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## **Supporting Material**

## Redox-regulated heterogeneous thresholds for ligand recruitment among InsP3R Ca2+ release channels

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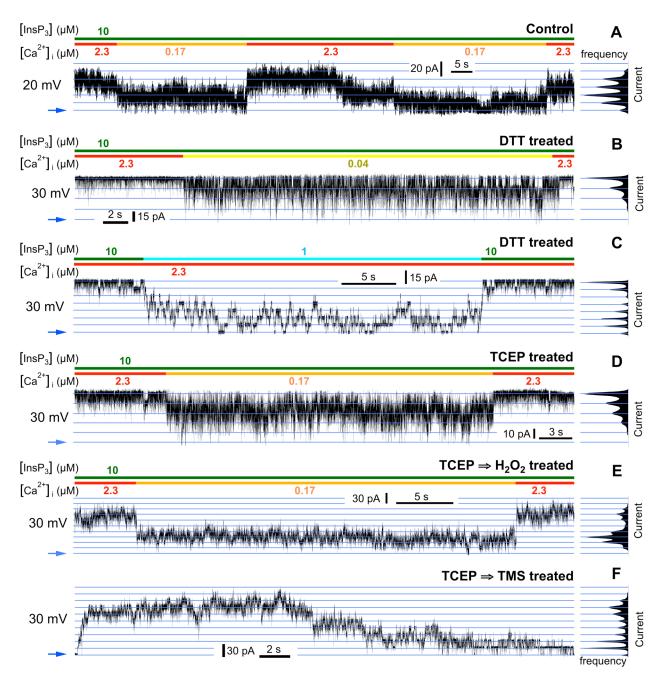


FIGURE S1 Current traces of homotetrameric recombinant  $InsP_3R-3$  channels in cyto-out membrane patches obtained from isolated DT40-KO-r-InsP<sub>3</sub>R-3 nuclei. (*A*) Current trace showing reversible changes in  $N_A$  when  $InsP_3R-3$  channels were alternately exposed to optimal (2.3 μM) and sub-optimal (170 nM)  $[Ca^{2+}]_i$ . (*B*) Current trace showing the lack of change in  $N_A$  when  $InsP_3R-3$  channels were alternately exposed to optimal and resting (40 nM)  $[Ca^{2+}]_i$  after DTT treatment (3 mM for > 60 min). (*C*) Current trace showing reversible decrease in  $N_A$  when DTT-treated channels were exposed to extreme reduction of  $[InsP_3]$  from 10 to 1 μM. (*D*) Current trace showing the absence of change in  $N_A$  when channels were alternately exposed to optimal and sub-optimal  $[Ca^{2+}]_i$  after TCEP treatment (6 mM for > 90 min). (*E*) Current trace showing the restoration of change in  $N_A$  when  $InsP_3R-3$  channels were alternately exposed to optimal and sub-optimal  $[Ca^{2+}]_i$  after treatment with TCEP (6 mM for 120 min) and then  $H_2O_2$  (5 mM for 60 min). (*F*) Current trace showing channel inactivation in constant presence of optimal  $[InsP_3]$  (10 μM) and  $[Ca^{2+}]_i$  (2.3 μM) after treatment with TCEP (6 mM for 120 min) and then TMS (100 μM for 60 min). At least 10 channels were activated by a jump in  $[InsP_3]$  from 0 to 10 μM at the beginning of the trace.