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1. Dataset assembly

2.1 Dataset of D₁R-selective compounds (ChEMBL)

Table S1. D₁R-selective compounds extracted from ChEMBL with in vitro determined biological activities for all investigated DR subtypes.

ID	SMILES	K _i [nM] ^a			Ref
		D ₁ R	D ₂ R	D ₃ R	
SC1	<chem>O=C(C=1C([H])=C([H])C([F])=C([H])C=1[H])N([H])C([H])([H])C([H])([H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])[C@@]4([H])C=3C([H])=C([H])C([H])=C([H])C=3C([H])([H])C([H])([H])C4([H])[H]</chem>	70	3090	1570	[46]
SC2	<chem>N#CC=1C([H])=C([H])C2=C(C=1[H])C(=C([H])N2[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([Cl])=C([H])C=3[H])C([H])([H])C4([H])[H]</chem>	550	4300	6400	[47]
SC3	<chem>N#CC=2C([H])=C([H])C=1C([H])=C(N([H])C=1C=2[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([Cl])=C([Cl])C=3[H])C([H])([H])C4([H])[H]</chem>	880	2000	33000	
SC4	<chem>[Br]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])C2=N[C@]([H])([C@]([H])(O2)C([H])([H])[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3[H])C([H])([H])C4([H])[H]</chem>	480	21000	10000	[48]
SC5	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C=3C([H])=NN4C=3C([H])=C([H])C([H])=C4C#C[H]</chem>	810	3700	4600	[49]
SC6	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])C([H])([H])N(C([H])([H])[H])C([H])([H])[H]</chem>	343	2740	10000	
SC7	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])C3([H])[H]</chem>	113	1006	2923	
SC8	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])C([H])([H])N3C([H])([H])C([H])([H])C([H])([H])C3([H])[H]</chem>	33.7	1015	7804	[50]
SC9	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])C([H])([H])N3C([H])([H])C([H])([H])OC([H])([H])C3([H])[H]</chem>	201	10000	10000	
SC10	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])C([H])([H])N3C([H])([H])C([H])([H])C([H])([H])C3([H])[H]</chem>	379	10000	1818	
SC11	<chem>O=C2C1=C(C([H])=C([Cl])C([H])=C1[Cl])N([H])C(=O)[C@@]2(C=4C([H])=C([H])C([H])=C(OC([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3[H])C=4[H])C([H])([H])[H]</chem>	120	10000	10000	[51]
SC12	<chem>[H]C=1C([H])=C([H])C3=C(C=1[H])C([H])([H])C=2C([H])=C(OC([H])([H])[H])C([H])=C([H])C=2C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C3([H])[H])C([H])([H])[H]</chem>	137	1396	23903	[52]
SC13	<chem>[H]C=1C([H])=C([H])C3=C(C=1[H])C([H])([H])C2=C([H])C(OC([H])([H])[H])=C(OC([H])([H])[H])C([H])=C2C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C3([H])[H])C([H])([H])[H]</chem>	579	1028	14830	
SC14	<chem>O=C1C3=C(N=C(N1C([H])([H])[H])N2C([H])([H])C([H])([H])OC([H])([H])C2([H])[H])C(=C([H])N3[H])C([H])([H])N5C([H])([H])C([H]</chem>	520	1100	2700	[53]

ID	SMILES	K _i [nM] ^a			Ref
		D ₁ R	D ₂ R	D ₃ R	
	<chem>)]([H])N(C=4C([H])=C([H])C([H])=C([H])C=4[H])C([H])([H])C5([H])</chem>				
SC15	<chem>O=C(C=1C([H])=C([H])C([F])=C([H])C=1[H])C([H])([H])C([H])([H])C([H])([H])N5[C@@]3([H])[C@]7([H])[C@@]2([H])C([H])([H])[C@]6([H])[C@]4([H])[C@@]2([H])[C@@]3([H])[C@]4([H])[C@@]5(O[H])[C@]67[H]</chem>	209	1724	1958	[54]
SC16	<chem>[H]C=1C([H])=C([H])C2=C(C=1[H])C4=C(N2[H])C([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3C([H])([H])C([H])([H])N([C@@]([H])C4([H])[H])C([H])([H])[H]</chem>	640	10000	10000	[55]
SC17	<chem>[H]C=1C([H])=C([H])C2=C(C=1[H])C4=C(N2[H])C([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3C([H])([H])N([C@@]([H])C4([H])[H])C([H])([H])O[H]C([H])([H])[H]</chem>	114	2923	4676	
SC18	<chem>O=C2C=1C([H])=C([H])C([H])=C([H])C=1C([H])([H])[C@@]2([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([Cl])=C([H])C=3[H])C([H])([H])C4([H])[H]</chem>	389.05	10000	4534	[56]
SC19	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C=4OC=3C([H])=C([H])C([H])=C([H])C=3C=4[H]</chem>	274	2507	1462	
SC20	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])[C@@]3([H])C=2C([H])=C(O[H])C([H])=C([H])C=2C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])[H]</chem>	207	1000	1000	
SC21	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])[C@@]3([H])C=2C([H])=C(O[H])C(=C([H])C=2C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])[H])N([H])[H]</chem>	334	1000	1000	[57]
SC22	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])[C@]3([H])C2=C([H])C(O[H])=C(C([H])=C2C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])[H])C4=C([H])C([H])=C([H])C(=C4[H])C([H])([H])[H]</chem>	216	1000	1000	
SC23	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])[C@@]3([H])C=2C([H])=C(O[H])C(=C([H])C=2C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])[H])C=4C([H])=C([H])C(O[H])=C([H])C=4[H]</chem>	349	1000	1000	
SC24	<chem>[H]/C4=C3\C1=C2/C(=C(/[H])C(OC([H])([H])[H])=C1OC([H])([H])[H])C([H])([H])C([H])([H])N([C@@]2([H])C([H])([H])C3=C([H])C(O[H])=C4OC([H])([H])[H])C([H])([H])[H]</chem>	373	inactive	>950	[58]
SC25	<chem>[H]/C1=C(\[H])C(OC([H])([H])[H])=C(OC([H])([H])[H])C4=C1C([H])([H])[C@@]3([H])C2=C([H])C(OC([H])([H])[H])=C(OC([H])([H])[H])C([H])=C2C([H])([H])C([H])([H])N3C4([H])[H]</chem>	153	1125	1371	[59]
SC26	<chem>[H]/C3=C(\[H])C(O/C2=N/C([H])=C([H])C=1OC([H])=C([H])C=12)=C([H])C(=C3/C5=C(/N=C(/[H])C4=NC([H])=C([H])N45)C([H])([H])[H])C([H])([H])[H]</chem>	110	8300	7900	
SC27	<chem>[F]C([F])([F])C=3C([H])=C(O/C2=N/C([H])=C([H])C=1C([H])=C([H])[S]C=12)C([H])=C([H])C=3/C5=C(\[H])C([H])=C([H])C4=NC([H])=C([H])N45</chem>	130	8700	6200	[60]
SC28	<chem>[F]C([F])([F])C=3C([H])=C(O/C2=N/C([H])=C([H])C=1OC([H])=C([H])C=12)C([H])=C([H])C=3/C5=C(\[H])C([H])=C([H])C4=NC([H])=C([H])N45</chem>	150	7600	10000	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC40	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C=4N=C3C([H])=C([H])C([H])=C(N3C=4[H])N5/N=C(/[H])C([H])=C5[H]</chem>	27000	590	1000	
SC41	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C=4N=C3C([H])=C([H])C([H])=C([H])N3C=4[H]</chem>	20000	990	3900	
SC42	<chem>[Cl]/C2=N/N1C([H])=C(N=C1C([H])=C2[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	7700	96	1400	
SC43	<chem>[F]C([H])([H])C([H])([H])OC=1C([H])=C([H])C([H])=C(C=1[H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([Cl])=C([H])C=2[H])C([H])([H])C3([H])[H]</chem>	5100	590	1100	
SC44	<chem>[F]C([H])([H])C([H])([H])OC=1C([H])=C([H])C=C([H])C=1OC([H])([H])[H]C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])[H])C([H])([H])C3([H])[H]</chem>	5400	310	1100	[67]
SC45	<chem>[F]C([H])([H])C([H])([H])O/C1=C(\[H])C(OC([H])([H])[H])=C(O C([H])([H])[H])C([H])=C1C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([Cl])=C([H])C=2[H])C([H])([H])C3([H])[H]</chem>	3500	760	3900	
SC46	<chem>O=C1N([H])C([H])([H])C([H])([H])N1C([H])([H])C([H])([H])N6C([H])([H])C([H])([H])C([H])(C2=C([H])N(C=3C([H])=C([H])C=C([H])C2=3)C=4/N=C(/[H])N(N=4)C([H])([H])[H])C=5C([H])=C([H])C([F])=C([H])C=5[H])C([H])([H])C6([H])[H]</chem>	4100	660	2900	
SC47	<chem>O=C1N(C([H])([H])C([H])([H])[C@]1([H])C([H])([H])C([H])([H])N6C([H])([H])C([H])([H])[C@@]([H])(C2=C([H])N(C=3C([H])=C([H])C=C([H])C2=3)C=4/N=C(/[H])N(N=4)C([H])([H])[H])C=5C([H])=C([H])C([F])=C([H])C=5[H])C([H])([H])C6([H])[H])C([H])([H])[H]</chem>	2600	190	1400	[68]
SC48	<chem>[Br]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])C2=N/C=C(/[H])[S]2)C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3[H])C([H])([H])C4([H])[H]</chem>	6400	190	20000	[53]
SC49	<chem>[H]C=3C([H])=C2C1=C(/N=C(/O[H])N1C([H])([H])[C@]([H])(N(C([H])([H])[H])C([H])([H])[H])C2([H])[H])C=3[H]</chem>	5000	20	10000	[69]
SC50	<chem>[F]C([H])([H])C([H])([H])C([H])([H])OC=1C([H])=C([H])C=C([H])C=1[H])N2N=N/C=C2/[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	38000	340	2800	
SC51	<chem>[F]C([H])([H])C([H])([H])OC([H])([H])C([H])([H])C([H])([H])OC=1C([H])=C([H])C=C([H])C=1[H])N2N=N/C=C2/[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	3600	440	2900	[70]
SC52	<chem>O=C(C=1C([H])=C([H])C([F])=C([H])C=1[H])C([H])([H])C([H])([H])C(O)N3C([H])([H])C([H])([H])C(O[H])(C=2C([H])=C([H])C([Cl])=C([H])C=2[H])C([H])([H])C3([H])[H]</chem>	19000	860	1700	[71]
SC53	<chem>[H]C=1C([H])=C([H])C2=C(C=1[H])C([H])([H])C([H])(C2([H])[H])C([H])([H])N5C([H])([H])C([H])([H])N(C=4C([H])=C([H])C=3OC([H])([H])C([H])([H])OC=3C=4[H])C([H])([H])C5([H])[H]</chem>	3000	738	2840	[72]

