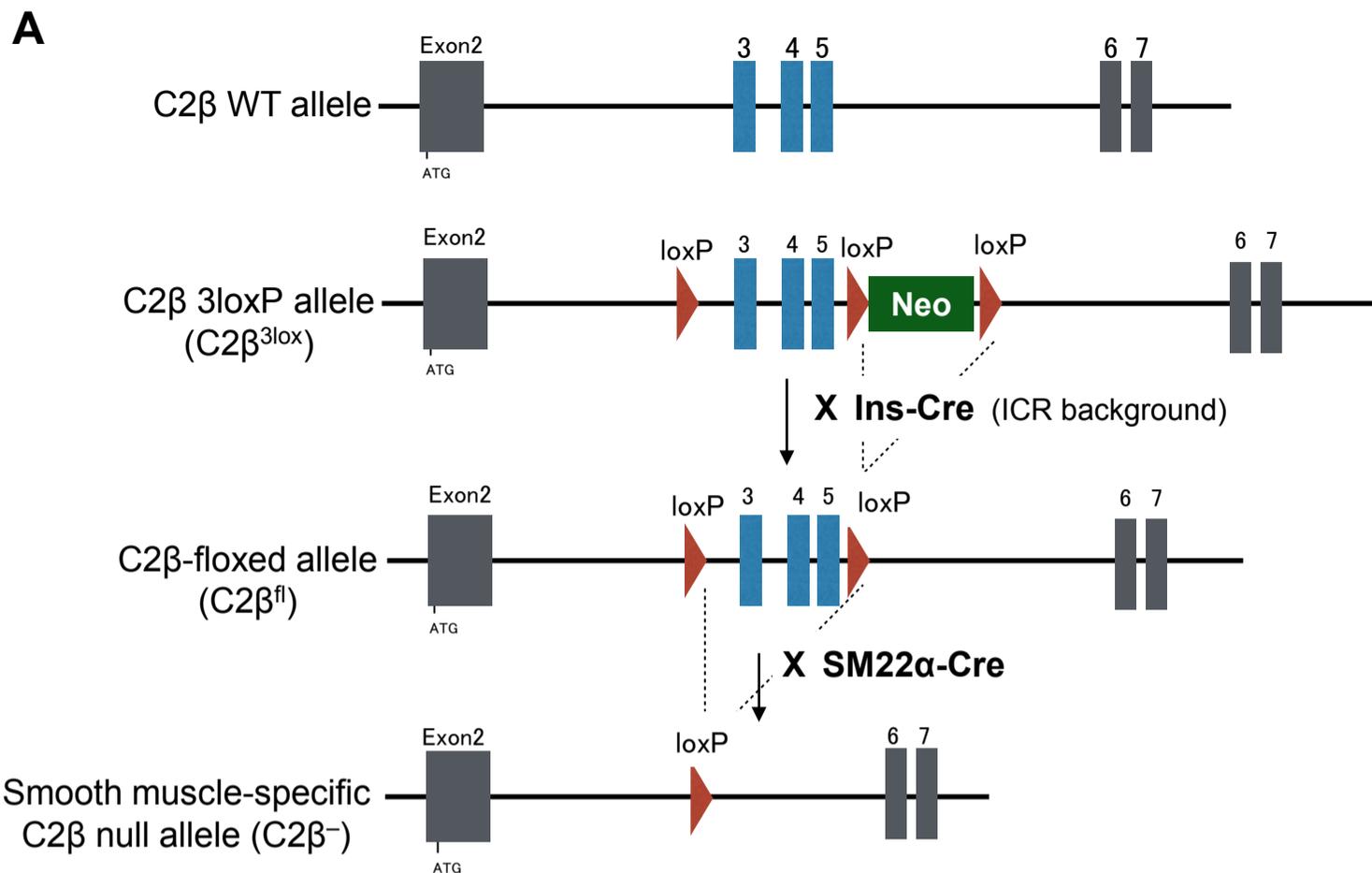


# Class II PI3Ks $\alpha$ and $\beta$ Are Required for Rho-Dependent Uterine Smooth Muscle Contraction and Parturition in Mice

