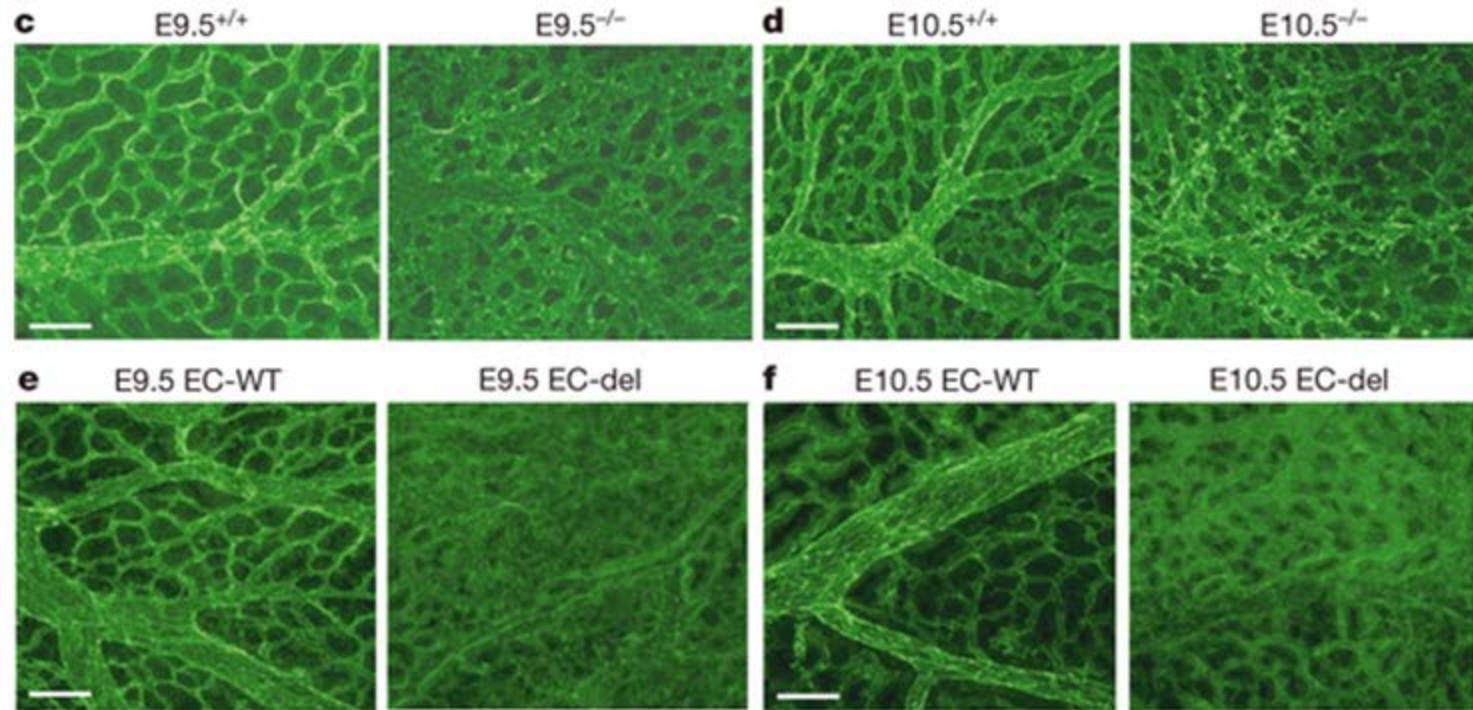
E10.5 -/-



Scale bar, 1 mm.

