

THE EFFECT OF TRIPLE CFTR MODULATOR THERAPY AND AZITHROMYCIN ON ION CHANNELS AND INFLAMMATION IN CYSTIC FIBROSIS

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Supplementary Data

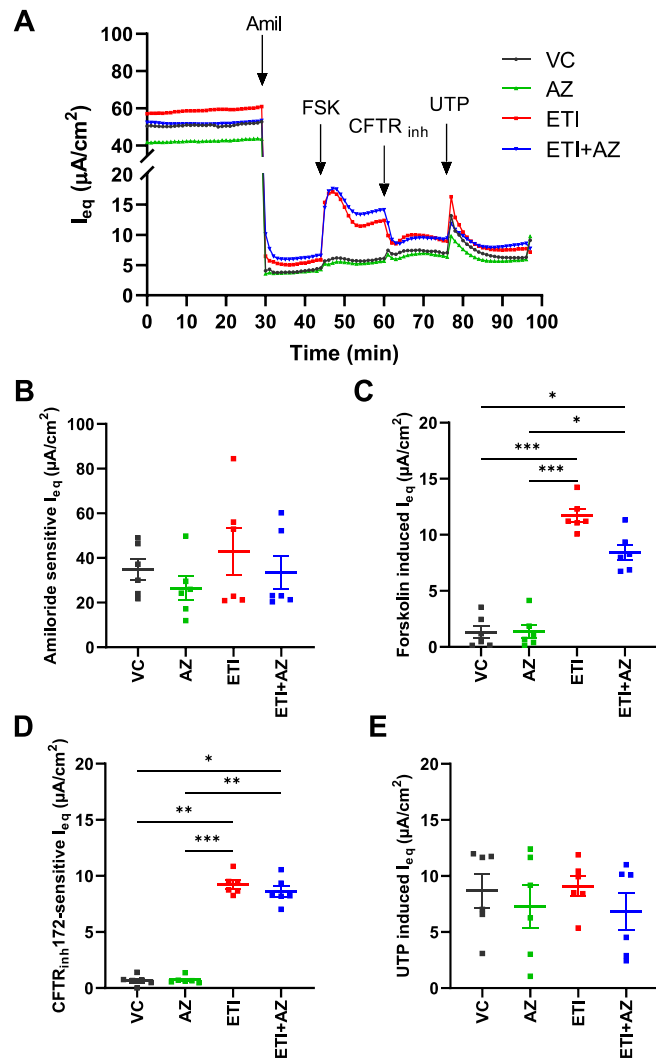


Figure S1: AZ has no synergistic/antagonist effect on ion channels upon combination with ETI in homozygous (F508del/F508del) CF HBE cells at 96 hours. (A) Representative traces of I_{eq} over time. CF HBEs primary cells grown at air liquid interface were pretreated basolaterally for **96 hours** with: vehicle (DMSO 0.08%; v/v) and AZ (10 $\mu\text{g}/\text{mL}$). ETI (ELX/TEZ/LUM; 3/18/1 μM) was added to all assigned wells 24 hours prior to experiment. Changes in I_{eq} were measured using the MTECC-24 system after the apical addition of amiloride (10 μM), Forskolin (20 μM), $\text{CFTR}_{inh}172$ (20 μM) and UTP (100 μM). Quantified (B) Amiloride sensitive I_{eq} ; (C) Forskolin induced I_{eq} ; (D) $\text{CFTR}_{inh}172$ sensitive I_{eq} and (E) UTP sensitive current responses ($\mu\text{A}/\text{cm}^2$) are shown. Data presented as the mean \pm SEM. Statistical analyses were performed using a Kruskal-Wallis statistical test with Dunn's multiple comparisons post-hoc test. * p value ≤ 0.05 . N=6 (3 filters each from two donors).

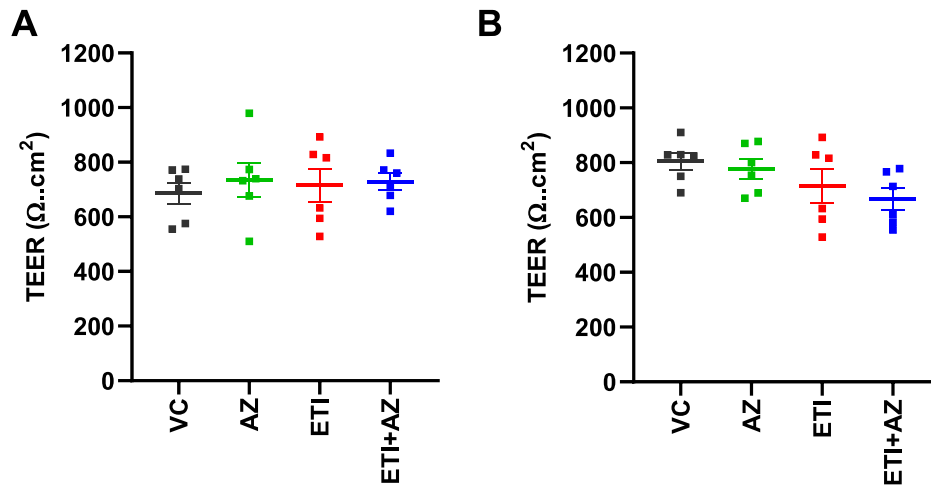


Figure S2: TEER assessment of homozygous (F508del/F508del) CF HBE cells over the course of the treatment period. CF HBEs primary cells grown at air liquid interface were pretreated basolaterally for (A) 24 and (B) 96 h with: vehicle (DMSO 0.08%; v/v) and AZ (10 $\mu\text{g}/\text{mL}$). ETI (ELX/TEZ/LUM; 3/18/1 μM) was added to all assigned wells 24 hours prior to measurement. TEER values ($\Omega\cdot\text{cm}^2$) were measured using the MTECC-24 system. Data presented as the mean \pm SEM. Statistical analyses were performed using a Kruskal-Wallis statistical test. No significant difference was reported. N=6 (3 filters each from two donors).