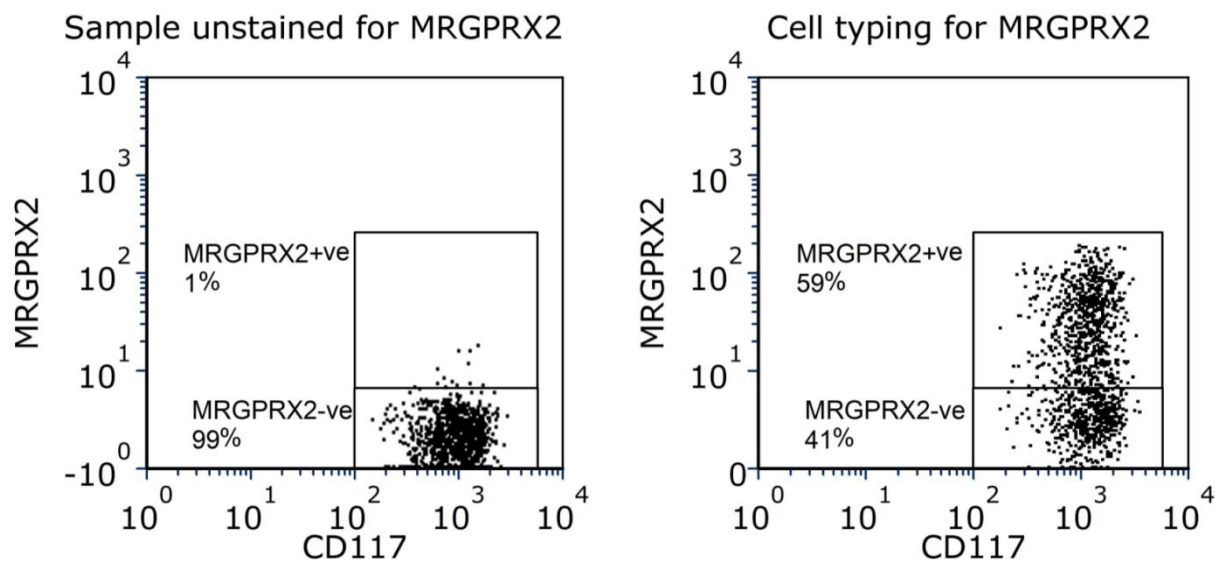


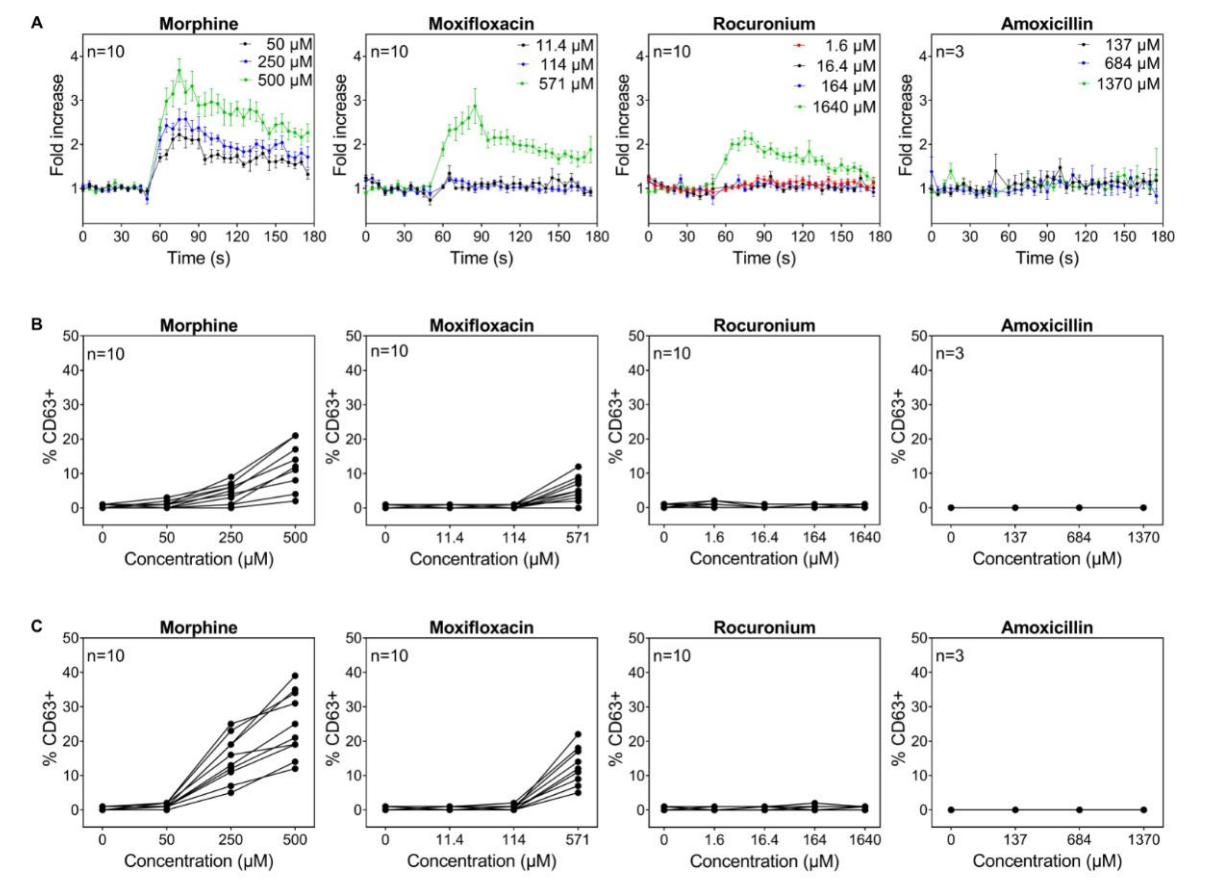
SUPPLEMENTARY MATERIAL

Figure 1. Representative plot for the MRGPRX2 expression on PBCMCs.

Peripheral blood cultured mast cells (PBCMCs) are defined