## Images of dissected *Piezo1<sup>+/+</sup>* and *Piezo1<sup>-/-</sup>* yolk sacs stained for CD31

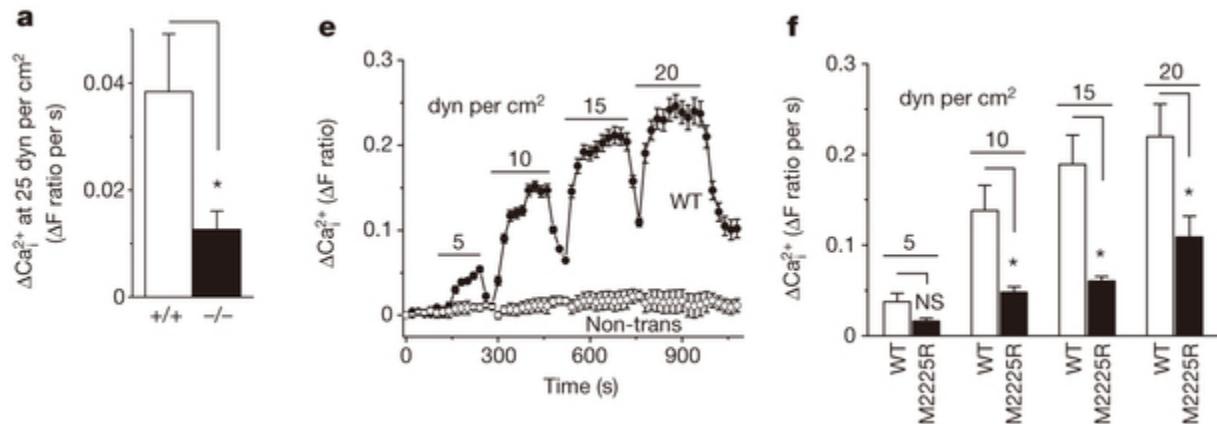
Fig. 1



Scale bars, 100 µm.