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC54	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C3=C([H])N([H])C=4/N=C(/[H])C([H])=C([H])C3=4</chem>	10000	960	2310	

^aAll K_i values retrieved from ChEMBL were determined in vitro utilizing radio-ligand binding assays.

2.3 Dataset of D₂-like-selective compounds (ChEMBL)

Table S3. D₂-like-selective compounds extracted from ChEMBL with in vitro determined biological activities for all investigated DR subtypes.

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC55	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C2=C([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C#CC=3C([H])=C([H])C([H])=NC=3[H]</chem>	5004	3.3	20	
SC56	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C2=C([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C#CC=3C([H])=C([H])C(=C([H])C=3[H])N([H])[H]</chem>	10000	0.67	24	[73]
SC57	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C2=C([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C#CC=3C([H])=C([H])C(=NC=3[H])N([H])[H]</chem>	10000	19	7	
SC58	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C2=C([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C#C/C3=N/C([H])=C(C([H])=C3[H])N([H])[H]</chem>	10000	1.9	2.6	
SC59	<chem>O=C(C=3C([H])=C([H])C=2C=1C([H])=C([H])C([H])=C([H])C=1C([H])([H])C=2C=3[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C([H])([H])N(C=4C([H])=C([H])C([H])=C([Cl])C=4[Cl])C([H])([H])C5([H])[H]</chem>	10000	217	1.4	[74]
SC60	<chem>O=C(C=3C([H])=C([H])C=2C=1C([H])=C([H])C([H])=C([H])C=1C=2C=3[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C([H])([H])N(C=4C([H])=C([H])C([H])=C([Cl])C=4[Cl])C([H])([H])C5([H])[H]</chem>	10000	262	0.9	
SC61	<chem>[Cl]C=2C([H])=C([H])C=1OC([H])([H])[C@@]([H])(OC=1C=2[H])C([H])([H])N([H])C([H])([H])[C@@]4([H])C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	1000	13.2	1.6	[75]
SC62	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])[C@@]3(O[H])C([H])([H])[C@@]2([H])N([C@]([H])(C([H])([H])C2([H])[H])C3([H])[H])C([H])([H])C4=C([H])N([H])C=5C([H])=C([H])C([H])=C([H])C4=5</chem>	1220	33.4	15.5	
SC63	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])[C@@]3(O[H])C([H])([H])[C@@]2([H])N([C@]([H])(C([H])([H])C2([H])[H])C3([H])[H])C([H])([H])C4=C([H])[S]C=5C([H])=C([H])C([H])=C([H])C4=5</chem>	3000	12.9	3.62	[76]
SC64	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])[C@@]3(O[H])C([H])([H])[C@@]2([H])N([C@]([H])(C([H])([H])C2([H])[H])C3([H])[H])C([H])([H])C4=C([H])OC=5C([H])=C([H])C([H])=C([H])C4=5</chem>	3930	1.06	0.71	
SC65	<chem>O=C1C([H])=NN(C(=O)N1C([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(/C3=C([H])C([H])=C([H])C=2C([H])=C([H])C(OC([H])([H])[H])=C([H])C=23)C([H])([H])C4([H])[H]</chem>	7396	22.01	62.8	[77]

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC66	<chem>O=C(C=1C([H])=C3C([H])=C([H])C=1C([H])([H])C([H])([H])C=2C([H])=C([H])C(=C([H])C=2[H])C([H])([H])C3([H])[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C([H])([H])N(C=4C([H])=C([H])C([H])=C([Cl])C=4[Cl])C([H])([H])C5([H])[H])</chem>	2800	220	3.6	[78]
SC67	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H])</chem>	2000	36.5	0.92	
SC68	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H])</chem>	2100	44.8	0.81	
SC69	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H])</chem>	4630	502	1.39	
SC70	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H])</chem>	10000	433	1.12	
SC71	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H])</chem>	10000	715	16.6	
SC72	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H])</chem>	10000	249	1.4	[79]
SC73	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])C([H])([H])C([H])([H])C4([H])[H])</chem>	4960	47.1	62.1	
SC74	<chem>O=C(/C2=C([H])C=1C([H])=C([F])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H])</chem>	10000	293	1.61	
SC75	<chem>O=C(/C2=C([H])C=1C([H])=C([F])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H])</chem>	5060	244	2.41	
SC76	<chem>O=C(/C2=C([H])C=1C([H])=C([I])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H])</chem>	10000	520	3.6	
SC77	<chem>O=C(/C2=C([H])C=1C([H])=C(OC([H])([H])[H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H])</chem>	10000	489	1.2	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC78	<chem>O=C(/C2=C(\ [H])C=1C([H])=C(OC([H])([H])[H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C4([H])[H]</chem>	10000	390	2.32	
SC79	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	10000	622	6.11	
SC80	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C4([H])[H]</chem>	10000	507	7.51	
SC81	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])[C@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	10000	581	7.71	
SC82	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C4([H])[H]</chem>	3690	430	7.5	
SC83	<chem>O=C(C=2[S]C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C4([H])[H]</chem>	6650	337	4.6	
SC84	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@]([H])(O[H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	10000	28.4	0.26	
SC85	<chem>O=C(C=2[S]C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])=C([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	1730	149	1.11	
SC86	<chem>O=C(C=2OC=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])=C([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	1460	76.5	2.11	
SC87	<chem>O=C2C=1C([H])=C([H])C([H])=C([H])C=1C([H])([H])C([H])([H])C([H])([H])N2C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	1737.8	36.3078	14.45	
SC88	<chem>O=C2C=1C([H])=C([H])C([H])=C([H])C=1C([H])([H])C([H])([H])C([H])([H])N2C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C4([H])[H]</chem>	2691.53	3.63078	1.585	[80]
SC89	<chem>O=C(C=1C([H])=C3C([H])=C([H])C=1C([H])([H])C([H])([H])C=2C([H])=C([H])C=C([H])C=2[H])C([H])([H])C3([H])[H]C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C([H])([H])C(O[H])(C=4C([H])=C([H])C([Cl])=C([H])C=4[H])C([H])([H])C5([H])[H]</chem>	9800	21	17	[81]