as $CD117^{+ve}CD203c^{+ve}$ cells.

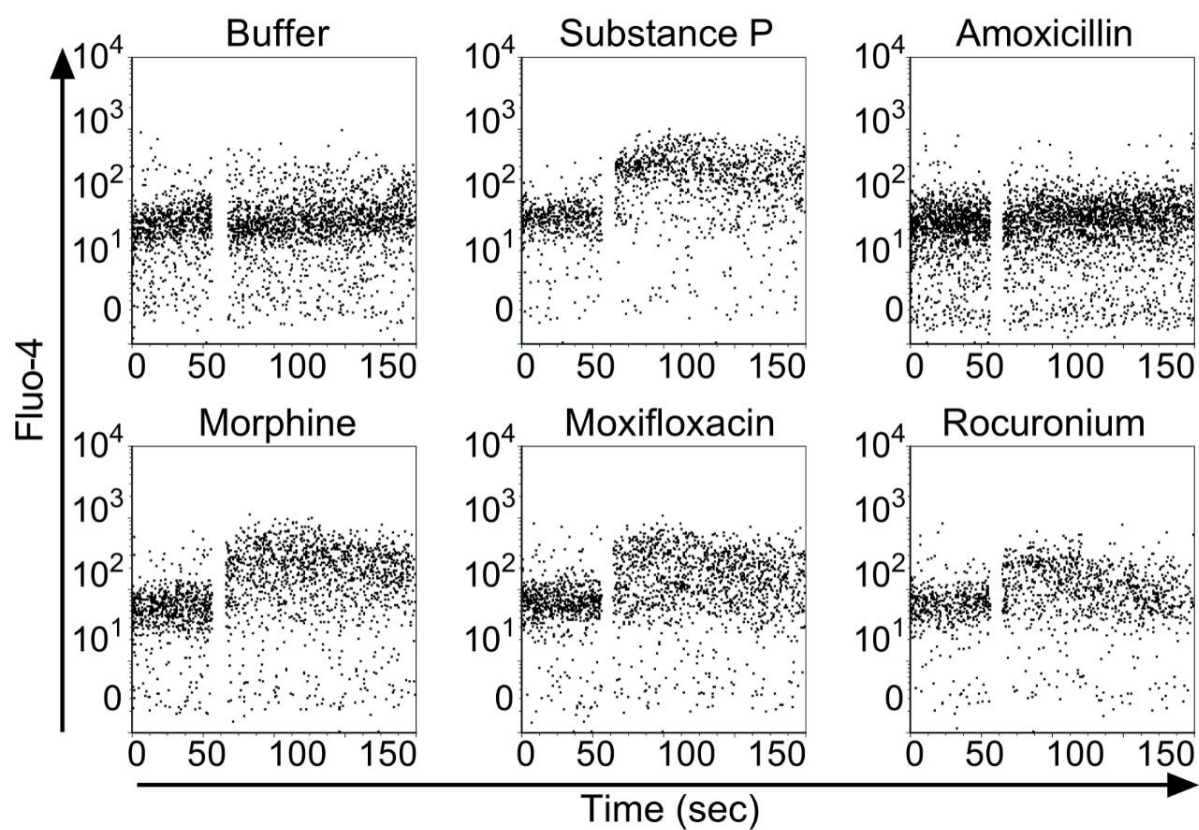
PBCMCs harbour two subpopulations: cells with surface expression of MRGPRX2 ($MRGPRX2^{+ve}$) and cells without expression of MRGPRX2 ($MRGPRX2^{-ve}$).

