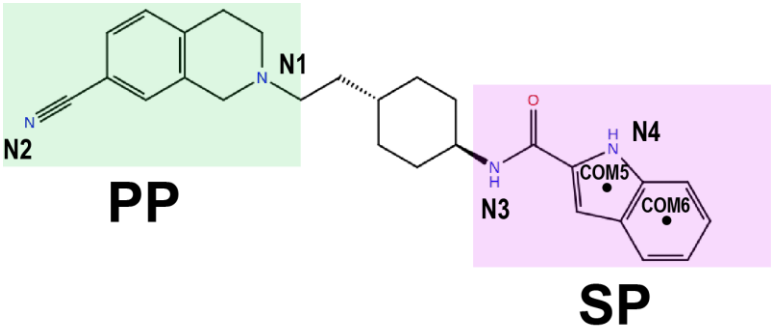
S1 Fig. Chemical structure of SB269652 and the input features for MSM analysis. The tetrahydroisoquinoline (PP) and indole-2-carboxamide moiety (SP) of SB269652 are shown in green and pink respectively. COM5 and COM6 refer to the centers of mass of the 5- and 6-member rings of the SP, respectively. The input features for MSM analysis include 12 distances from the nitrogen atoms of SB269652 to the indicated receptor Cβ atoms, 2 intramolecular distances of SB269652, and the projections of the vectors from COM5 or COM6 to N4 along the axis perpendicular to membrane (see Methods).



Index	Atom name	SB269652 atom
1	$Asp^{3.32}$ Cβ	N1
2	$Trp^{7.39}$ Cβ	N1
3	$Leu^{3.33}$ Cβ	N2
4	$Phe^{5.38}$ Cβ	N2
5	$Val^{5.39}$ Cβ	N2
6	$Ser^{5.42}$ Cβ	N2
7	$Ser^{5.43}$ Cβ	N2
8	$Ser^{5.46}$ Cβ	N2
9	$Val^{2.61}$ Cβ	N3
10	$Leu^{2.64}$ Cβ	N3
11	$Ser^{7.36}$ Cβ	N3
12	$Leu^{2.64}$ Cβ	N4
13	SB269652 N4	N3
14	SB269652 N4	O
15	SB269652 N4	COM5
16	SB269652 N4	COM6