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC90	<chem>O=C(C=2[S]C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N6C([H])([H])C([H])([H])N(C=3C([H])=C5C([H])=C([H])C=3C([H])([H])C([H])([H])C=4C([H])=C([H])C=C([H])C=4[H])C([H])([H])C5([H])[H]C([H])([H])C6([H])[H]</chem>	4000	83	1.6	
SC91	<chem>O=C(C=2[S]C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C(C([H])=C([H])C=3C([H])([H])[H])C([H])([H])[H]C([H])([H])C4([H])[H]</chem>	2400	84	2.5	
SC92	<chem>O=C(C=1C([H])=C(C([H])=C([H])C=1C([H])([H])[H])C([H])([H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])[H]C([H])([H])C3([H])[H]</chem>	2100	22	4.7	
SC93	<chem>O=C(C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])[H]C([H])([H])C3([H])[H]</chem>	1100	27	0.34	[82]
SC94	<chem>[F]C([H])([H])C([H])([H])C([H])([H])OC=3C([H])=C2C1=C(C([H])=C([H])C(O[H])=C1O[H])C([H])([H])[C@]4([H])C2=C(C=3[H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])C([H])([H])[H]</chem>	5000	28	430	
SC95	<chem>[F]C([H])([H])C([H])([H])C([H])([H])OC=3C([H])=C2C1=C(C([H])=C([H])C([H])=C1O[H])C([H])([H])[C@]4([H])C2=C(C=3[H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])C([H])([H])[H]</chem>	5000	0.54	100	
SC96	<chem>[F]C([H])([H])C([H])([H])OC=3C([H])=C2C1=C(C([H])=C([H])C([H])=C1O[H])C([H])([H])[C@]4([H])C2=C(C=3[H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])C([H])([H])[H]</chem>	5000	3.5	410	[69]
SC97	<chem>[F]C([H])([H])C([H])([H])C([H])([H])OC=3C([H])=C2C1=C(C([H])=C([H])C([H])=C1O[H])C([H])([H])[C@]4([H])C2=C(C=3[H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])[H]</chem>	5000	3.2	240	
SC98	<chem>[F]C([H])([H])C([H])([H])OC=3C([H])=C2C1=C(C([H])=C([H])C([H])=C1O[H])C([H])([H])[C@]4([H])C2=C(C=3[H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])[H]</chem>	5000	0.83	550	
SC99	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N3N=N/C(=C3/[H])C([H])([H])O/C4=C(\[H])C([H])=C([H])C7=C4/C5=C(\[H])C([H])=C([H])C6=C5[C@@([H])(N(C([H])([H])C6([H])[H])C([H])([H])C([H])([H])C([H])([H])[H])C7([H])[H]</chem>	2700	661	2.67	
SC100	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C=3N=NN(C=3[H])C([H])([H])C([H])([H])C([H])([H])O/C4=C(\[H])C([H])=C([H])C7=C4/C5=C(\[H])C([H])=C([H])C6=C5[C@@([H])(N(C([H])([H])C6([H])[H])C([H])([H])C([H])([H])C([H])([H])[H])C7([H])[H]</chem>	1900	206	8.14	[83]
SC101	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])O/C3=C(\[H])C([H])=C([H])C6=C3/C4</chem>	1174	174	189	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
	<chem>=C(\[H])C([H])=C([H])C5=C4[C@@]([H])(N(C([H])([H])C5([H])</chem> <chem>[H])C([H])([H])C([H])([H])C([H])([H])[H])C6([H])[H]</chem>				
SC102	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])(</chem> <chem>[H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H]</chem> <chem>)C([H])([H])C([H])([H])O/C3=C(\[H])C([H])=C([H])C6=C3/C4</chem> <chem>=C(\[H])C([H])=C([H])C5=C4[C@]([H])(N(C([H])([H])C5([H])[</chem> <chem>H])C([H])([H])C([H])([H])C([H])([H])[H])C6([H])[H]</chem>	1307	141	23.4	
SC103	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])[C@]2([H])N(C([H])([H]</chem> <chem>)[C@]4([H])[C@@]2([H])[C@@]([H])(N3C([H])([H])C([H])([H])</chem> <chem>C([H])([H])C3([H])[H])[C@]4([H])C([H])([H])O[H])C([H])([H])</chem> <chem>C=5C([H])=C([H])C([H])=C([H])C=5[H]</chem>	22000	250	280	[84]
SC104	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])</chem> <chem>N(C([H])([H])C([H])([H])C=2C([H])=C([H])C([H])=C(O[H])C=2</chem> <chem>[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	10000	64.4	21.8	
SC105	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])</chem> <chem>N(C([H])([H])C2([H])C([H])([H])C2([H])[H])C([H])([H])C([H])</chem> <chem>([H])C3=C([H])C([H])=C([H])C(O[H])=C3[H]</chem>	10000	118	168	[85]
SC106	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])</chem> <chem>N(C([H])([H])C2([H])C([H])([H])C([H])([H])C2([H])[H])C([H])</chem> <chem>([H])C([H])([H])C3=C([H])C([H])=C([H])C(O[H])=C3[H]</chem>	10000	450	282	
SC107	<chem>O=C1C([H])=NN(C(=O)N1C([H])([H])[H])C([H])([H])C([H])([</chem> <chem>H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([</chem> <chem>H])=C([H])C([H])=C([H])C=2OC([H])([H])C([H])([H])[H])C([H]</chem> <chem>)([H])C3([H])[H]</chem>	10000	37.1	22	[86]
SC108	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])C([H])([H])[H])=C(C=1[H]</chem> <chem>)N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H]</chem> <chem>)C=3[S]C([H])=C([H])C=3[H]</chem>	10000	162	112	
SC109	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])C([H])([H])[H])=C(C=1[H]</chem> <chem>)N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H]</chem> <chem>)C=3C([H])=C([H])C([H])=NC=3[H]</chem>	10000	255	42	
SC110	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([</chem> <chem>H])N(C([H])([H])C2([H])[H])C([H])([H])C=3C([H])=C([H])C([</chem> <chem>H])=NC=3[H]</chem>	10000	327	41	
SC111	<chem>[H]C=1C([H])=C([H])C2=C(C=1[H])C([H])=C([H])C([H])=C2C([</chem> <chem>H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=</chem> <chem>C([H])C=3OC([H])([H])C([H])([H])[H])C([H])([H])C4([H])[H]</chem>	10000	264	59	
SC112	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([</chem> <chem>H])N(C([H])([H])C2([H])[H])C([H])([H])C4=C([H])C([H])=C([</chem> <chem>H])C=3C([H])=C([H])C([H])=C([H])C=34</chem>	10000	351	92	[87]
SC113	<chem>[H]C=2C([H])=C([H])C=1[S]C([H])=C(C=1C=2[H])C([H])([H])N</chem> <chem>4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3</chem> <chem>OC([H])([H])C([H])([H])[H])C([H])([H])C4([H])[H]</chem>	10000	219	45.1	
SC114	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([</chem> <chem>H])N(C([H])([H])C2([H])[H])C([H])([H])C3=C([H])[S]C=4C([H]</chem> <chem>)=C([H])C([H])=C([H])C3=4</chem>	10000	383	52.5	
SC115	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])N3C([H])([</chem> <chem>H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([</chem> <chem>H])C([H])([H])[H])C([H])([H])C3([H])[H]</chem>	10000	217	42.7	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC116	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3[H]</chem>	10000	242	41	
SC117	<chem>[H]C=1C([H])=C([H])C2=C(C=1[H])C(=C([H])N2[H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])C([H])([H])[H])C([H])([H])C4([H])[H]</chem>	10000	473	103	
SC118	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C3=C([H])N([H])C=4C([H])=C([H])C([H])=C([H])C3=4</chem>	10000	502	123	
SC119	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N3N=N/C(=C3/[H])C=4C([H])=C([H])C(=C([H])C=4[H])N([H])[H]</chem>	38430	198.8	2.7	[88]
SC120	<chem>[I]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])N([H])C([H])([H])[C@]4([H])OC=3C([H])=C(O[H])C([H])=C([H])C=3C([H])([H])C4([H])[H]</chem>	3960	80.5	23	[89]
SC121	<chem>[H]/C1=C(\[H])C(O[H])=C([H])C=2O[C@]([H])(C([H])([H])C([H])([H])C1=2)C([H])([H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	10900	8.19	547	
SC122	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3[S]4</chem>	1897	219	31	[56]
SC123	<chem>O=C(N([H])[C@@]1([H])C([H])([H])C([H])([H])[C@]([H])(C([H])([H])C1([H])[H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([Cl])C=2[Cl])C([H])([H])C3([H])[H])N(C([H])([H])[H])C([H])([H])[H]</chem>	2100	0.47	0.27	
SC124	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])O/C4=C(\[H])C3=C([H])C([H])=NN3C([H])=C4[H]</chem>	2900	34	7	
SC125	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])OC=4/C3=C(\[H])C([H])=NN3C([H])=C([H])C=4[H]</chem>	1800	25	10	
SC126	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])OC=4C([H])=C([H])C3=C([H])C([H])=NN3C=4[H]</chem>	1200	4.2	2.3	[90]
SC127	<chem>[Cl]C=1C([H])=C([H])C([H])=C(C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])O/C3=N/N4/C(=C3/[H])C([H])=C([H])C([H])=C4[H]</chem>	1100	9.4	3.5	
SC128	<chem>O=C([H])C=1C([H])=NN2C=1C([H])=C([H])C([H])=C2OC([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	1600	51	6.4	
SC129	<chem>[Cl]/C1=C(\[H])C([H])=C([H])C(=C1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	4300	1.8	2.3	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
	<chem>)]C([H])([H])O/C4=C(\ [H])C3=C(C([H])=NN3C([H])=C4[H])C([H])=NO[H]</chem>				
SC130	<chem>N#C/C1=N/N4/C(=C1/[H])C([H])=C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([Cl])C=2[Cl])C([H])([H])C3([H])[H])C([H])=C4[H]</chem>	3500	8.9	1.9	
SC131	<chem>[F]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N2[C@@]4([H])C([H])([H])C([H])([H])[C@]2([H])C([H])([H])[C@@](O[H])(C=3C([H])=C([H])C([Cl])=C([H])C=3[H])C4([H])[H]</chem>	10000	33	9.7	