Figure 2. Dose-response curves for changes in intracellular calcium and CD63 expression in PBCMCs.



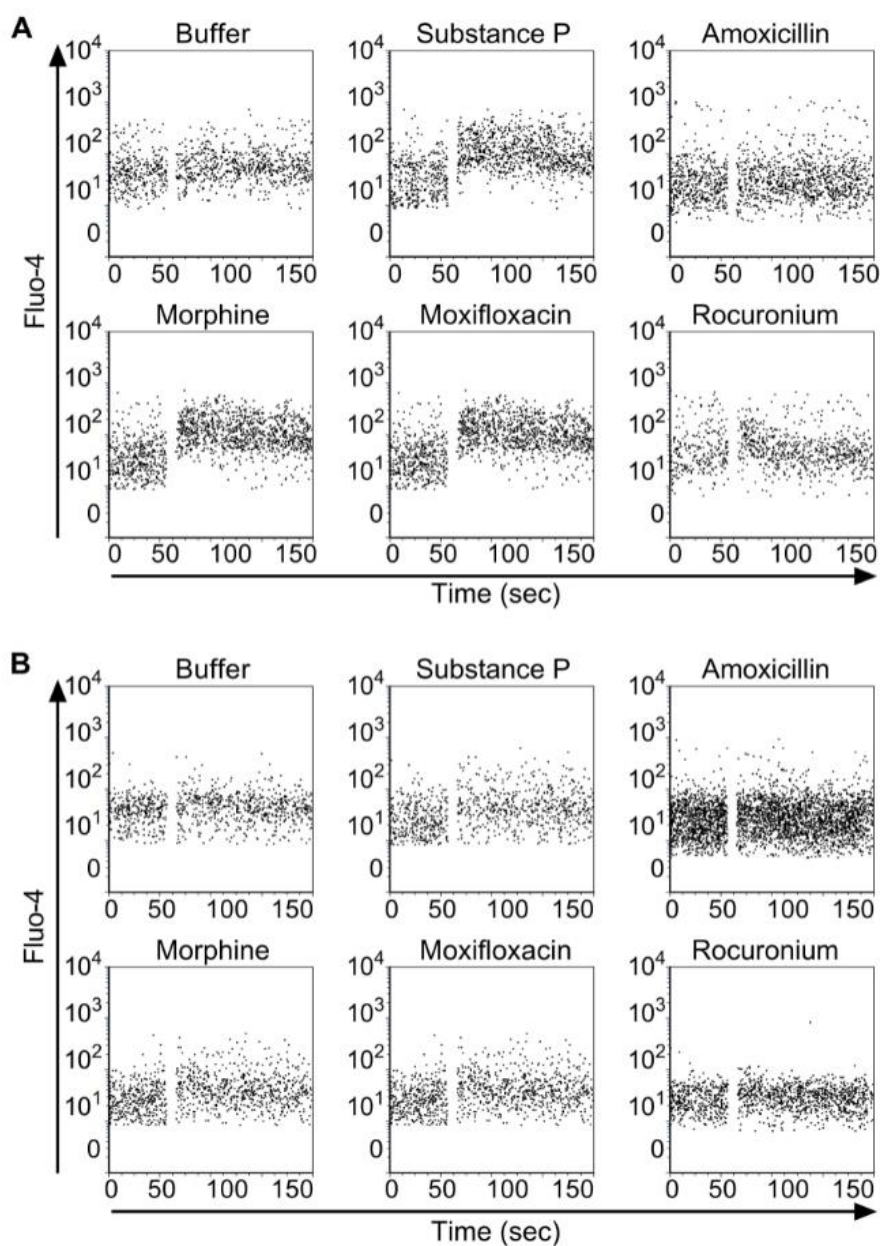
(A) Dose-response curves of intracellular calcium levels. **(B)** Dose-response curves of CD63 up-regulation after 3 min of stimulation or **(C)** after 20 min of stimulation. Attempts to increase the rocuronium concentration revealed to be cytotoxic.

