



Figure S1. GnaqRQ Mouse Gene Targeting Approach

Homologous recombination was used to replace the endogenous exon 4 of *Gnaq* with a construct allowing for Cre recombinasedependent expression of p.R183Q GNAQ. The following elements were present in the targeting construct: 1) a cDNA block of mouse *Gnaq* exons 4-7 encoding the WT form of the protein, 2) a neomycin resistance cassette (green box) flanked by FRT sites (yellow triangles) for FRT-mediated recombination, 3) loxP sites (blue triangles) for Cre mediated recombination of the WT *Gnaq* cDNA block and the FRT site, and 4) a cDNA block of mouse *Gnaq* exons 4-7 encoding the p.R183Q mutant form of the protein. The 3'UTR and poly(A) site from Bovine growth hormone 1 (BGH1, pink block denoted pA) was ligated to the 3' end of exon 7 to provide a way to distinguish transcripts from the conditional allele from the endogenous *Gnaq*.

Figure S2



Figure S2. *Gnaq* transcript levels in embryos from *Gnaq*RQ^{fl/wt} x *Gnaq*RQ^{fl/wt}; β -actin-Cre+ cross. qRT-PCR results from e14.5 embryonic mouse tissue from *Gnaq*RQ^{fl/wt} and *Gnaq*RQ^{fl/wt}; β -actin-Cre+ littermates using allele specific primers revealed that *Gnaq* conditional allele transcripts were significantly higher than transcripts from the endogenous allele in both groups (p*** = 0.0002 and p** = 0.007 conditional *vs* endogenous allele, for *Gnaq*RQ^{fl/wt} and *Gnaq*RQ^{fl/wt}; β -actin-Cre+, respectively, two tailed paired *t*-test) with a greater increase in *Gnaq*RQ^{fl/wt}; β -actin-Cre+ (p^{###}<0.001 *Gnaq*RQ^{fl/wt}; β -actin-Cre+ vs *Gnaq*RQ^{fl/wt}, two tailed unpaired *t*-test).