SC132	<chem>[F]C=4C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])N1[C@@]3([H])C([H])([H])C([H])([H])[C@]1([H])C([H])([H])[C@@](O[H])(C=2C([H])=C([H])C([Cl])=C([H])C=2[H])C3([H])[H])=C([H])C=4[H]</chem>	10000	22	4.1	
SC133	<chem>[F]C=1C([H])=C([H])C([H])=C([H])C=1[S]C([H])([H])C([H])([H])C([H])([H])N2[C@@]4([H])C([H])([H])C([H])([H])[C@]2([H])C([H])([H])[C@@](O[H])(C=3C([H])=C([H])C([Cl])=C([H])C=3[H])C4([H])[H]</chem>	4093	4.8	1.3	[91]
SC134	<chem>O=C(C=1C([H])=C([H])C([H])=C([H])C=1OC([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C3C=2C([H])=C([H])N3[S](=O)(=O)C=4C([H])=C([H])C([H])=C([H])C=4[H])C([H])([H])C5([H])[H])N([H])[H]</chem>	1000	6.3	0.1	
SC135	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C=2C([H])=C([H])C([H])=C([H])C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@]3([H])C([H])([H])C([H])=C(C#C[H])C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	13000	9.4	1	
SC136	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C=2C([H])=C([H])C([H])=C([H])C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@@]3([H])C([H])([H])C([H])=C(C#C[H])C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	18000	14	4.3	
SC137	<chem>O=C(C=2C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N=NN(C=1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@]3([H])C([H])([H])C([H])=C(C#C[H])OC3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	2600	1.6	0.63	
SC138	<chem>O=C(C=2C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N=NN(C=1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@@]3([H])C([H])([H])C([H])=C(C#C[H])OC3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	3900	4.9	2.2	[92]
SC139	<chem>O=C2C([H])=C([H])C=1/C(=C(/[H])C([H])=C(O[H])C=1N2[H])C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([H])C(=O)C=4C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])C=3N=NN(C=3[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])=C(OC([H])([H])[H])C=4[H]</chem>	4600	24	1.6	
SC140	<chem>O=C(C=2C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N=NN(C=1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@]3([H])C([H])([H])C([H])=C(C#C[H])OC3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	23000	18	1.2	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
	<chem>=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3[H])=C(OC([H])([H])[H])C=4[H]</chem>				
SC153	<chem>O=C(N([H])C2=C([H])C1=C([H])C([H])=NN1C([H])=C2[H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	1700	46	97	
SC154	<chem>O=C([H])C=1C([H])=NN4C=1C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])N(C=2C([H])=C([H])C([H])=C([H])C=2OC([H])([H])[H])C([H])([H])C3([H])[H])=C4[H]</chem>	1400	7.8	17	
SC155	<chem>O=C(N([H])C2=C([H])C1=C(C([H])=NN1C([H])=C2[H])C([H])=NO[H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	2900	0.57	2.2	
SC156	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])OC=4C([H])=C([H])C3=C(C([H])=NN3C=4[H])C([H])=NO[H]</chem>	1000	4.2	2.3	
SC157	<chem>O=C([H])C=1C([H])=NN4C=1C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@]3([H])C([H])([H])C=2C([H])=C([H])C([H])=C(O[H])C=2C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H])=C4[H]</chem>	1400	2.4	1.6	
SC158	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])O/C4=C([H])C3=C([H])C([H])=NN3C([H])=C4[H]</chem>	2800	18	20	[97]
SC159	<chem>O=C([H])C=1C([H])=NN2C=1C([H])=C(C([H])=C2[H])N([H])C(=O)C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	1200	1.4	70	
SC160	<chem>O=C([H])C=1C([H])=NN4C=1C([H])=C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([C@]3([H])C([H])([H])C=2C([H])=C([H])C([H])=C(O[H])C=2C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H])C([H])=C4[H]</chem>	1300	2.3	1.1	
SC161	<chem>[H]/C1=C([H])C4=C(C(O[H])=C1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])O/C3=C([H])C2=C([H])C([H])=NN2C([H])=C3[H])C4([H])[H]</chem>	3700	96	13	
SC162	<chem>[H]/C1=C([H])C4=C(C(O[H])=C1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])OC=3C([H])=C([H])C2=C([H])C([H])=NN2C=3[H])C4([H])[H]</chem>	2100	0.63	0.71	
SC163	<chem>[H]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])OC=4C([H])=C([H])C3=C([H])C([H])=NN3C=4[H]</chem>	1100	1.8	5	
SC164	<chem>[H]/C1=C([H])C4=C(C(O[H])=C1[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])O/C3=C([H])C2=C([H])C([H])=NN2C([H])=C3[H])C4([H])[H]</chem>	1800	2.2	0.59	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
	<chem>C([H])([H])C([H])([H])C([H])([H])O/C5=C([H])C4=C(C([H])=NN4C([H])=C5[H])C([H])=NO[H]</chem>				
SC188	<chem>O=C1N([H])C=2C(OC1([H])[H])=C(C([H])=C([H])C=2O([H])N3C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	20000	120	5.9	
SC189	<chem>O=C2C([H])=C([H])C=1/C(=C([H])C([H])=C(O[H])C=1N2([H])N3C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])[H]</chem>	20000	780	380	
SC190	<chem>O=C([H])C=1C([H])=NN5C=1C([H])=C(OC([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=2C([H])=C([H])C(O[H])=C3OC([H])([H])C([H])([H])OC=23C([H])([H])C4([H])[H])C([H])=C5[H]</chem>	11000	5.7	24	
SC191	<chem>O=C1N([H])C=2C(OC1([H])[H])=C(C([H])=C([H])C=2O([H])N3C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])[H]</chem>	20000	630	48	
SC192	<chem>O=C2C([H])=C([H])C=1/C(=C([H])C([H])=C(O[H])C=1N2([H])N3C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])O/C5=C([H])C4=C(C([H])=NN4C([H])=C5[H])C(=O)[H]</chem>	15000	1	2.3	
SC193	<chem>O=C/C1=N/N2/C(=C1/[H])C([H])=C([H])C([H])=C2([H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H])C([H])([H])C4([H])[H]</chem>	1600	270	3.5	[100]
SC194	<chem>O=C/C1=N/N2/C(=C1/[H])C([H])=C([H])C([H])=C2([H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N/C3=N/C(=N/C(=C3/[H])C([F])([F])[F])C(C([H])([H])C([H])([H])C([H])[H])C([H])([H])C4([H])[H]</chem>	7500	430	1.2	
SC195	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])N(C([H])([H])[H])C([H])([H])[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])[C@]([H])(C=2C([H])=C([H])C([H])=C(O[H])C=2[H])C3([H])[H]</chem>	10000	345	27	[101]
SC196	<chem>[F]C([F])([F])C=1C([H])=C(C([H])=C([H])C=1[Cl])C2(O[H])C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C=3C([H])=C([H])C(OC([H])([H])[H])=C([H])C=3[H])C=4C([H])=C([H])C(OC([H])([H])[H])=C([H])C=4[H]</chem>	10000	417	437	[102]
SC197	<chem>N#CC=1C([H])=C([H])C([H])=C(C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N([H])C(=O)C=3C([H])=C([H])C(=C([H])C=3[H])C=4C([H])=C([H])[S]C=4[H]</chem>	10000	743	0.5	[103]
SC198	<chem>[F]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])[C@@]2([H])C([H])([H])[C@]2([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])[S]/C4=N/N=C(/C=3C([H])=C([H])C([H])=C([H])C=3[H])N4C([H])([H])[H])C([H])(C([H])([H])[H])C([H])([H])[H]</chem>	5000	512.9	104.7	[104]
SC199	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([Cl])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])[H]</chem>	1981	408	23.4	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC200	<chem>[Cl]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])[C@]2([H])C([H])([H])[C@]2([H])C([H])([H])N(C([H])([H])C([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[S]/C4=N/N=C(/C=3C([H])=C([H])C([H])=C([H])C=3[H])N4C([H])([H])[H]</chem>	1509	238	31.6	
SC201	<chem>[Cl]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])[C@@]2([H])C([H])([H])[C@@]2([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[S]/C4=N/N=C(/C=3C([H])=C([H])C([H])=C([H])C=3[H])N4C([H])([H])[H]</chem>	1918	233	36.7	
SC202	<chem>[F]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])[C@@]2([H])C([H])([H])[C@]2([H])C([H])([H])N(C([H])([H])C([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[S]/C4=N/N=C(/C=3C([H])=C([H])C([H])=C([H])C=3[H])N4C([H])([H])[H]</chem>	1724.6	275.4	44	
SC203	<chem>[H]C2([H])C=1N=C([S]C=1C([H])([H])[C@@]([H])(N([H])C([H])([H])C([H])([H])C([H])([H])[H])C2([H])[H])N([H])[H]</chem>	10000	743.5	0.9	[105]
SC204	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]3([H])C([H])([H])[C@]3([H])C([H])([H])N4C([H])([H])[C@@]([H])(OC([H])([H])[C@]4([H])C([H])([H])[H])C=5C([H])=C([H])C(=NC=5[H])N([H])[H]</chem>	5630	87.8	1.85	
SC205	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])[C@@]3([H])C([H])([H])C([H])([H])[C@]([H])(C([H])([H])C3([H])[H])C([H])([H])C([H])([H])N4C([H])([H])[C@@]([H])(OC([H])([H])[C@]4([H])C([H])([H])[H])C=5C([H])=C([H])C(=NC=5[H])N([H])[H]</chem>	100000	104	21.6	[106]
SC206	<chem>[H]/C1=C(\ [H])C(O[H])=C([H])C2=C1OC([H])([H])[C@]3([H])N(C([H])([H])C([H])([H])O[C@]23[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	100000	28	1.67	

