

Supplemental table 2. Structure Activity relationship (SAR) studies of compounds related to 5848633 and 5923764, the related hits from the STF3A cell based Wnt signaling inhibitor screen.

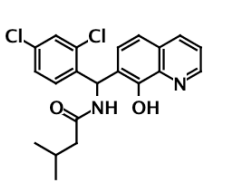
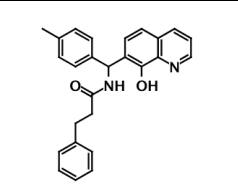
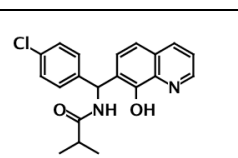
LD₅₀ values for the reporter cell line after 24 hours treatment with a subset of these compounds were also determined. These studies indicate that additions to the phenyl ring and a wide array of amide-linked groups have little or no detrimental effect on signal inhibition. By contrast, additions such as chloride or nitrosyl groups on C5 of the 8 hydroxyquinolone group or variations in linker length between the phenyl ring and the 8 hydroxyquinolone significantly reduced efficacy.

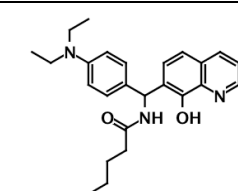
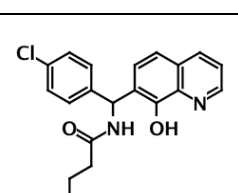
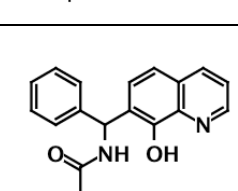
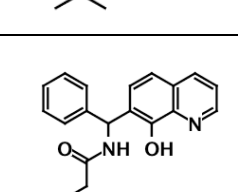
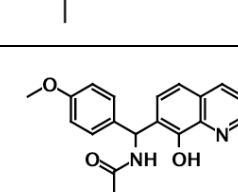
Therapeutic Index was calculated as IC₅₀ divided by LD₅₀ at 24 hours for STF3A cells.

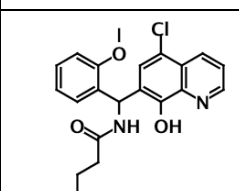
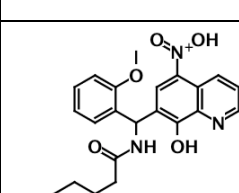
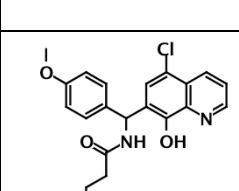
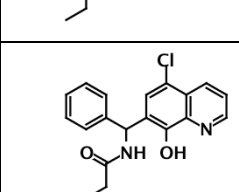
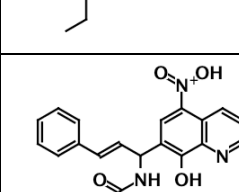
compound ID	structure	IC ₅₀ (μM); Therapeutic index
5923764		0.1 ND
5843979		0.2 17
5839152 (HQBA)		0.3 108
5837493		0.3 19
5845469		0.3 7
5848633		0.3 8
5842373		0.6 6

6385237		0.6 ND
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compound ID	structure	IC ₅₀ (μM); Therapeutic index
5837313		0.6 7
5925700		0.7 ND
5842334		0.7 6
6997141		0.7 ND
5847638		0.8 23

7151628		0.9 ND
7160597		1.4 ND
5925778		2.2 ND

compound ID	structure	IC ₅₀ (μM); Therapeutic index
6965682		4.5 ND
5923955		5.3 ND
5925622		6.3 ND
5924635		10.0 ND
5927674		16.0 ND

compound ID	structure	IC ₅₀ (μM); Therapeutic index
7171545		18.0 ND
6951252		20.5 ND
7125178		30.0 ND
7185515		63.0 ND
6997750		>56.0 ND