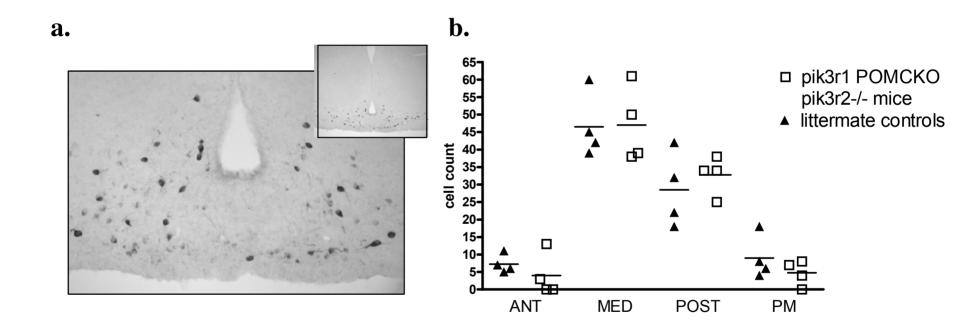
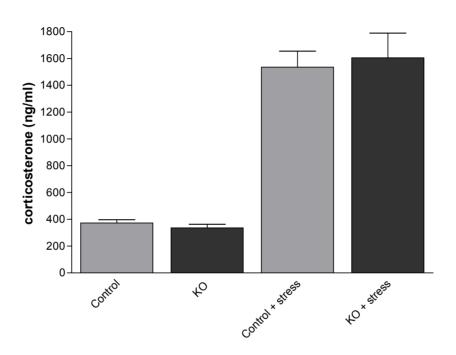
## Supplemental Table 1

	Size (pF)	Vm (mV)	Firing (Hz)	Input Resistance (MΩ)
Wildtype	13.8 ± 0.5	-4 .7 ± 0.8	1.2 ± 0.1	879.3 ± 9 .47
pik3r1 POMCKO	13.3 ± 0.9	-4 .1 ± 1.2	1.3 ± 0.2	
pik3r1 POMCKO pik3r2-/-	12.7 ± 0.5	-4 .7 ± 1.6	1.3 ± 0.1	1050 ± 102.5

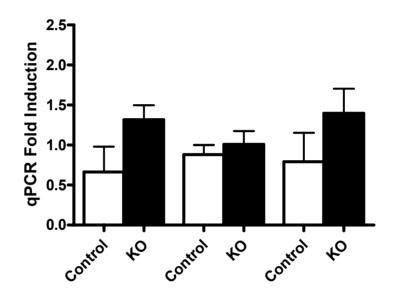
## **Supplemental Figure 1**



## **Supplemental Figure 2**



## **Supplemental Figure 3**



Supplemental Figure 1. POMC cell morphology and numbers are similar in control and *pik3r1 POMCKO pik3r2*\_/\_ hypothalami. (A) Immunohistochemical stain for β-endorphin in a *pik3r1 POMCKO pik3r2*\_/\_ medial hypothalamus. (B) POMC cell counts in *pik3r1 POMCKO pik3r2*\_/\_ (open squares) and littermate controls lacking POMC-cre expression (filled triangles) in mouse hypothalami at four levels (n=4). B-endorphin staining cells were counted on a single side per section. ANT = Bregma -2.3 mm, Interaural 6.7 mm, MED = Bregma -3.14 mm, Interaural 5.86 mm, POST = Bregma -3.8 mm, Interaural 5.2 mm, PM (premammillary) = Bregma -4.52 mm, Interaural 4.48 mm.

Supplemental Figure 2. Normal corticosterone secretion in pik3r1 POMCKO pik3r2—mice. (A) Corticosterone levels (ng/ml) of male littermate control (n = 6) and pik3r1 POMCKO pik3r2—mice (n = 8) were studied under basal conditions after three days of individual housing. (B) Corticosterone levels in male mice after 30 minutes of housing with novel males. Difference between basal and stressed: p<0.0001 for both groups. No significant difference exists between the pik3r1 POMCKO pik3r2—mice and littermate controls under either condition. Data are given as mean  $\pm$  SEM.

Supplemental Figure 3. Hypothalamic neuropeptide expression in young mice. (B) NPY, AgRP and POMC expression in hypothalami from 8 week old male control or pik3r1  $POMCKO\ pik3r2$ —mice (n = 4) as measured by qPCR.