^aAll K_i values retrieved from ChEMBL were determined in vitro utilizing radio-ligand binding assays.

2.4 Dataset of D₃R-selective compounds (ChEMBL)

Table S4. D₃R-selective compounds extracted from ChEMBL with in vitro determined biological activities for all investigated DR subtypes.

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC207	<chem>O=C(C=1C([H])=C([Cl])C(=C([H])C=1OC([H])([H])[H])N([H])[H])N([H])C([H])([H])[C@]2([H])C([H])([H])C([H])([H])N(C2([H])[H])C([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3[H]</chem>	11000	1400	570	[62]
SC208	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C2=N/C(=C(/[H])[S]2)C([H])([H])N([H])C([H])([H])[C@]3([H])C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])C=4C([H])=C([H])C([H])=C([H])C=4[H]</chem>	8850	7900	660	[63]
SC209	<chem>[Br]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])C2=N/C(=C(/[H])O2)C([H])([H])N([H])C([H])([H])C([H])([H])[C@]3([H])C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])C=4C([H])=C([H])C([H])=C([H])C=4[H]</chem>	8950	1070	120	
SC210	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C3=C([H])C([H])=C([H])N3[H]</chem>	5400	3000	570	[107]

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC211	<chem>O=C(C=2[S]C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	8800	3600	0.5	[108]
SC212	<chem>O=C(C=2C([H])=C([Br])C=1C([H])=C([H])C([H])=C([H])C=1C=2OC([H])([H])[H]N([H])C([H])([H])C([H])([H])[C@]3([H])C([H])([H])C([H])([H])N(C3([H])[H])C([H])([H])C=4C([H])=C([H])C([H])=C([H])C=4[H]</chem>	1700	2000	580	[65]
SC213	<chem>O=C(C=2OC=1C([H])=C([H])C([I])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	3300	3200	5.7	[109]
SC214	<chem>O=C(C=2C([H])=C([H])C(C#CC=1C([H])=C([H])C([H])=C([H])C=1[H])=C([H])C=2[H])N([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	5500	4700	29	
SC215	<chem>O=C(C=2C([H])=C([H])C([H])=C(C#CC=1C([H])=C([H])C([H])=C([H])C=1[H])C=2[H])N([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[Cl])C([H])([H])C4([H])[H]</chem>	3100	1200	68	[110]
SC216	<chem>O=C(C=2C([H])=C([H])C(C#CC=1C([H])=C([H])C([H])=C([H])C=1[H])=C([H])C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H]C([H])([H])C4([H])[H]</chem>	16000	1400	10	
SC217	<chem>O=C(C=2C([H])=C([H])C([H])=C(C#CC=1C([H])=C([H])C([H])=C([H])C=1[H])C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])[H]C([H])([H])C4([H])[H]</chem>	5900	1200	22	
SC218	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[Cl])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C=4N=C3C([H])=C([H])C(=NN3C=4[H])N5C([H])([H])C([H])([H])N(C([H])([H])C5([H])[H])C([H])([H])[H]</chem>	13000	8000	880	[66]
SC219	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C([H])([H])N2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C4=C([H])C=3C([H])=C([H])C([H])=C(OC([H])([H])[H])C=3N4C([H])([H])C5([H])[H]</chem>	10000	1012	33.5	
SC220	<chem>O=C(/C2=N/C([H])=C1C([H])=C([H])C([H])=C([H])C1=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([H])C([H])=C([Cl])C=3[H])C([H])([H])C4([H])[H]</chem>	5500	1039	0.27	[111]
SC221	<chem>O=C3/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1N2C([H])([H])C([H])([H])N3C([H])([H])C([H])([H])C([H])([H])C([H])([H])N6C([H])([H])C5=C([H])C=4C([H])=C([H])C([H])=C(OC([H])([H])[H])C=4N5C([H])([H])C6([H])[H]</chem>	10000	3530	8.6	
SC222	<chem>O=C(/C2=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C([H])([H])N2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C4=N/C3=C([H])C([H])=C([H])C(OC([H])([H])[H])=C3[C@]4([H])C([H])([H])C5([H])[H]</chem>	10000	6475	6	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC236	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3[S]4</chem>	1273	2321	214	
SC237	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3O4</chem>	1172	3962	180	
SC238	<chem>[F]C([F])([F])C=3C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])N2C([H])([H])C([H])([H])N(/C1=N/C([H])=C([H])C([H])=C1[H])C([H])([H])C2([H])[H])=C([H])C=3[H]</chem>	2344	1092	355	[56]
SC239	<chem>[Cl]C=1C([H])=C([H])C(=C([H])C=1[H])N2C([H])([H])C([H])([H])N(C([H])([H])C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3O4</chem>	7939	3108	119	
SC240	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])C@@]2([H])OC([H])([H])C([H])([H])N(C2([H])[H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3N4[H]</chem>	100000	100000	10.8	
SC241	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])C@]2([H])OC([H])([H])C([H])([H])N(C2([H])[H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3N4[H]</chem>	100000	100000	25.9	[115]
SC242	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])C@@]2([H])OC([H])([H])C([H])([H])N(C2([H])[H])C([H])([H])C4=NC=3C([H])=C([H])C([H])=C([H])C=3N4[H]</chem>	100000	100000	15.7	
SC243	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])C([H])([H])C=3C([H])=C([H])C=2OC([H])([H])OC=2C=3[H])C([H])([H])C([H])([H])C=4C([H])=C([H])C(OC([H])([H])C([H])=C(OC([H])([H])C([H])C=4[H]</chem>	1682	1729	498	[116]
SC244	<chem>[H]C=1C([H])=C([H])C(=C([H])C=1[H])C([H])([H])C([H])([H])C([H])([H])OC=2C([H])=C([H])C(=C([H])C=2OC([H])([H])C([H])([H])C([H])([H])C([H])([H])N([H])C([H])([H])C@@]([H])(O[H])C=3C([H])=C([H])C(O[H])=C(O[H])C=3[H]</chem>	2900	1900	660	[96]
SC245	<chem>O=C2C([H])=C([H])C=1/C(=C(/[H])C([H])=C(O[H])C=1N2[H])C@]([H])(O[H])C([H])([H])N([H])C([H])([H])C([H])([H])C=4C([H])=C([H])C(OC([H])([H])C([H])([H])C([H])([H])C=3C([H])=C([H])C([H])=C([H])C=3[H])=C(OC([H])([H])C([H])C=4[H]</chem>	6800	2900	790	
SC246	<chem>[H]/C2=C(\ [H])C(O[H])=C1OC([H])([H])C([H])([H])O/C1=C2 \ N3C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])C([H])</chem>	7700	3200	790	
SC247	<chem>[H]/C2=C(\ [H])C(O[H])=C1OC([H])([H])C([H])([H])O/C1=C2 \ N3C([H])([H])C([H])([H])N([H])C([H])([H])C3([H])[H]</chem>	6000	6900	110	[99]
SC248	<chem>O=C1N([H])C2=C(OC1([H])[H])C(=C([H])C([H])=C2O[H])N3C([H])([H])C([H])([H])C([H])([H])N([H])C([H])([H])C3([H])[H]</chem>	20000	14000	510	
SC249	<chem>O=C1N([H])C=2C(OC1([H])[H])=C(C([H])=C([H])C=2O[H])N3C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])[H]</chem>	20000	6400	42	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC250	<chem>O=C2C([H])=C([H])C=1/C(=C/[H])C([H])=C(O[H])C=1N2([H])N3C([H])([H])C([H])([H])N(C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	20000	1900	420	
SC251	<chem>O=C1N([H])C2=C(OC1([H])[H])C(=C([H])C([H])=C2O[H])N3C([H])([H])C([H])([H])N([H])C([H])([H])C3([H])[H]</chem>	20000	8300	160	
SC252	<chem>O=C(/C1=N/N2/C(=C1/[H])C([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N5C([H])([H])C([H])([H])[N+]⁴(C=3C([H])=C([H])C([H])=C([H])C=3OC([H])([H])C4([H])[H])C([H])([H])C5([H])[H]</chem>	5700	6300	200	[100]
SC253	<chem>O=C(C=2[S]C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N/C3=N/C(=N/C(=C3/[H])C([F])([F])[F])C(C([H])([H])[H])C([H])([H])[H]C([H])([H])C([H])([H])C4([H])[H]</chem>	10000	4000	2.4	[100]
SC254	<chem>[H]/C1=C(\ [H])C(=C([H])C(O[H])=C1[H])[C@@]2([H])C([H])([H])C([H])([H])N(C2([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1278	3555	162	[101]
SC255	<chem>O=C(/C1=C(\ [H])C([H])=NC=2C([H])=C([H])C([H])=C([H])C1=2)N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C3=C([H])C(O[H])=C(OC([H])([H])[H])C([H])=C3C([H])([H])C4([H])[H]</chem>	1810	inactive	22	[117]
SC256	<chem>O=C(C=1C([H])=C([H])C([H])=C([H])C=1[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N3C([H])([H])C2=C([H])C(O[H])=C(OC([H])([H])[H])C([H])=C2C([H])([H])C3([H])[H]</chem>	1600	inactive	10	
SC257	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C=2C([H])=C([H])[S]C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N4C([H])([H])C([H])([H])N(C=3C([H])=C([Cl])C([H])=C([Cl])C=3[H])C([H])([H])C4([H])[H]</chem>	10000	2049	0.44	[103]
SC258	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([F])C=4[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1308	4823	26.7	
SC259	<chem>[F]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])[C@@]2([H])C([H])([H])[C@@]2([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])C3([H])[H])C([H])([H])C([H])([H])C([H])([H])[S]/C5=N/N=C(/C=4C([H])=C([H])C([H])=C([H])C=4[H])N5C([H])([H])[H]</chem>	1078	5000	64.6	
SC260	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([F])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1758	3144	4	[104]
SC261	<chem>[F]C=1C([H])=C([H])C(OC([H])([H])[H])=C(C=1[H])[C@@]2([H])C([H])([H])[C@]2([H])C([H])([H])N(C([H])([H])C([H])([H])C([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[S]/C4=N/N=C(/C=3C([H])=C([H])C([H])=C([H])C=3[H])N4C([H])([H])[H]</chem>	1668	1161.4	87.1	
SC262	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1711	5000	43	

