S52-1 Role of TRPC3/NCX1 in synpathetic regulation of vascular tone

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the molecular mechanisms underlying α_1 -AR-mediated vasoconstriction remain obscure. We found that phenylephrine-induced cytosolic Ca²⁺ elevation and contraction were greater in mesenteric arteries from TRPC3- or NCX1-transgenic mice (TG). In these mice, a bolus injection of norepinephrine elicited ST elevation and AV block (coronary spasm), which were suppressed by

SEA0400 (NCX1 inhibitor). When we crossed TRPC3-TG with NCX1-knokout mice or NCX1-TG with dominant negative TRPC3-TG, their offspring mice did not exhibit α_1 -AR-induced hypervasoreactivity. Coimmunoprecipitation, sucrose gradient fractionation, and immunolocalization experiments revealed that NCX1 and TRPC3 are interactively enriched in caveolar raft domains of vascular myocytes. These findings indicate that TRPC3/NCX1 coupling plays a pivotal role in regulating arterial tonus via α_1 -AR.

 α_1 -adrenoceptor (AR) contributes to the sympathetic regulation of various arteries. However,