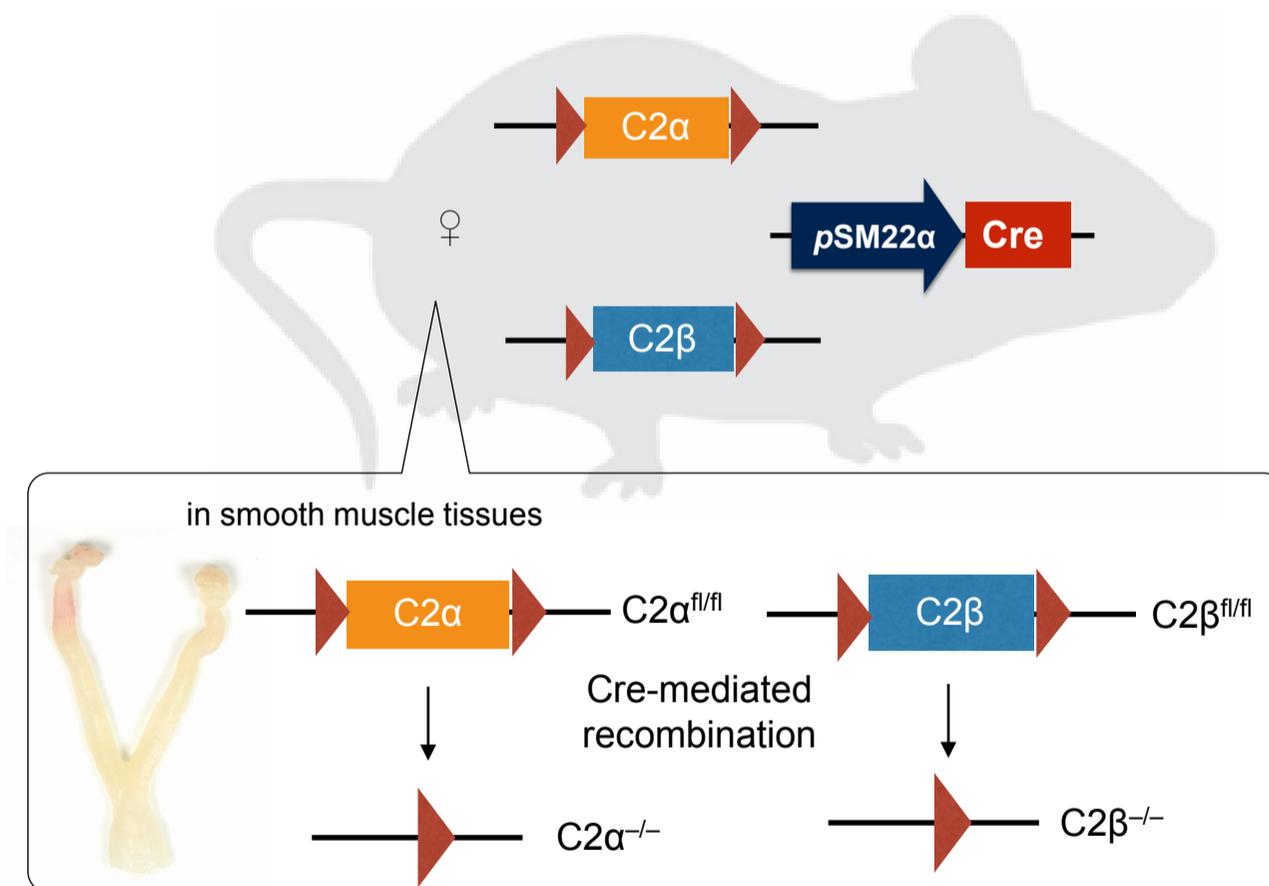
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