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
	<chem>=C([H])C=4[H]C([F])([F])[F]C([H])([H])C([H])([H])C([H])([H])[H]</chem>				
SC263	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([F])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	2906	5000	45.4	
SC264	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([Cl])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	992	3494	11.1	
SC265	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([F])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	967	1083.9	14.5	
SC266	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([Cl])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	5000	2105	39.4	
SC267	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([Cl])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	930	1116	26.9	
SC268	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([F])=C([H])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	3631	5000	106.3	
SC269	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C(=C([H])C=4[H]C([F])([F])[F]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1423	5000	23.4	
SC270	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([F])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	4898	1349	4.1	
SC271	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@@]3([H])C([H])([H])[C@]3([H])C=4C([H])=C([H])C([H])=C([Cl])C=4[Cl]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	2344	1023	5.3	
SC272	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	940	4012	15.5	

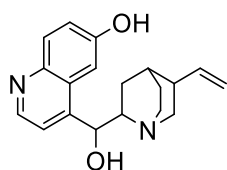
ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
	<chem>H)([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([Cl])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>				
SC273	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([Cl])C=4[Cl])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1647	2362	1.2	
SC274	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([Cl])=C([H])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1308	5000	17.6	
SC275	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([F])=C([H])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	992	5623	55.8	
SC276	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([F])=C([H])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1457	5934	45.7	
SC277	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([Cl])C=4[Cl])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1047	1148	20.8	
SC278	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C(=C([H])C=4[H]C([F])([F])[F])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1278	5000	33.1	
SC279	<chem>O=C(/C2=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@]3([H])C([H])([H])[C@@]3([H])C=4C([H])=C([H])C([H])=C([F])C=4[H]C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1105	3494	18.8	
SC280	<chem>O=C(C=1C([H])=C([H])C(=C([H])C=1[H])C2=NC([H])=C([H])C([H])=C2[H])N([H])C([H])([H])C([H])([H])C([H])([H])C([H])([H])N(C([H])([H])[C@@]3([H])C([H])([H])[C@]3([H])C=4C([H])=C([F])C([H])=C([H])C=4OC([H])([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	1071	1230	3.8	
SC281	<chem>O=C(C=2C([H])=C([H])C=1C([H])=C([H])C([H])=C([H])C=1C=2[H])N([H])[C@]3([H])C([H])([H])[C@](O[H])(C3([H])[H])C([H])([H])C([H])([H])N([C@]5([H])C([H])([H])C=4[S]C(=NC=4C([H])([H])C5([H])[H])N([H])[H])C([H])([H])C([H])([H])C([H])([H])[H]</chem>	inactive	3902	30	[105]
SC282	<chem>[H]C=4C([H])=C3C1=C(C([H])=C([H])C=2N=C([S]C1=2)N([H])[H])C([H])([H])[C@]5([H])C3=C(C=4[H])C([H])([H])C([H])([H])N5C([H])([H])[H]</chem>	1150	1740	381	[118]
SC283	<chem>[H]/C1=C(\ [H])C(=NC([H])=C1[C@@]2([H])OC([H])([H])[C@@]([H])(N(C2([H])[H])C([H])([H])C([H])([H])C([H])([H])[H])C([H])([H])[H])N([H])[H]</chem>	100000	1740	185	[106]

ID	SMILES	K _i [nM] ^a			Ref.
		D ₁ R	D ₂ R	D ₃ R	
SC284	<chem>O=C(/C=C(\ [H])C=1C([H])=C([H])C([H])=C([H])C=1N2[H])N([H])C([H])([H])C([H])([H])[C@@]([H])(O[H])C([H])([H])N3C([H])([H])[C@@]([H])(OC([H])([H])[C@]3([H])C([H])([H])[H])C=4C([H])=C([H])C(=NC=4[H])N([H])[H]</chem>	100000	1010	423	

^aAll KI values retrieved from ChEMBL were determined in vitro utilizing radio-ligand binding assays.

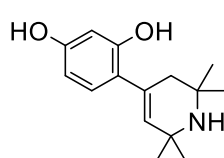
2. In vitro screening– overview of 2D structures and NDF values of all compounds

Table S5. Overview of all compounds selected for further in vitro investigations. 2D structures and NDF ± standard deviation (SD) are shown for each compound (n = 4). All NDF values for D₂R were obtained in the previously published article of Zell et al [42]. NDF, normalized decreased fluorescence.



