

LRRC8 channel complexes counterbalance K_{ATP} channels to mediate swell-secretion coupling in mouse pancreatic β -cells

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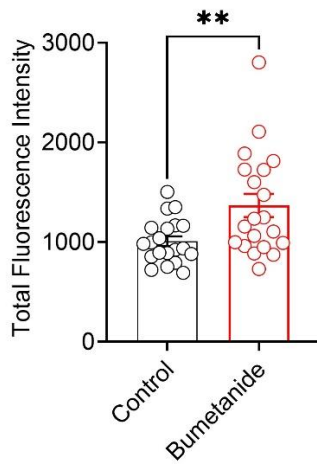
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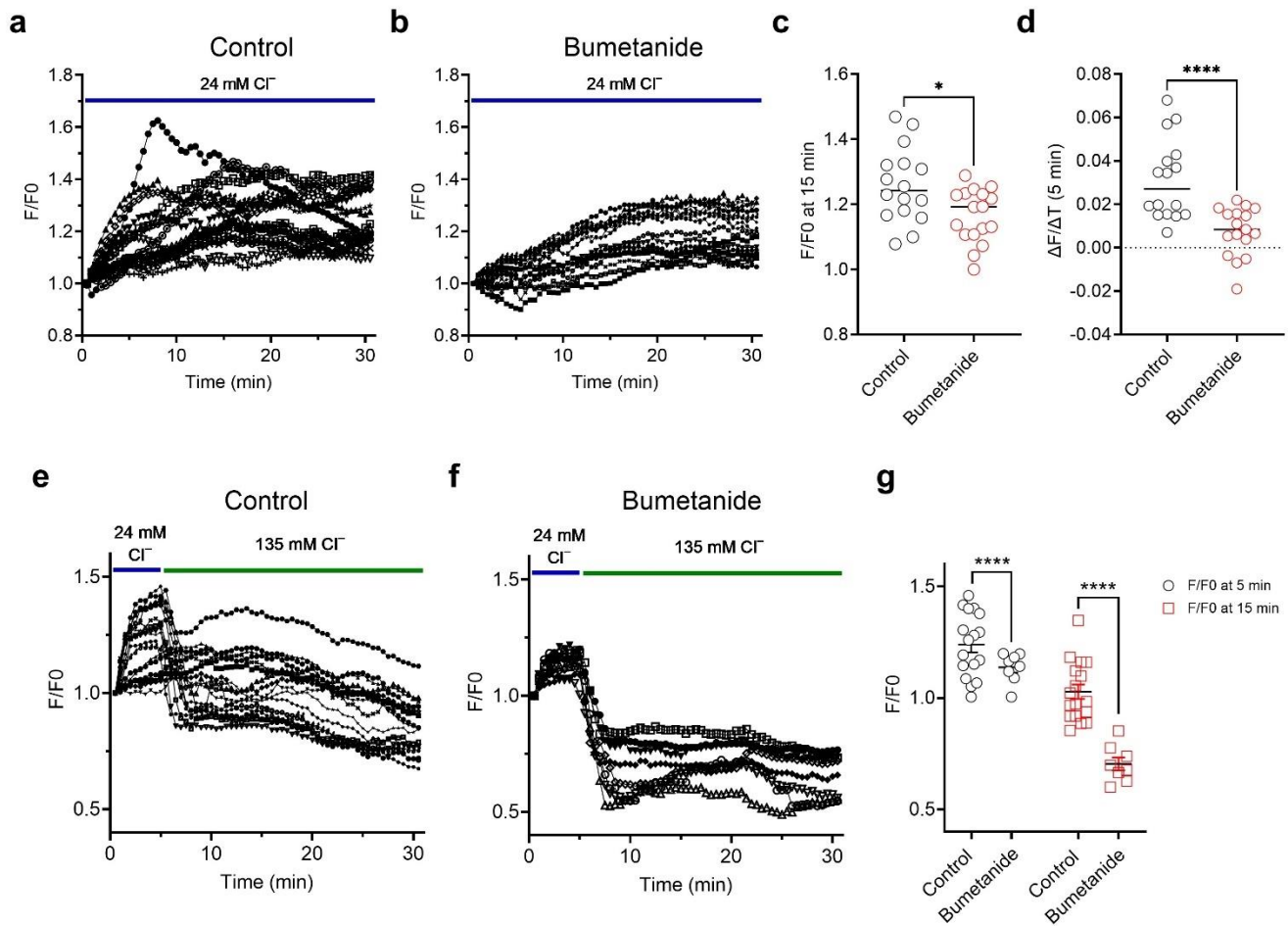
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Supplementary Figure 1. Bumetanide treatment reduces intracellular Cl^- concentration in β -cells.

Total MQAE fluorescence intensity in β -cells under control conditions ($n = 20$) and after treatment with $20 \mu\text{M}$ bumetanide ($n = 20$). Data are presented as mean $\pm$ S.E.M. Statistical significance for all data was determined using two-tailed unpaired Student's t-test (** $P < 0.01$).



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35 **Supplementary Figure 2. Bumetanide pre-treatment lowers β -cell intracellular Cl^- as compared to**
 36 **control thereby reducing Cl^- efflux and increasing Cl^- influx. a,b** Normalized MQAE fluorescence in
 37 control (n = 16) and bumetanide-treated β -cells (n = 17) during perfusion with a low- Cl^- solution (24 mM).
 38 **c** Mean normalized MQAE fluorescence in control and bumetanide-treated β -cells after 15 minutes of
 39 perfusion with a low- Cl^- solution (24 mM). **d** Rate of change of MQAE fluorescence ($\Delta F/\Delta T$) over 5
 40 minutes in control and bumetanide-treated β -cells upon perfusion with a low- Cl^- (24 mM) solution. **e,f**
 41 Normalized fluorescence intensity during the transition from media to a low- Cl^- solution (24 mM), and
 42 then to high- Cl^- solution (135 mM) in control (n = 17) versus bumetanide-treated β -cells (n = 8). **g** Mean
 43 normalized MQAE fluorescence in control and bumetanide-treated β -cells after 5 minutes of perfusion
 44 with a low- Cl^- solution (24 mM) and 15 minutes of perfusion with a high- Cl^- solution (135 mM). Statistical
 45 significance for all data was determined using two-tailed unpaired Student's t-test (*P < 0.05; ****P
 46 < 0.0001).