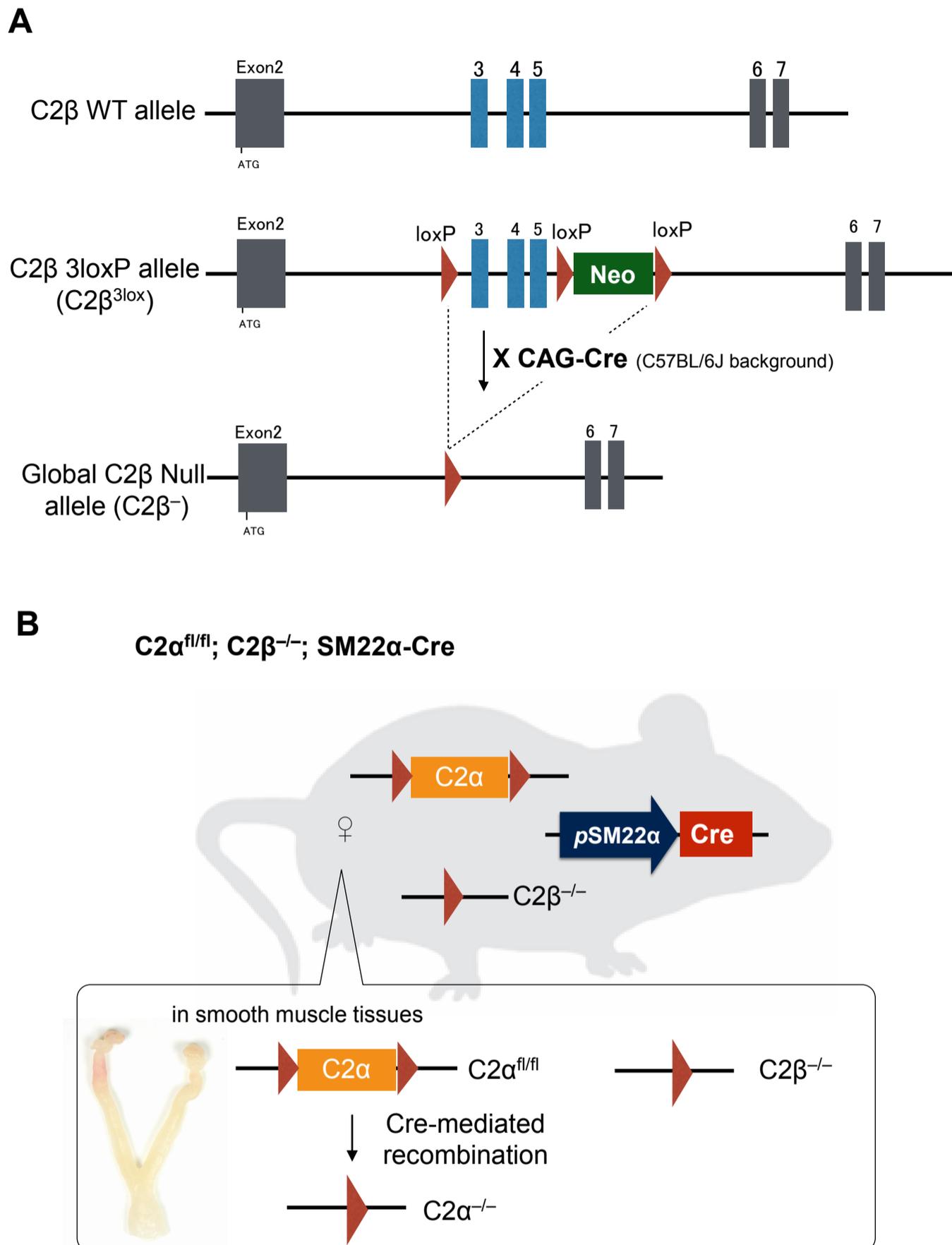