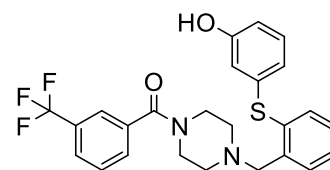
SC285

NDF (D₁R) = 0.83 ± 0.06
NDF (D₂R) = 1.13 ± 0.37
NDF (D₃R) = 1.17 ± 0.11



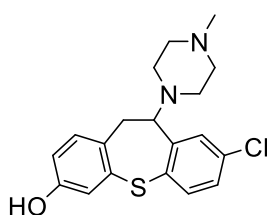
SC286

NDF (D₁R) = 0.88 ± 0.20
NDF (D₂R) = 1.05 ± 0.45
NDF (D₃R) = 1.04 ± 0.10



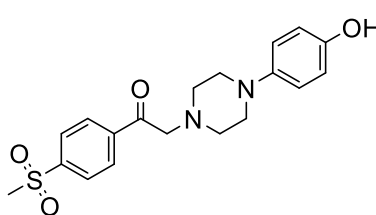
SC287

NDF (D₁R) = 0.91 ± 0.21
NDF (D₂R) = 0.90 ± 0.25
NDF (D₃R) = 1.00 ± 0.17



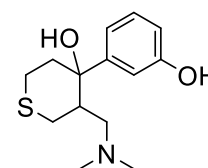
2

NDF (D₁R) = 5.41 ± 1.97
NDF (D₂R) = 40.41 ± 1.39
NDF (D₃R) = 4.16 ± 1.08



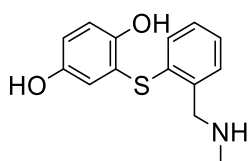
SC288

NDF (D₁R) = 3.21 ± 3.27
NDF (D₂R) = 1.01 ± 0.38
NDF (D₃R) = 1.58 ± 0.58



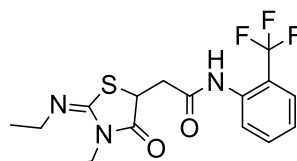
SC289

NDF (D₁R) = 1.66 ± 1.16
NDF (D₂R) = 0.90 ± 0.30
NDF (D₃R) = 1.24 ± 0.25



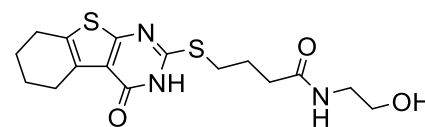
SC290

NDF (D₁R) = 3.50 ± 3.16
NDF (D₂R) = 0.91 ± 0.36
NDF (D₃R) = 1.81 ± 1.04



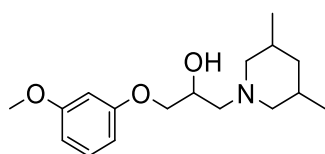
SC291

NDF (D₁R) = 0.81 ± 0.08
NDF (D₂R) = 0.84 ± 0.23
NDF (D₃R) = 1.02 ± 0.24



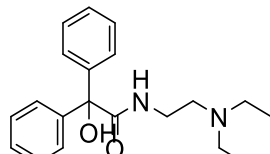
SC292

NDF (D₁R) = 0.80 ± 0.21
NDF (D₂R) = 0.95 ± 0.43
NDF (D₃R) = 0.92 ± 0.13



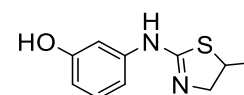
SC293

NDF (D₁R) = 0.85 ± 0.14
NDF (D₂R) = 1.02 ± 0.39
NDF (D₃R) = 1.04 ± 0.11



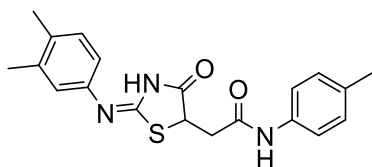
SC294

NDF (D₁R) = 2.72 ± 2.66
NDF (D₂R) = 1.01 ± 0.50
NDF (D₃R) = 1.49 ± 0.54



SC295

NDF (D₁R) = 1.08 ± 0.42
NDF (D₂R) = 0.95 ± 0.43
NDF (D₃R) = 1.26 ± 0.25

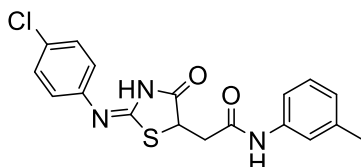


SC296

NDF (D₁R) = 1.02 ± 0.36

NDF (D₂R) = 0.94 ± 0.32

NDF (D₃R) = 0.93 ± 0.12

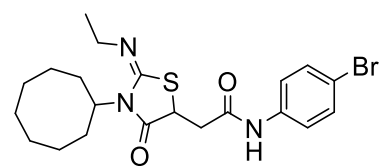


SC297

NDF (D₁R) = 0.89 ± 0.24

NDF (D₂R) = 0.96 ± 0.38

NDF (D₃R) = 1.04 ± 0.11

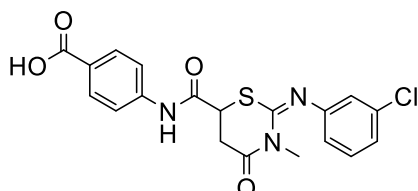


SC298

NDF (D₁R) = 1.21 ± 0.43

NDF (D₂R) = 1.00 ± 0.35

NDF (D₃R) = 0.85 ± 0.13

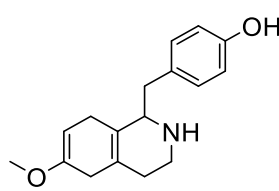


SC299

NDF (D₁R) = 0.94 ± 0.13

NDF (D₂R) = 0.82 ± 0.25

NDF (D₃R) = 1.05 ± 0.36

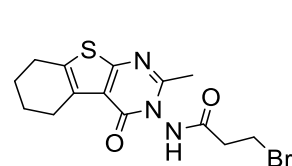


SC300

NDF (D₁R) = 2.72 ± 0.84

NDF (D₂R) = 1.14 ± 0.56

NDF (D₃R) = 2.16 ± 0.40

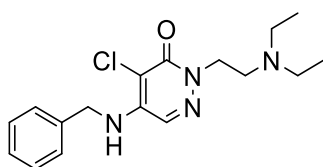


SC301

NDF (D₁R) = 0.78 ± 0.22

NDF (D₂R) = 0.99 ± 0.35

NDF (D₃R) = 0.97 ± 0.11

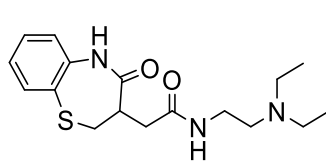


SC302

NDF (D₁R) = 1.04 ± 0.28

NDF (D₂R) = 0.99 ± 0.39

NDF (D₃R) = 1.26 ± 0.14

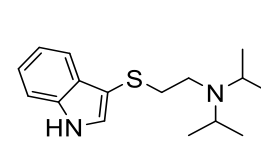


SC303

NDF (D₁R) = 1.00 ± 0.33

NDF (D₂R) = 1.02 ± 0.36

NDF (D₃R) = 1.50 ± 0.24

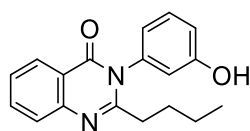


SC304

NDF (D₁R) = 0.88 ± 0.17

NDF (D₂R) = 0.98 ± 0.32

NDF (D₃R) = 1.20 ± 0.11

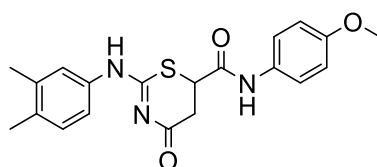


SC305

NDF (D₁R) = 0.90 ± 0.26

NDF (D₂R) = 0.93 ± 0.29

NDF (D₃R) = 1.03 ± 0.14

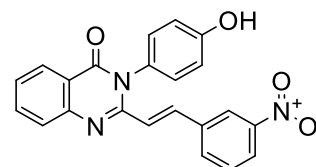


SC306

NDF (D₁R) = 1.05 ± 0.44

NDF (D₂R) = 0.94 ± 0.36

NDF (D₃R) = 1.09 ± 0.30

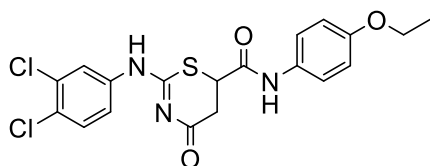


SC307

NDF (D₁R) = 0.93 ± 0.16

NDF (D₂R) = 0.86 ± 0.24

NDF (D₃R) = 0.93 ± 0.15

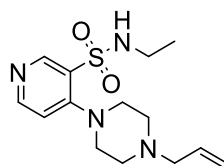


SC308

NDF (D₁R) = 0.87 ± 0.20

NDF (D₂R) = 2.04 ± 1.34

NDF (D₃R) = 1.64 ± 0.29

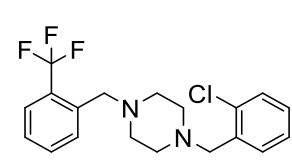


SC309

NDF (D₁R) = 0.96 ± 0.34

NDF (D₂R) = 1.10 ± 0.46

NDF (D₃R) = 2.03 ± 0.18

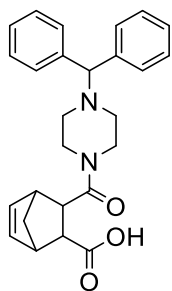


SC310

NDF (D₁R) = 0.99 ± 0.34

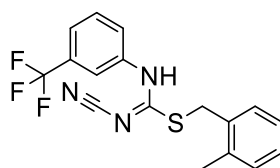
NDF (D₂R) = 1.06 ± 0.41

NDF (D₃R) = 1.54 ± 0.33



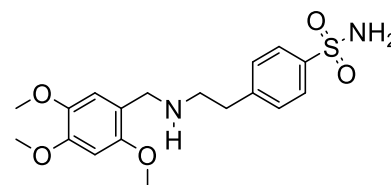
SC311

NDF (D₁R) = 1.09 ± 0.43
 NDF (D₂R) = 1.02 ± 0.35
 NDF (D₃R) = 1.67 ± 0.44



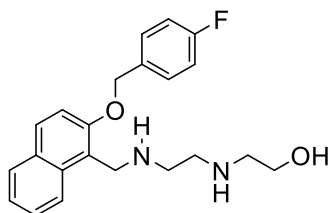
SC312

NDF (D₁R) = 0.97 ± 0.40
 NDF (D₂R) = 1.10 ± 0.44
 NDF (D₃R) = 1.23 ± 0.27



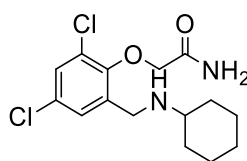
SC313

NDF (D₁R) = 0.88 ± 0.33