Figure 3. Representative plots for intracellular calcium imaging in PBCMC.



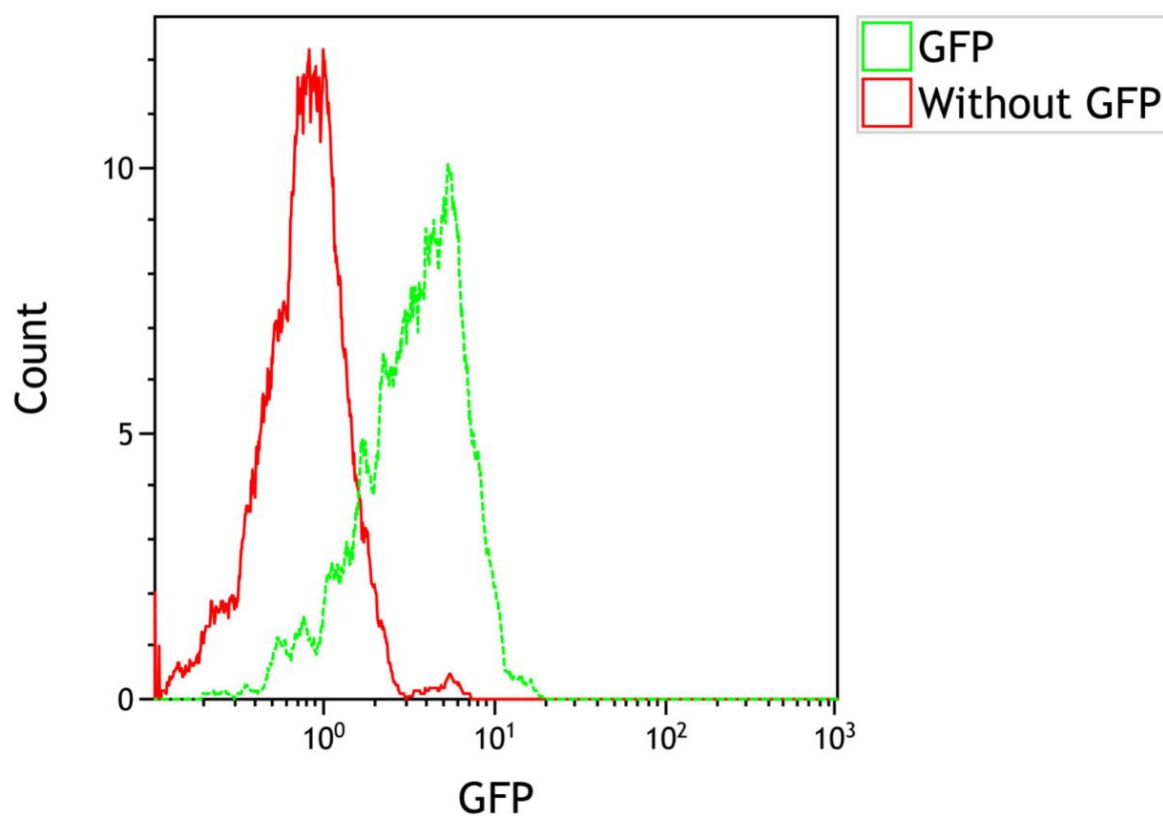
PBCMCs were, after 50 sec, stimulated with buffer, substance P (74 μ M), amoxicillin (1370 μ M), morphine (500 μ M), moxifloxacin (571 μ M) or rocuronium (1640 μ M).

Figure 4. Representative plot for intracellular calcium imaging in MRGPRX2^{+ve} (A) and MRGPRX2^{-ve} (B) subpopulations.