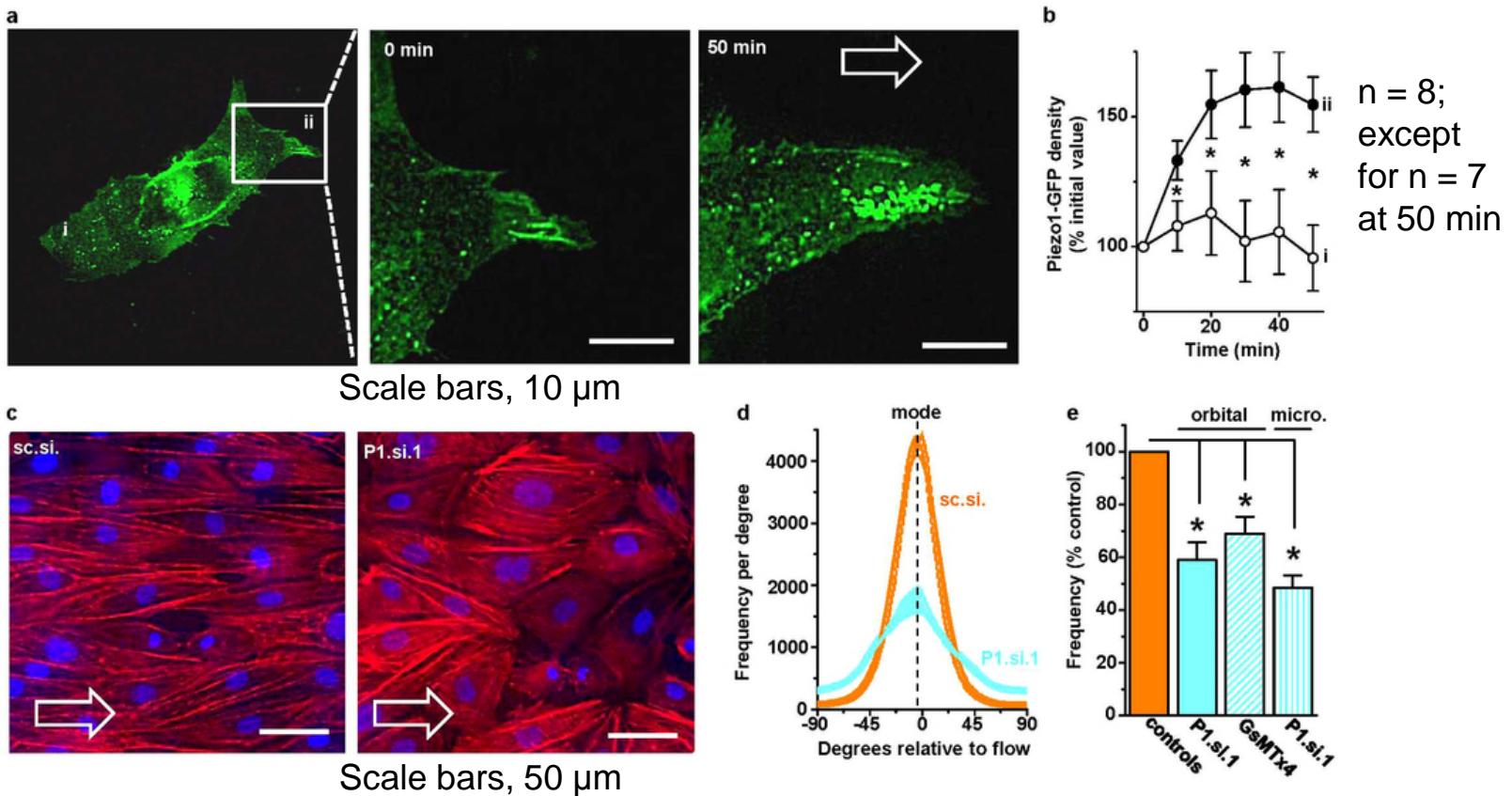
# Piezo1 in shear stress sensing

Fig. 2



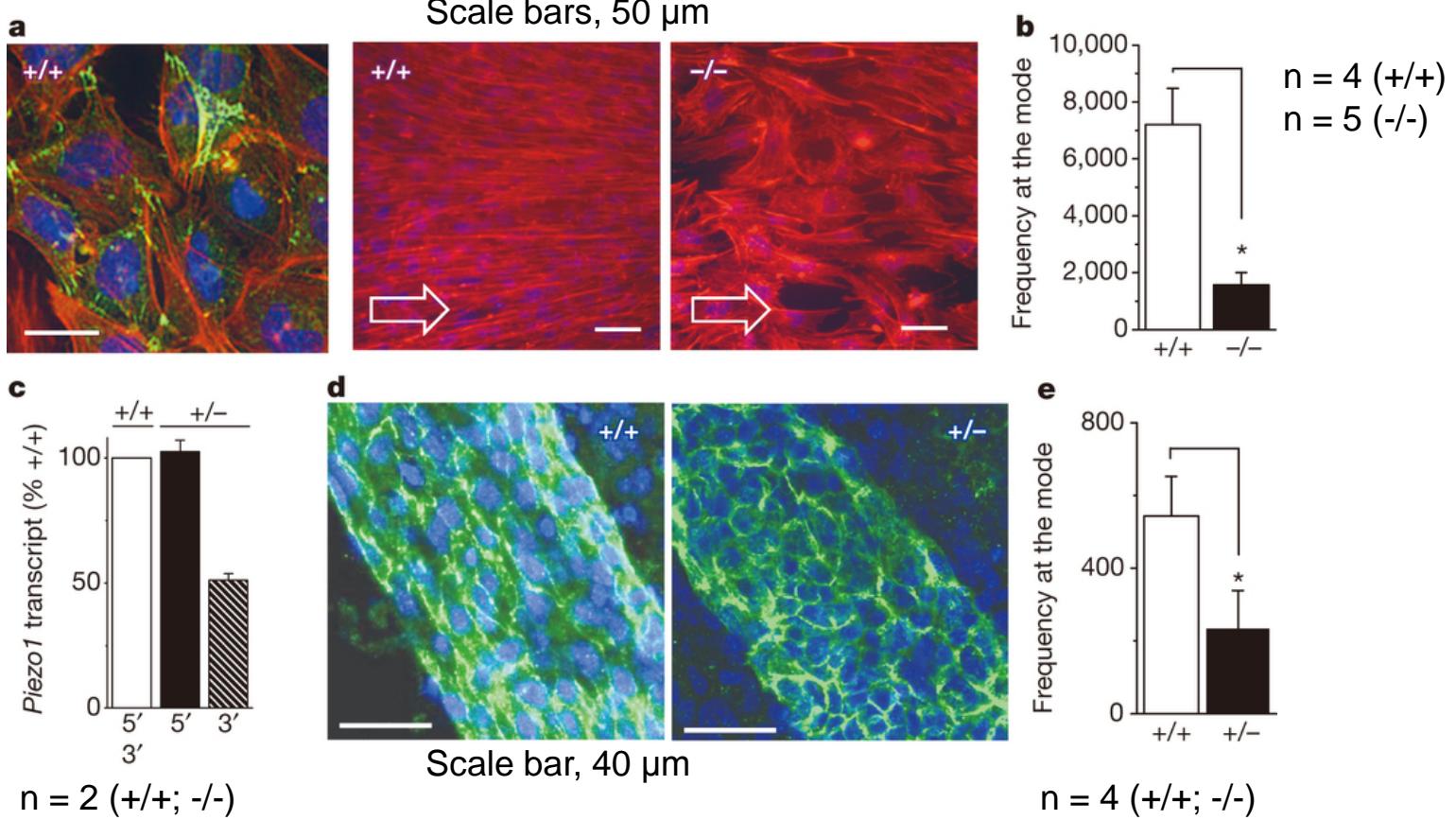
# Shear-stress-evoked redistribution of Piezo1 and the role of Piezo1 in alignment of endothelial cells to the direction of shear stress

