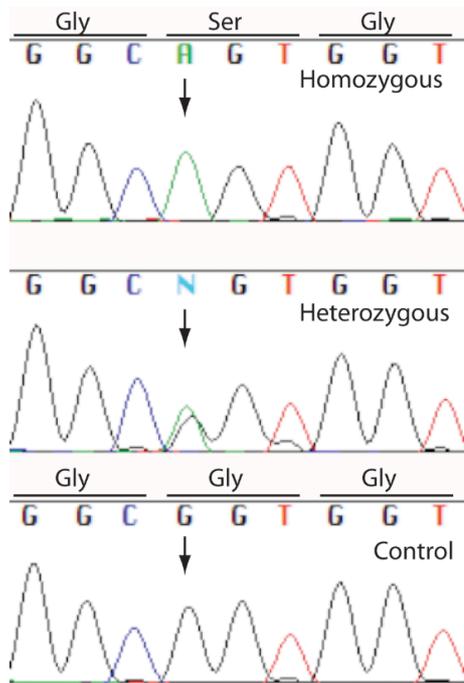
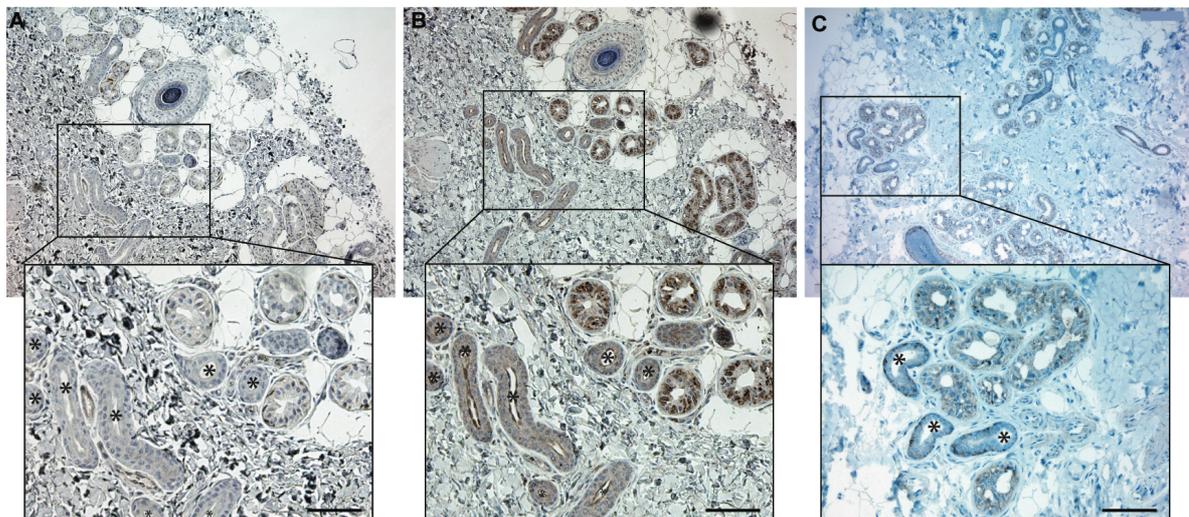