**B** **smDKO:  $C2\alpha^{fl/fl}$ ;  $C2\beta^{fl/fl}$ ; SM22 $\alpha$ -Cre**



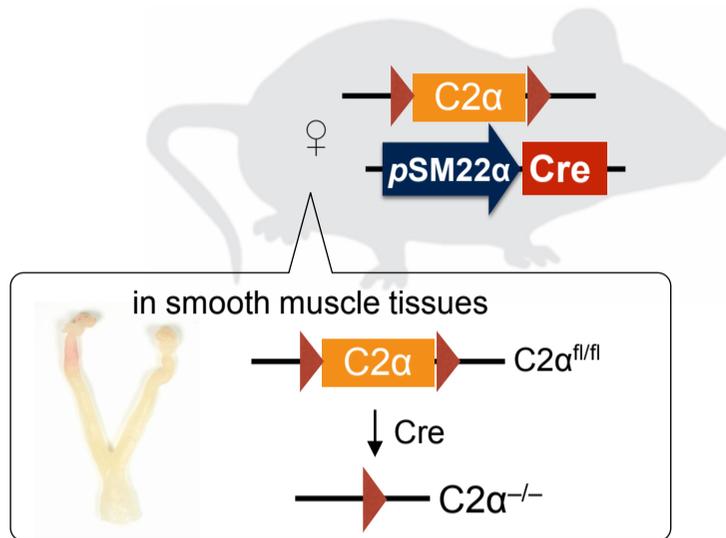
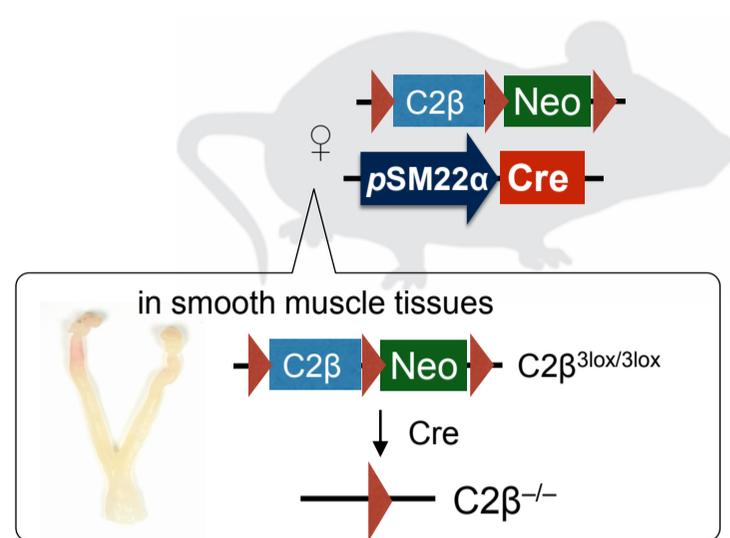
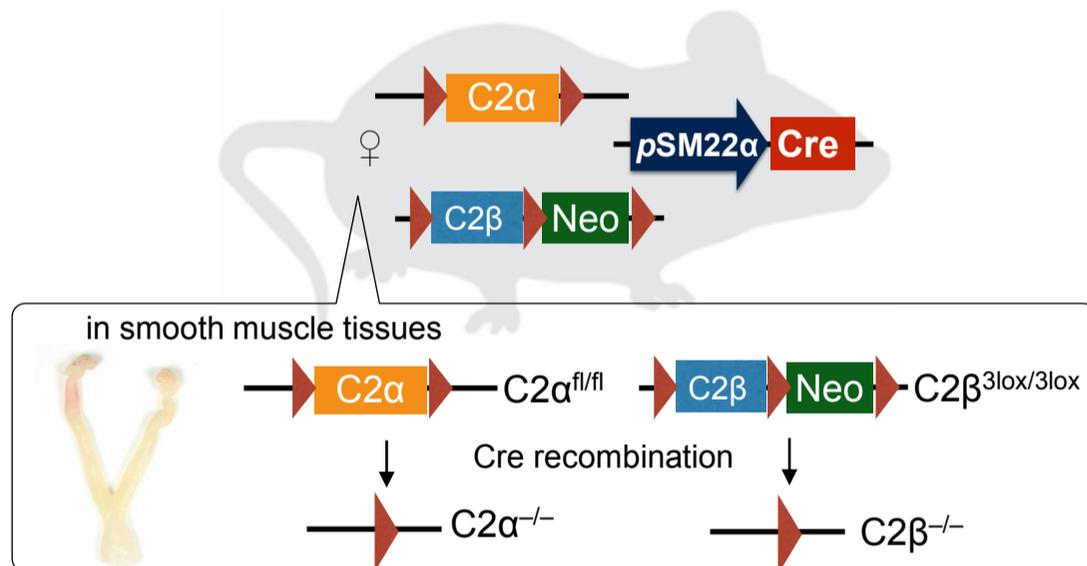
**Supplementary Figure 1. Generation of C2 $\beta$  floxed mice and smooth muscle-specific deletion of C2 $\alpha$  and C2 $\beta$ .**

A. C2 $\beta$  3loxP allele has one loxP before exon 3 and the two loxPs flanking the neomycin cassette (Neo). To produce C2 $\beta$ -floxed allele, the Neo was specifically deleted by mating with Ins-Cre mice (ICR strain), which expressed Cre recombinase (Cre) downstream of the insulin promoter in fertilized eggs. C2 $\beta$ -floxed allele was specifically deleted in smooth muscle by Cre that are expressed by SM22 $\alpha$  promoter. B. In smDKO mice, both C2 $\alpha$ - and C2 $\beta$ -floxed alleles are specifically deleted in smooth muscle by SM22 $\alpha$  promoter-driven expression of Cre.



**Supplementary Figure 2. Generation of  $C2\beta$ -null mice and smooth muscle-specific deletion of  $C2\alpha$ -floxed allele in  $C2\beta$ -null mice.**

A.  $C2\beta$  3loxP allele-carrying mice were crossed with mice that expressed Cre downstream of the universal CAG promoter, leading to generation of mice with  $C2\beta$ -null allele. B.  $C2\alpha$ -floxed allele is specifically deleted in smooth muscle by SM22 $\alpha$  promoter-driven expression of Cre.

**A****smC2 $\alpha$ KO: C2 $\alpha$ <sup>fl/fl</sup>; SM22 $\alpha$ -Cre****B****smC2 $\beta$ KO: C2 $\beta$ <sup>3lox/3lox</sup>; SM22 $\alpha$ -Cre****C****C2 $\alpha$ <sup>fl/fl</sup>; C2 $\beta$ <sup>3lox/3lox</sup>; SM22 $\alpha$ -Cre**

**Supplementary Figure 3 Smooth muscle-specific single deletion of C2 $\alpha$ -floxed allele (smC2 $\alpha$ KO) (A) and C2 $\beta$  3loxP allele (smC2 $\beta$ KO) (B), and deletion of both C2 $\alpha$ -floxed allele and C2 $\beta$  3loxP allele (C) by SM22 $\alpha$  promoter-driven expression of Cre.**