



Supplementary Figure

The L467F-F508del Complex Allele Hampers Pharmacological Rescue of Mutant CFTR by Elexacaftor/Tezacaftor/Ivacaftor in Cystic Fibrosis Patients: The Value of the Ex Vivo Nasal Epithelial Model to Address Non-Responders to CFTR-Modulating Drugs

Elvira Sondo ^{1,†}, Federico Cresta ^{2,†}, Cristina Pastorino ³, Valeria Tomati ¹, Valeria Capurro ¹, Emanuela Pesce ¹, Mariateresa Lena ¹, Michele Iacomino ¹, Ave Maria Baffico ⁴, Domenico Coviello ⁴, Tiziano Bandiera ⁵, Federico Zara ^{1,3}, Luis J. V. Galletta ^{6,7}, Renata Bocciardi ^{1,3}, Carlo Castellani ^{2,‡,*} and Nicoletta Pedemonte ^{1,‡,*}

¹ UOC Genetica Medica, IRCCS Istituto Giannina Gaslini, 16147 Genova, Italy; elvirasondo@gaslini.org (E.S.); valeriatomati@gaslini.org (V.T.); valeriapapurro@gaslini.org (V.C.); emanuela.pesce@yahoo.it (E.P.); mariateresalena@gaslini.org (M.L.); micheleiacomino@gaslini.org (M.I.); federico.zara@unige.it (F.Z.); renata.bocciardi@unige.it (R.B.)

² UOSD Centro Fibrosi Cistica, IRCCS Istituto Giannina Gaslini, 16147 Genova, Italy; federicocresta@gaslini.org

³ Department of Neurosciences, Rehabilitation, Ophthalmology, Genetics, Maternal and Child Health (DINOEMI), University of Genoa, 16132 Genova, Italy; cristinapastorino22@gmail.com

⁴ UOC Laboratorio di Genetica Umana, IRCCS Istituto Giannina Gaslini, 16147 Genova, Italy; avemariabaffico@gaslini.org (A.M.B.); domenicooviello@gaslini.org (D.C.)

⁵ D3-PharmaChemistry, Fondazione Istituto Italiano di Tecnologia, 16163 Genova, Italy; tiziano.bandiera@iit.it

⁶ Telethon Institute of Genetics and Medicine (TIGEM), 80078 Pozzuoli, Italy; l.galletta@tigem.it

⁷ Department of Translational Medical Sciences (DISMET), University of Naples Federico II, 80131 Naples, Italy

* Correspondence: carlocastellani@gaslini.org (C.C.); nicolettapedemonte@gaslini.org (N.P.)

† These authors contributed equally as first authors.

‡ These authors contributed equally as last authors.

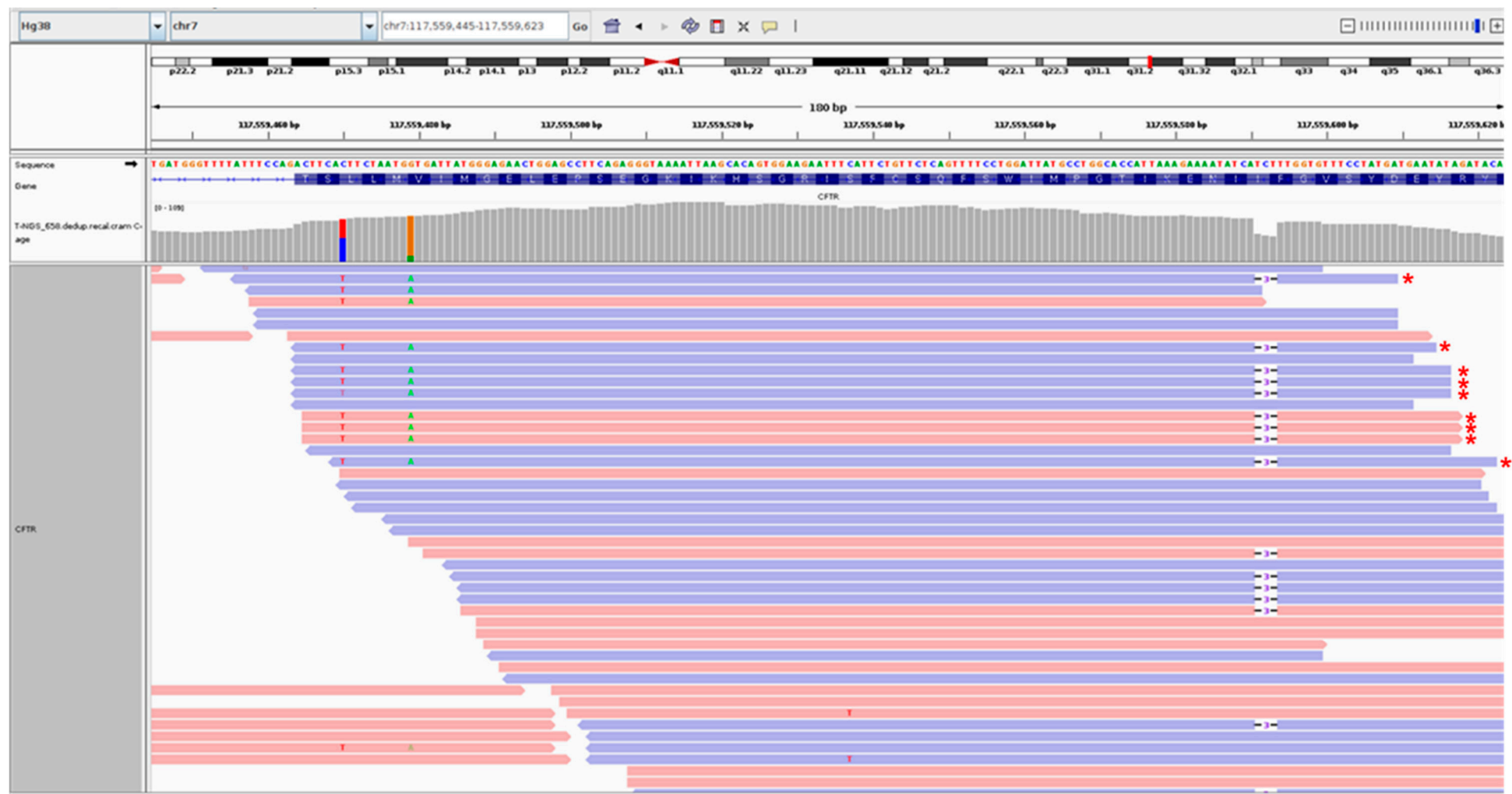


Figure S1. The L467/F508del complex allele and the Gaslini exome genomic database. Genomic window spanning a region of the *CFTR* exon 11 generated by the Integrative Genomics Viewer (IGV) and displaying the BAM alignment track mapped read-segments per reference sequence (GRCh38/hg38). Red asterisks indicate the reads showing the mismatch positions corresponding to the presence of the L467F (T, red) and F508del (-3-) variants in cis in the same sequence fragment. A further mismatch (A, green) in cis with the two variants corresponds to the presence of the V470M polymorphism