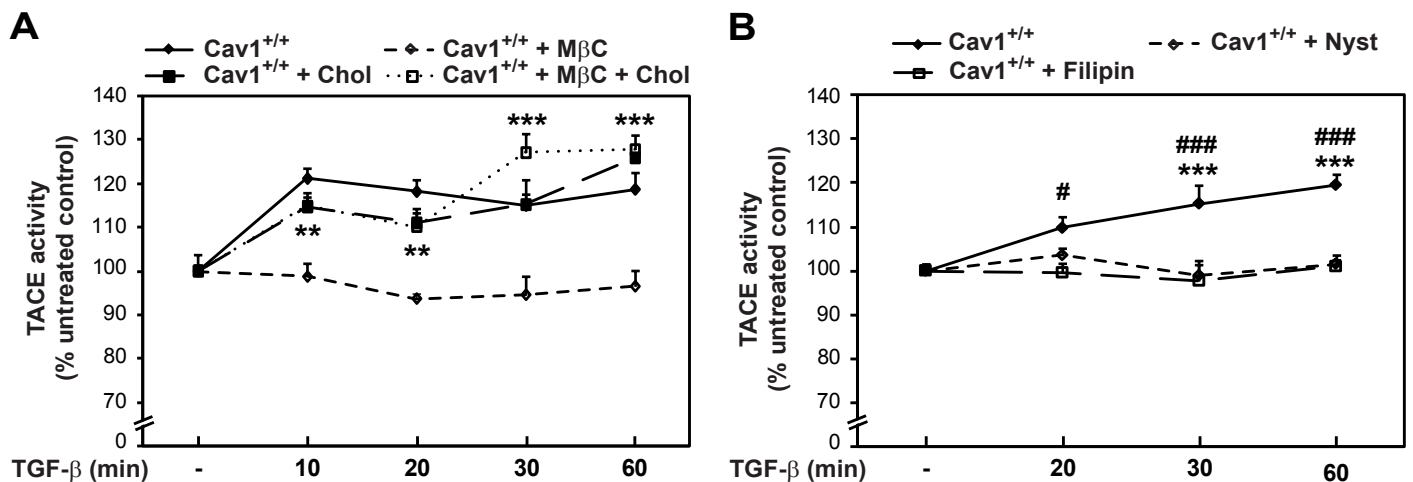




Supplementary Figure 6. Lipid raft integrity is necessary for TGF- β -mediated activation of TACE/ADAM17. Cav1^{+/+} hepatocytes treated with TGF- β (2 ng/ml) at different times after previous FBS starvation (16 h). Metalloprotease TACE/ADAM17 activity expressed as percentage of activity/untreated control. A: Cells were treated in the absence (Cav1^{+/+}) or presence (Cav1^{+/+} + M β C) of M β C for 30 minutes. After this time, in half of the cells in each treatment group, medium was removed and fresh medium with cholesterol (1 mM) was added (Cav1^{+/+} + Chol; Cav1^{+/+} + M β C + Chol). After 1 hour, cells were treated with TGF- β and collected for TACE/ADAM17 activity analysis at the times indicated in the figure (N=3). B: Cells were treated with the cholesterol-binding agents nystatin (Nyst, 25 μ M) and filipin (0.5 μ M) added 30 minutes before TGF- β treatment (N=3). Results expressed as mean $\pm$ SEM. Statistical comparison using two-way ANOVA with Bonferroni post-hoc test: *p<0.05; **p<0.01; ***p<0.001 (A: (*) Cav1^{+/+} + M β C + Col vs. Cav1^{+/+} + M β C; B: (*) Cav1^{+/+} vs. Cav1^{+/+} + Nyst and (#) Cav1^{+/+} vs. Cav1^{+/+} + Filipin).