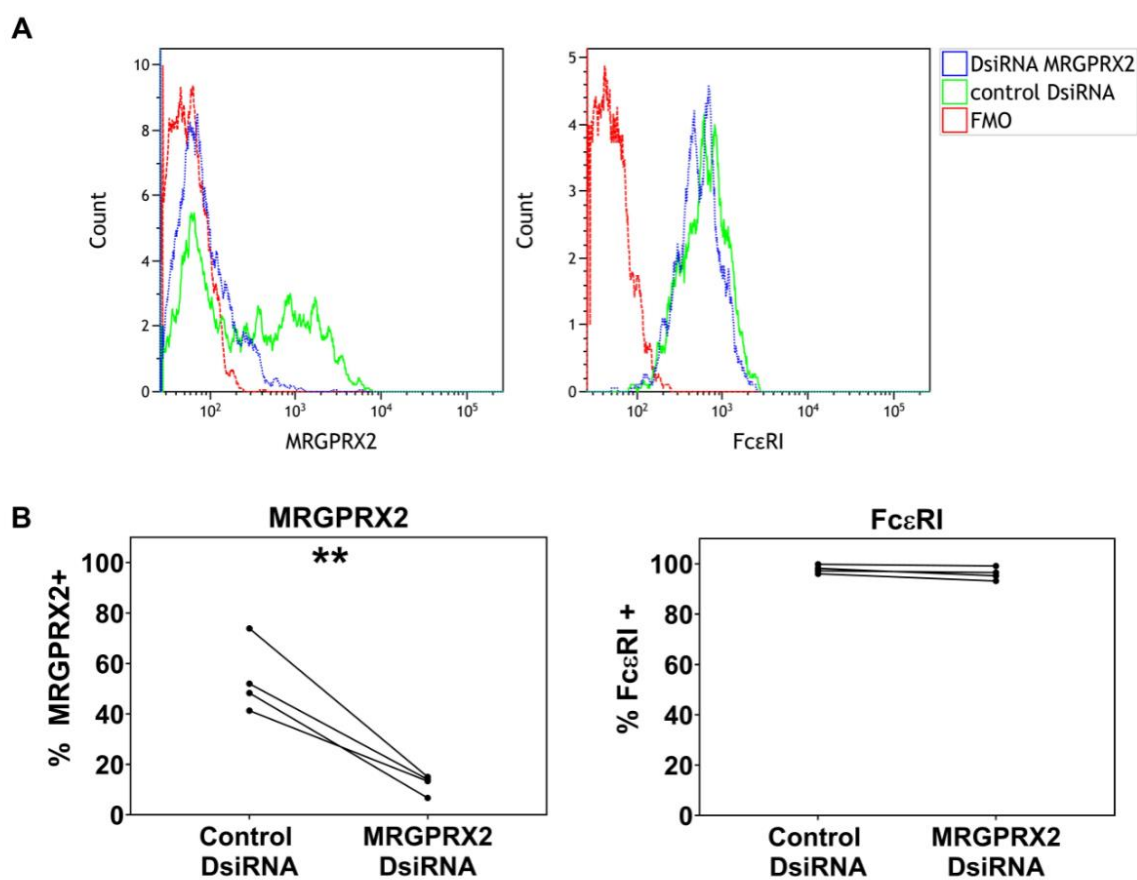
PBCMC samples were (after 50 sec) incubated with buffer, the natural ligand of MRGPRX2 substance P (74 μ M), amoxicillin (1370 μ M), morphine (500 μ M), moxifloxacin (571 μ M) or rocuronium (1640 μ M).

Figure 5. Representative plot of the transfection efficiency using enhanced green fluorescent protein (EGFP) mRNA.