Ext. Fig. 6



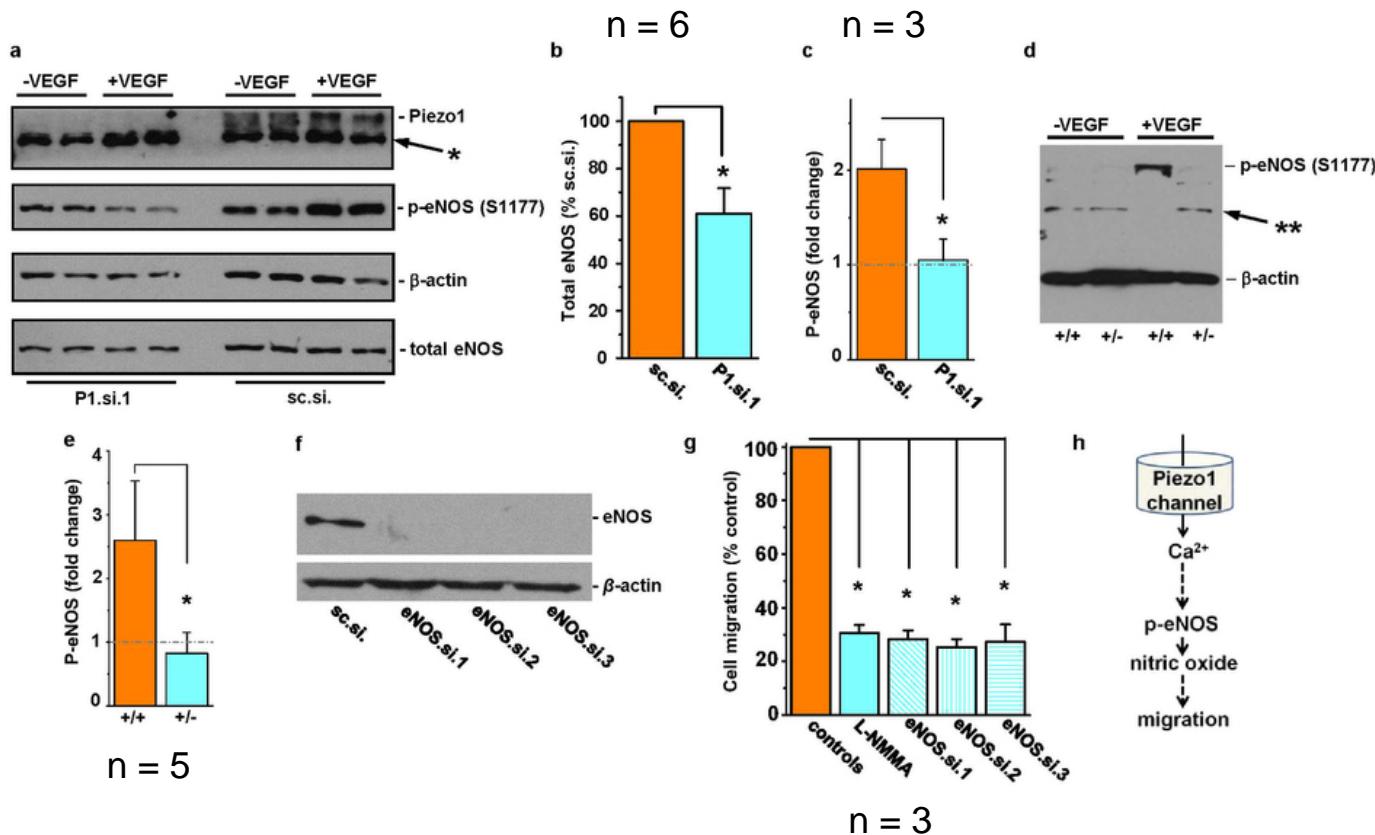
# The role of Piezo1 in endothelial cell alignment

