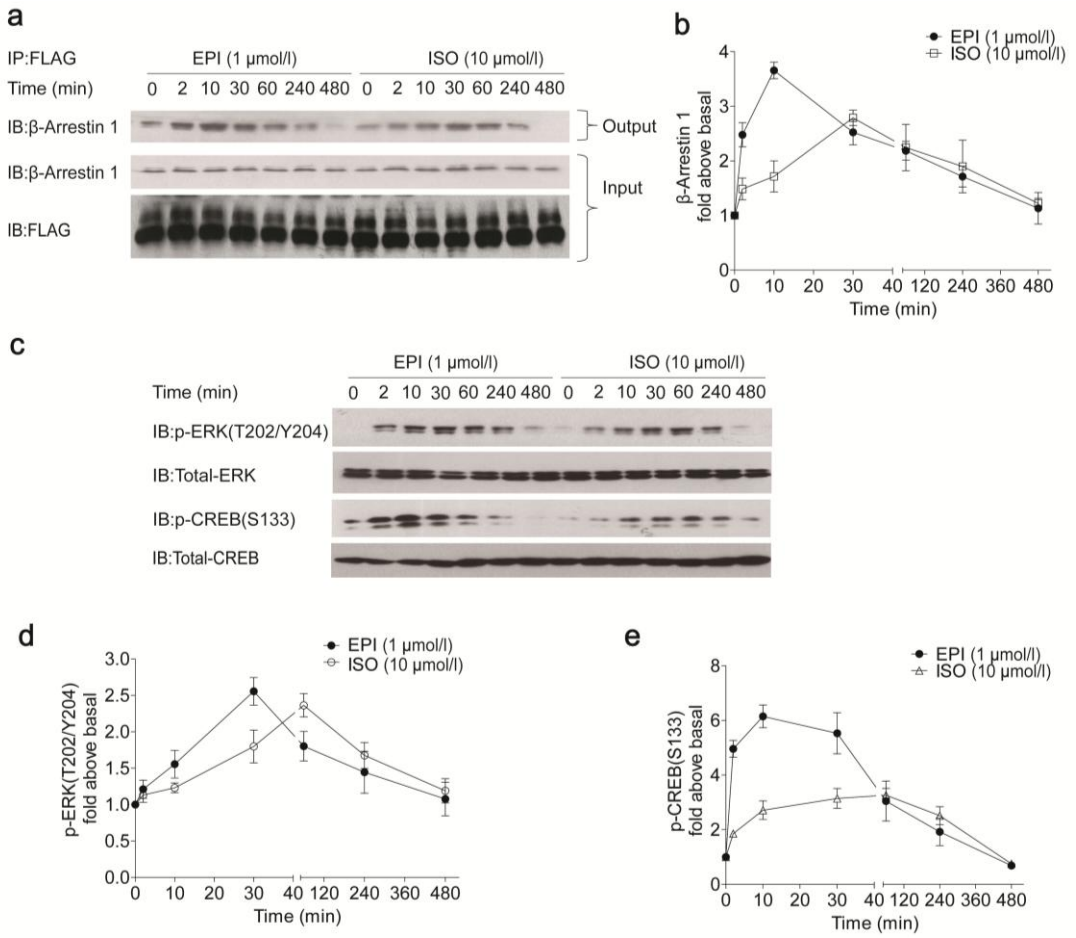
***ESM*-Fig. 2 The time course of the effects of EPI or ISO on the β2-AR/β-arrestin1 complex formation, ERK phosphorylation and CREB phosphorylation.**



***ESM*-Fig. 2 The time course of the effects of EPI or ISO on the β2-AR/β-arrestin1 complex formation, ERK phosphorylation and CREB phosphorylation.**

(a). Detection of EPI/ISO induced β2-AR/β-arrestin1 complex formation. TGP52 cells expressing Flag-tagged β2-ARs were incubated for the indicated times in the presence of EPI or ISO. The receptor-β-arrestin1 complexes were stabilized by covalent cross-linking with DSP and then immunoprecipitated by a Flag antibody. The formation of the β2-AR/β-arrestin1 complex was detected by an anti-β-arrestin1 antibody (ACT1).

(b). Signals from (a) were quantified and normalized to the zero time point. Data are the means ± SEM of three independent experiments.

(c). The time course of the effects of epinepherine or Isoproterenol on ERK phosphorylation or CREB phosphorylation. The TGP52 cells were incubated with epinephrine or isoproterenol for the indicated times; the phosphorylation of ERK at pT^{202}/pY^{204} and CREB at pS^{133} was detected by specific antibodies.

(d-e). Signals for phosphorylation of ERK at pT^{202}/pY^{204} (d) or REB at pS^{133} (e) were quantified and normalized to the zero time point. Data are the means ± SEM of at least three independent experiments.