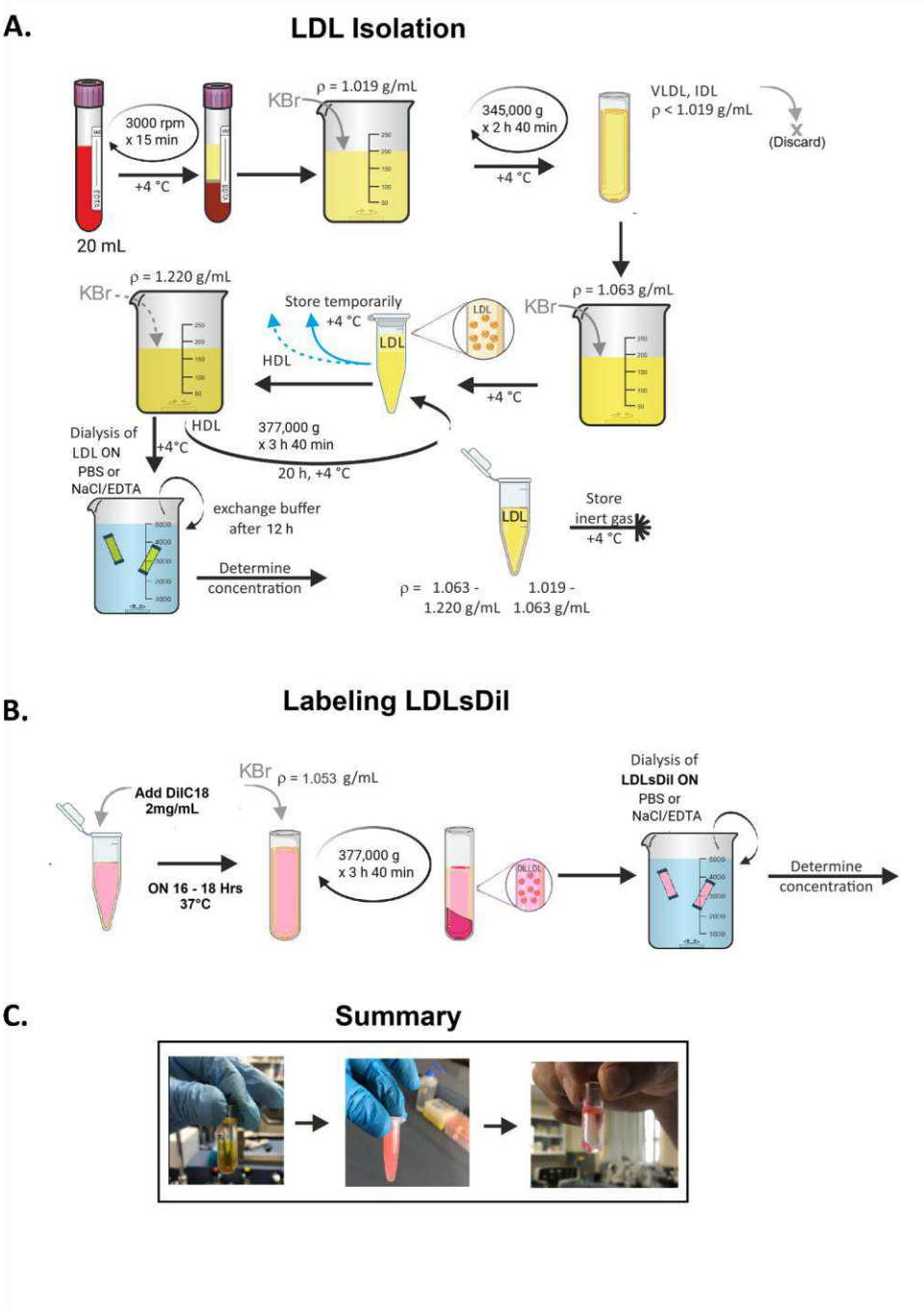
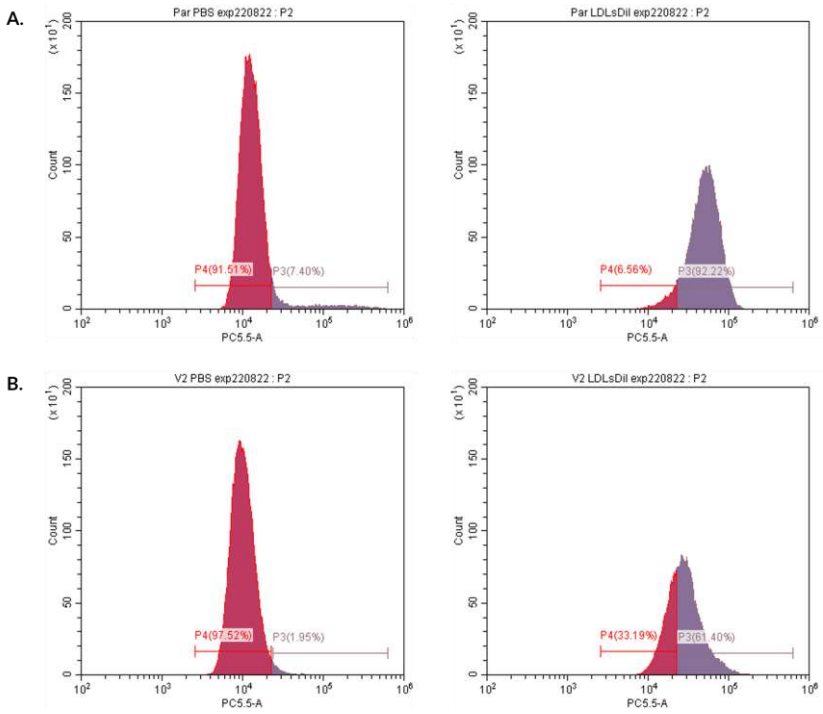
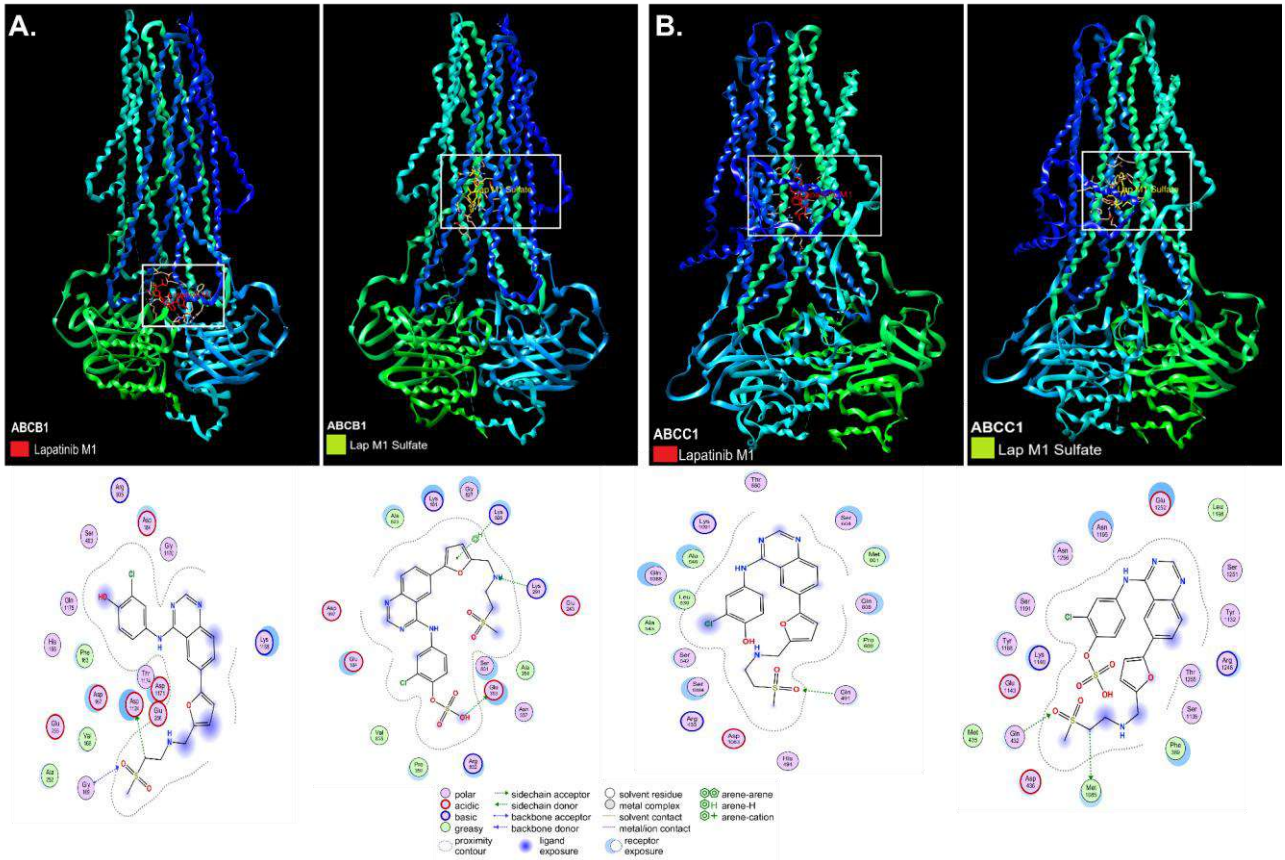
Supplementary Material



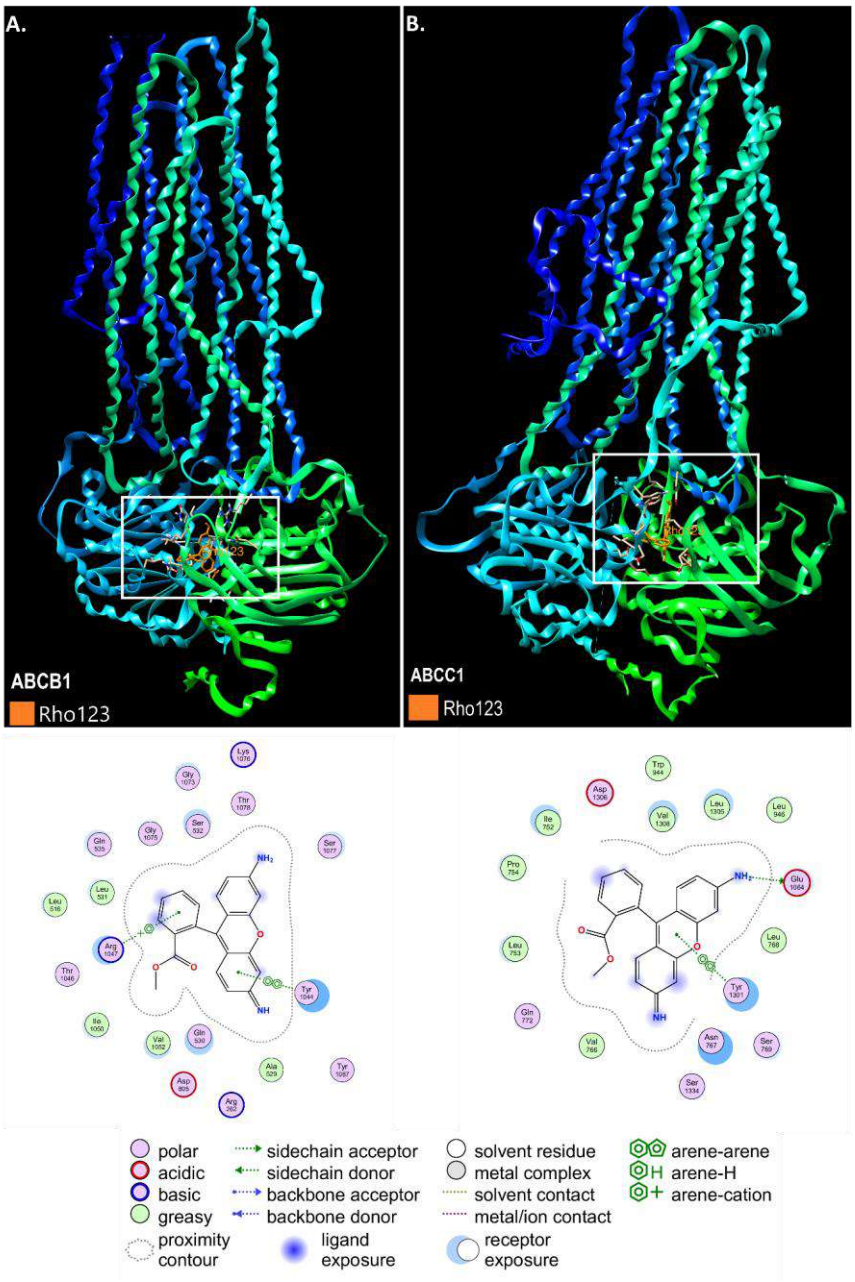
Sup. fig. 1. Isolation and labeling of LDLs with the DilC18 probe. A. The procedure for the isolation of lipoproteins is observed using the differential ultracentrifugation technique; an S140-AT 2555 rotor was used (Taken and modified from Axmann et al., 2019). **B.** Labeling of nLDLs subsequent to their isolation. **C.** A summary of the process with real pictures.