Fig. 3



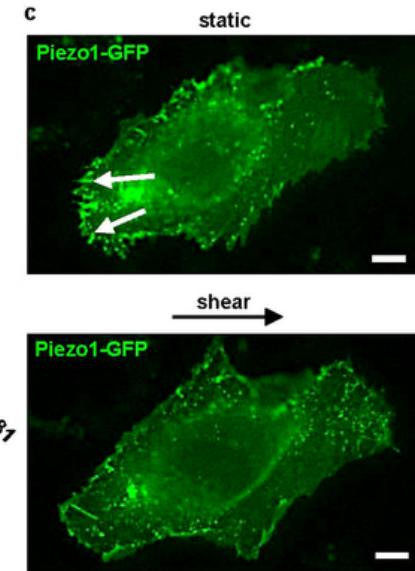
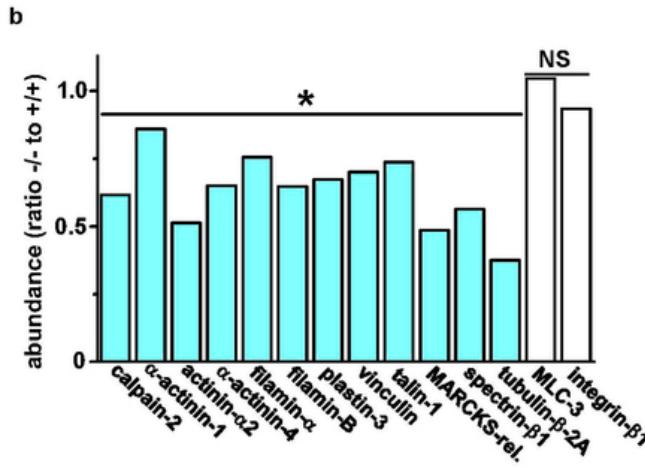
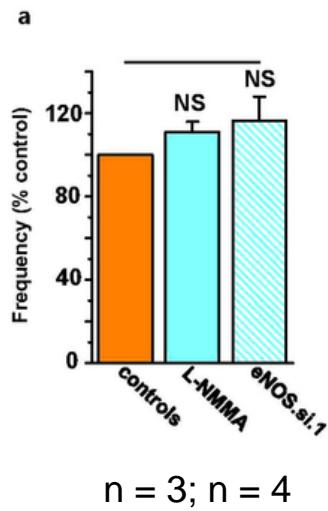
# Coupling to endothelial nitric oxide synthase

Ext. Fig. 7



# Endothelial cell alignment to shear stress lacks dependency on nitric oxide but is coupled to calpain

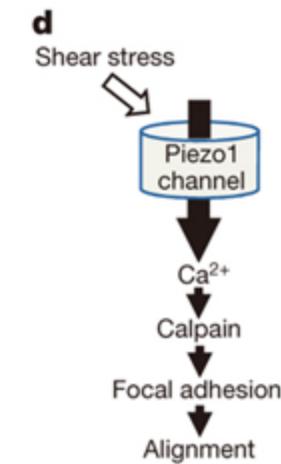
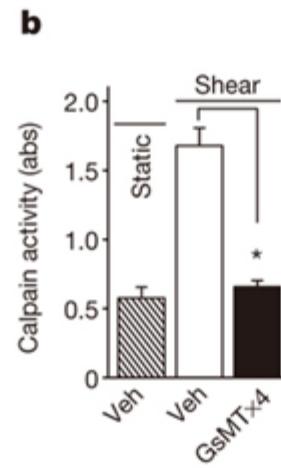
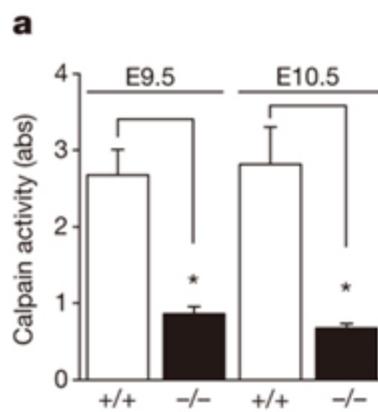
Ext. Fig. 8



Scale bar, 10  $\mu$ m  
n = 4

## Piezo1 coupling to calpain

Fig. 4



Thank you!