### Supplementary figures and legends

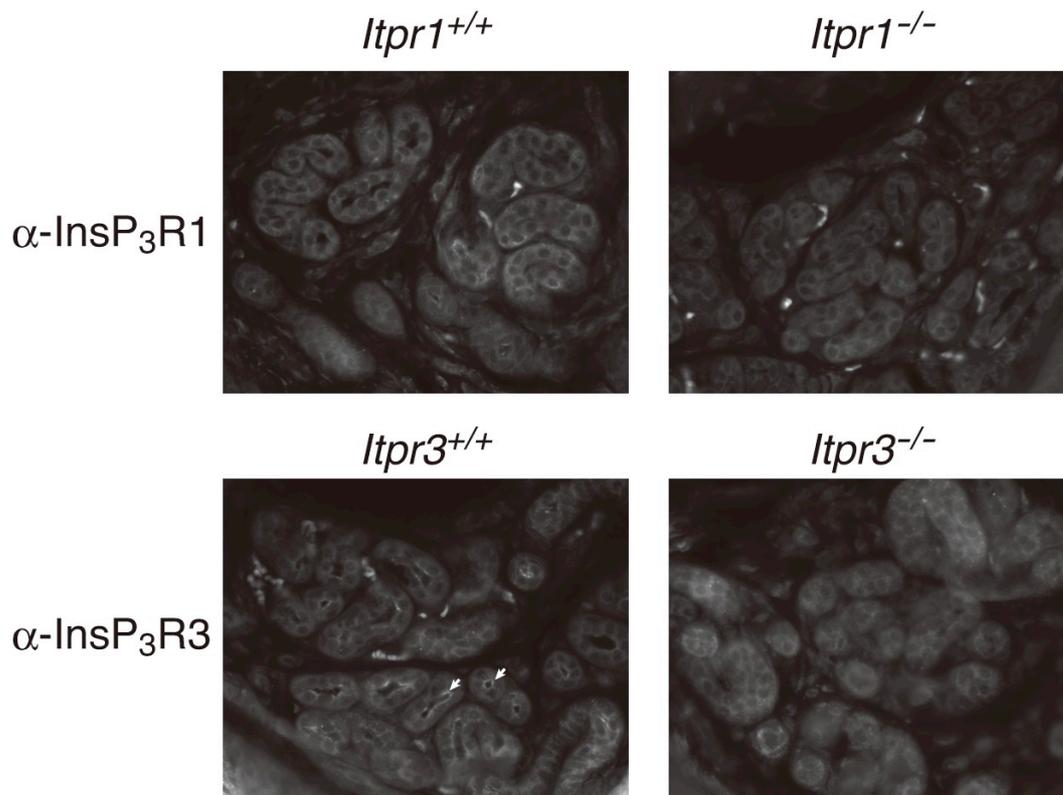


**Supplementary Figure 1.** Chromatograms illustrating the *ITPR2* gene mutation c.7492G>A (arrows) in an affected homozygous individual (top) and a heterozygous parent (middle). The w.t. sequence of a normal control individual is shown at the bottom. The corresponding codons are indicated above each chromatogram.



**Supplementary Figure 2.** Immunoreactivity for InsP<sub>3</sub>R1, InsP<sub>3</sub>R2 and InsP<sub>3</sub>R3 in normal human eccrine sweat glands. (A) InsP<sub>3</sub>R1 is not detectable in the sweat glands (box enlarged). (B) InsP<sub>3</sub>R2 is predominantly expressed in the secretory portion (box

enlarged) with some staining in the excretory ducts (\*) showing a concentration in subcellular regions lining the ducts. (C)  $\text{InsP}_3\text{R3}$  show strong staining of the basal ductal (peripheral) cell layer of the excretory duct (\*), and a weaker staining in cells of the secretory part (box enlarged). Stain: Polyclonal Rabbit anti-human  $\text{InsP}_3\text{R1}$  1:1000 (HPA 016487, Sigma), Polyclonal rabbit anti-human  $\text{InsP}_3\text{R2}$  1:1000 (AB9074, Millipore) and Polyclonal rat anti-mouse  $\text{InsP}_3\text{R3}$  1:500 (LC3, (26)). Original magnification x10. Size bar: 20  $\mu\text{m}$ .



**Supplementary Figure 3. Immunohistochemistry of  $\text{InsP}_3\text{R1}$  and  $\text{InsP}_3\text{R3}$  in mouse sweat glands.** Upper panel shows the immunohistochemistry of  $\text{InsP}_3\text{R1}$  in  $Itpr1^{+/+}$  and  $Itpr1^{-/-}$  mouse sweat glands. Lower panel shows the immunohistochemistry of  $\text{InsP}_3\text{R3}$  in  $Itpr3^{+/+}$  and  $Itpr3^{-/-}$  mice. Strong  $\text{InsP}_3\text{R3}$  signals at the apical site of clear cells were sometimes observed (arrows). Stain: Polyclonal rat anti-mouse  $\text{InsP}_3\text{R1}$  (4C11 (28)) and polyclonal rabbit anti-mouse  $\text{InsP}_3\text{R3}$  (LC3 (26)).

**Supplementary video 1. The pseudo-colored image of  $\text{Ca}^{2+}$  signals in mouse sweat glands in response to acetylcholine.** The isolated sweat glands from wild-type mice were sequentially stimulated with 30, 100, 300, 1000 nM of acetylcholine.