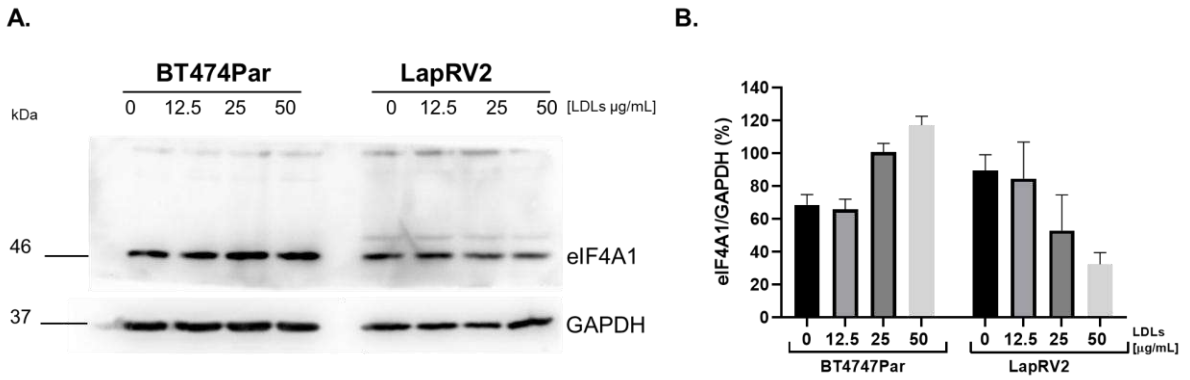
Sup. fig. 2. Histograms of DiI-LDL internalization (0–50 µg/mL) and incubation of 24 h. (**A–B**) percentages comparing Parental and Variant 2 (LapR V2). **A**. Left histogram corresponds to BT474 parental control cells, i.e., without treatment. Right histogram corresponds to parental cells treated with LDLs labeled with the fluorescent probe DiI. **B.** Left histogram corresponds to resistant cells (LapR V2) without any treatment, using only 1X PBS as the vehicle. Right histogram corresponds to the internalization percentages obtained for LapRV2 resistant cells labeled with the probe D282. Using PBS 1X as a vehicle control.



Sup. fig. 3. Molecular docking of Lapatinib M1 and Lapatinib M1 Sulfate and the ABC transporter**s.** **A.** Figure corresponding to the ligand Interaction of ABCB1 transporter (6c0v) + Lapatinib M1 and Lap M1 Sulfate. **B.** Figures corresponding to the Ligand Interaction ABCC1 (6uy0) + Lapatinib M1 and Lap M1 Sulfate. In both transporters Lapatinib M1 is shown in red and Lapatinib M1 Sulfate in yellow-green.



Sup. fig. 4. Molecular docking of Rhodamine 123 and the ABC transporters. A-B. Figures corresponding to the Ligand Interaction of ABCB1 transporter (PDB:6c0v) and Ligand Interaction of ABCC1 (6uy0) vs Rhodamine123 as a control. Rho123 are shown in orange.



Sup fig. 5. **eIF4A1 expression under LDLs treatments A.** Representative Western blots showing protein expression of levels of eIF4A1 in BT474Par and LapRV2 cells. **B.** Densitometry analysis of eIF4A1 under LDL increasing concentrations (0-50 µg/mL) for 48 h. Data represent the mean ± SD of three independent experiments, and GAPDH was used as a loading control.