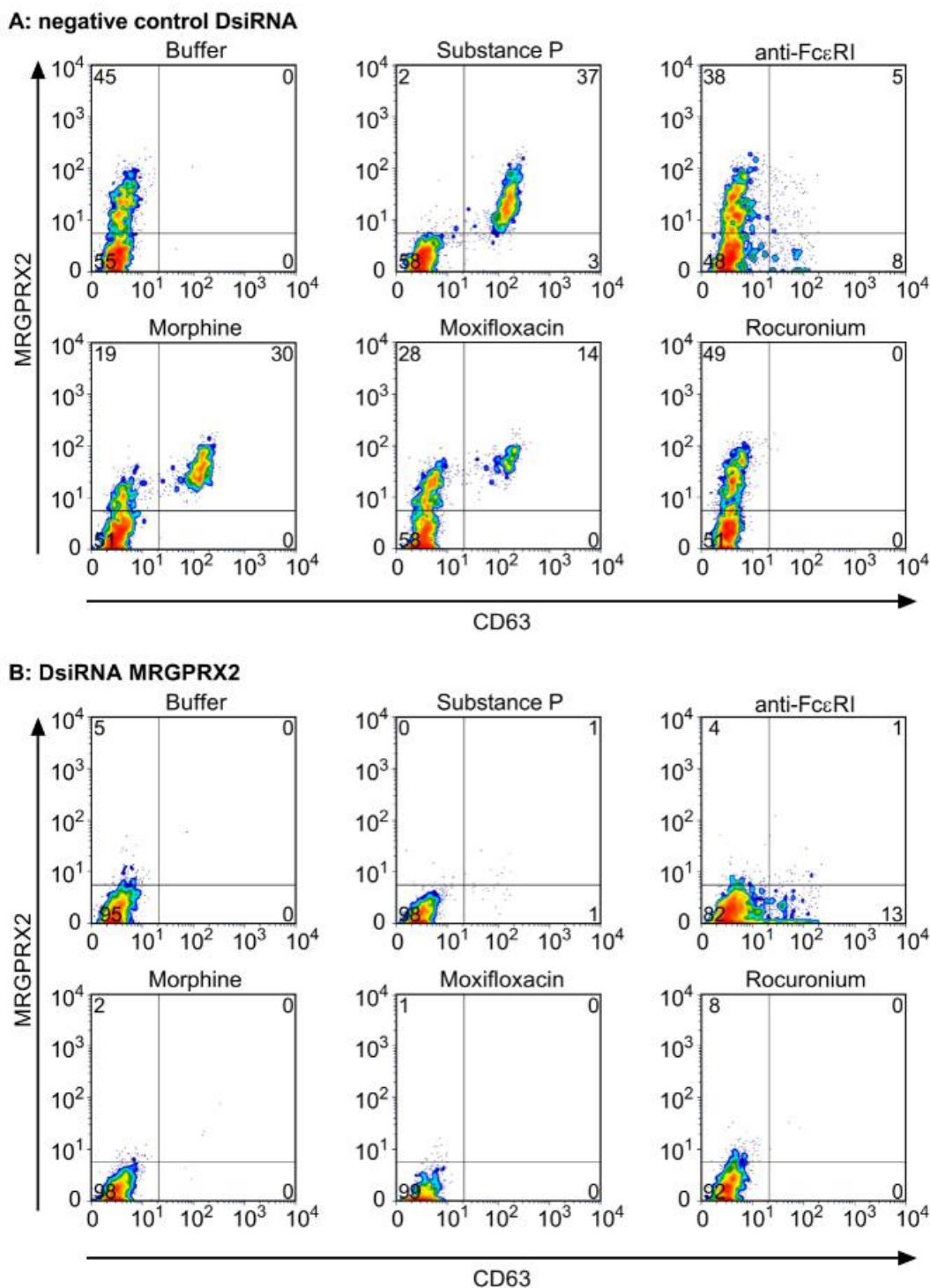
PBCMCs are electroperated with (green dot line) or without (red full line) EGFP mRNA as a surrogate for transfection efficiency of RNA electroporation. 24h after EGFP expression is analyzed using flow cytometry.

Figure 6. Silencing of the MRGPRX2-receptor after electroporation with anti-MRGPRX2 DsiRNA.



(A) Representative plots or (B) a comparison of the surface expression of MRGPRX2 or FcεRI between PBCMC electroporated with non-targeting DsiRNA or DsiRNA specific for MRGPRX2. In all experiments, $n=4$. $p < 0.01^{**}$.

Figure 7. Representative plots of CD63 up-regulation after silencing of MRGPRX2.



Cells were electroporated with a negative control(A) or target specific DsiRNA(B).

Thereafter, cells were incubated with buffer, substance P (74 μ M), anti-Fc ϵ RI(2.5 μ g/mL),

morphine (500 μ M), moxifloxacin (571 μ M)orrocuronium (1640 μ M).