

Supplementary information Table S3. Summary of cAMP inhibition assay of S1PR5.

Receptor	cAMP inhibition assay				Expression level (%WT)
	EC ₅₀ (nM)	Fold	EC ₅₀ (nM)	fold	
S1PR5	siponimod		SEW2871		
WT	4.18±0.24	1.0	120.4±89.3	1.0	100
R139 ^{3.56} A	ND	ND	-	-	62.64±15.9
R245 ^{6.29} A	ND	ND	-	-	65.89±14.1
R133 ^{3.50} A	ND	ND	-	-	74.55±15.5
R140 ^{1CL2} A	ND	ND	-	-	88.11±11.8
P142 ^{1CL2} L	3.02±1.80	0.72	-	-	81.72±16.2
N20D	ND	ND	-	-	74.43±18.5
N20A	ND	ND	-	-	56.75±13.3
N20Q	ND	ND	-	-	49.66±10.3
Q287 ^{7.37} R	17.9±11.3	4.3	42.86±22.02	0.35	77.54±12.5
S1PR5-DC345	5.21±1.96	1.2	-	-	135.9±16.7

ND means not detected. “-” means no measurement.