 NDF (D₂R) = 0.94 ± 0.32
 NDF (D₃R) = 1.12 ± 0.15



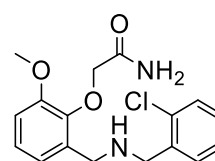
SC314

NDF (D₁R) = 4.17 ± 1.08
 NDF (D₂R) = 1.68 ± 0.90
 NDF (D₃R) = 3.00 ± 0.61



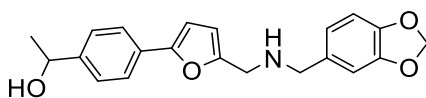
SC315

NDF (D₁R) = 1.18 ± 1.01
 NDF (D₂R) = 0.95 ± 0.32
 NDF (D₃R) = 1.06 ± 0.20



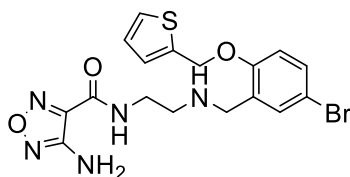
SC316

NDF (D₁R) = 0.93 ± 0.30
 NDF (D₂R) = 1.01 ± 0.44
 NDF (D₃R) = 1.16 ± 0.16



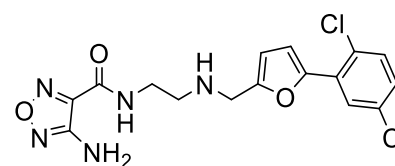
SC317

NDF (D₁R) = 1.06 ± 0.47
 NDF (D₂R) = 0.93 ± 0.42
 NDF (D₃R) = 1.07 ± 0.15



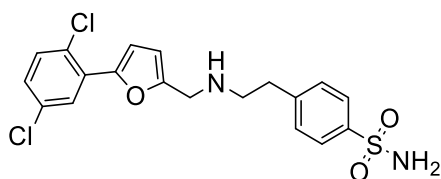
SC318

NDF (D₁R) = 1.81 ± 0.42
 NDF (D₂R) = 1.14 ± 0.57
 NDF (D₃R) = 2.39 ± 0.48



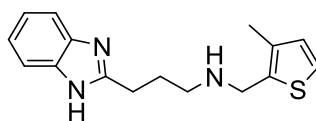
SC319

NDF (D₁R) = 0.89 ± 0.26
 NDF (D₂R) = 2.41 ± 2.01
 NDF (D₃R) = 3.64 ± 1.08



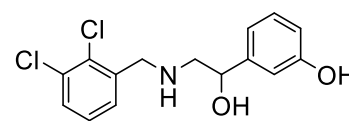
3

NDF (D₁R) = 1.78 ± 1.35
 NDF (D₂R) = 3.99 ± 2.58
 NDF (D₃R) = 3.96 ± 1.10



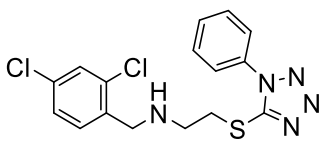
SC320

NDF (D₁R) = -
 NDF (D₂R) = -
 NDF (D₃R) = -



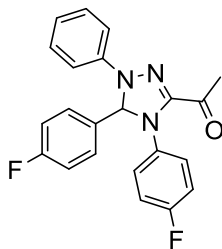
4

NDF (D₁R) = 0.90 ± 0.31
 NDF (D₂R) = 0.90 ± 0.38
 NDF (D₃R) = 2.75 ± 0.61



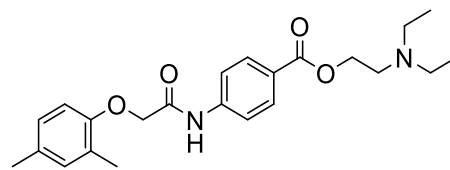
SC321

NDF (D₁R) = 1.00 ± 0.42
 NDF (D₂R) = 0.97 ± 0.36
 NDF (D₃R) = 1.26 ± 0.24



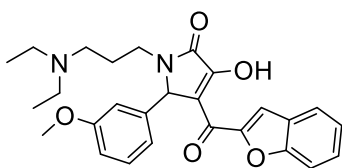
SC322

NDF (D₁R) = 0.86 ± 0.24
 NDF (D₂R) = 0.90 ± 0.25
 NDF (D₃R) = 1.01 ± 0.23



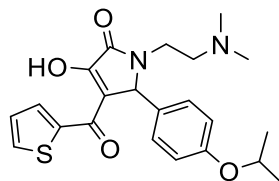
SC323

NDF (D₁R) = 1.43 ± 0.58
 NDF (D₂R) = 0.95 ± 0.35
 NDF (D₃R) = 1.55 ± 0.23



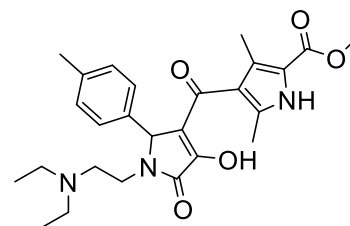
SC324

NDF (D₁R) = -
 NDF (D₂R) = -
 NDF (D₃R) = -



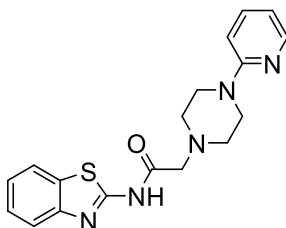
SC325

NDF (D₁R) = 1.02 ± 0.46
 NDF (D₂R) = 0.94 ± 0.46
 NDF (D₃R) = 1.22 ± 0.27



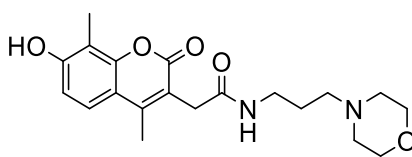
SC326

NDF (D₁R) = 0.87 ± 0.33
 NDF (D₂R) = 0.95 ± 0.42
 NDF (D₃R) = 1.06 ± 0.36



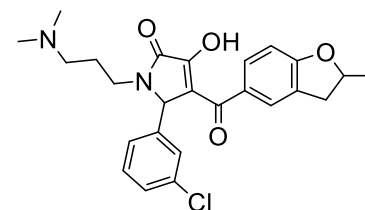
SC327

NDF (D₁R) = 0.92 ± 0.39
 NDF (D₂R) = 0.90 ± 0.39
 NDF (D₃R) = 1.00 ± 0.20



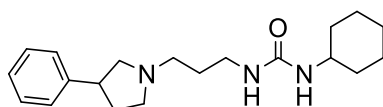
SC328

NDF (D₁R) = 0.43 ± 0.15
 NDF (D₂R) = 0.99 ± 0.43
 NDF (D₃R) = 1.13 ± 0.31



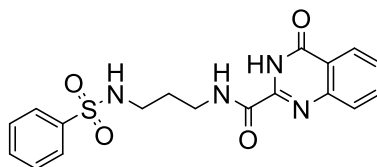
SC329

NDF (D₁R) = -
 NDF (D₂R) = -
 NDF (D₃R) = -



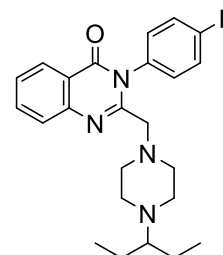
SC330

NDF (D₁R) = 1.88 ± 0.72
 NDF (D₂R) = 1.35 ± 0.68
 NDF (D₃R) = 3.31 ± 0.92



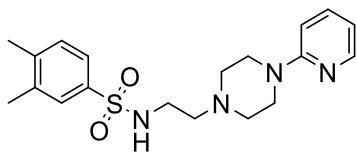
SC331

NDF (D₁R) = 0.95 ± 0.17
 NDF (D₂R) = 0.90 ± 0.35
 NDF (D₃R) = 1.14 ± 0.40



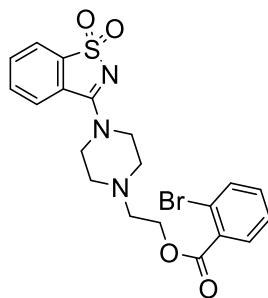
SC332

NDF (D₁R) = 0.97 ± 0.40
 NDF (D₂R) = 0.90 ± 0.36
 NDF (D₃R) = 1.18 ± 0.28



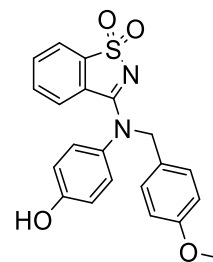
SC333

NDF (D₁R) = 1.13 ± 0.52
 NDF (D₂R) = 2.98 ± 1.77
 NDF (D₃R) = 2.49 ± 0.47



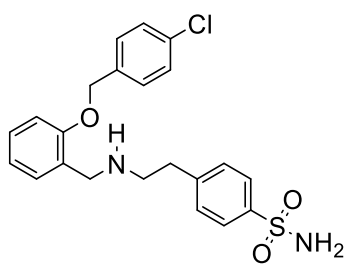
SC334

NDF (D₁R) = 0.95 ± 0.35
 NDF (D₂R) = 0.96 ± 0.37
 NDF (D₃R) = 1.12 ± 0.30



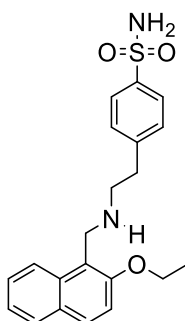
SC335

NDF (D₁R) = 0.93 ± 0.39
 NDF (D₂R) = 1.03 ± 0.42
 NDF (D₃R) = 1.40 ± 0.54



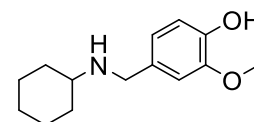
SC336

NDF (D₁R) = 1.46 ± 0.71
 NDF (D₂R) = 1.07 ± 0.46
 NDF (D₃R) = 2.76 ± 0.63



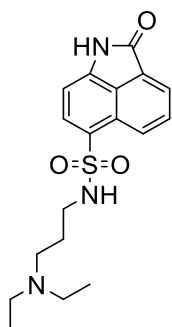
SC337

NDF (D₁R) = 1.12 ± 0.56
 NDF (D₂R) = 0.99 ± 0.40
 NDF (D₃R) = 1.24 ± 0.22



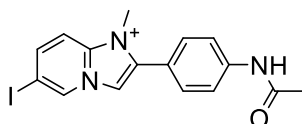
SC338

NDF (D₁R) = 1.14 ± 0.77
 NDF (D₂R) = 0.93 ± 0.38
 NDF (D₃R) = 1.15 ± 0.35



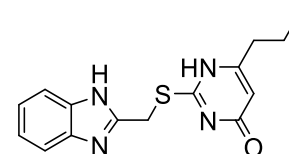
SC339

NDF (D₁R) = 0.59 ± 0.17
 NDF (D₂R) = 0.94 ± 0.41
 NDF (D₃R) = 1.11 ± 0.48



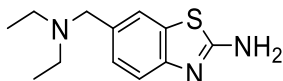
SC340

NDF (D₁R) = 0.83 ± 0.35
 NDF (D₂R) = 1.41 ± 1.07
 NDF (D₃R) = 1.10 ± 0.29



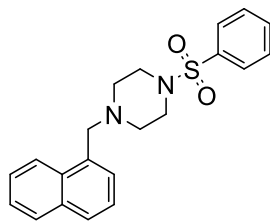
SC341

NDF (D₁R) = 0.83 ± 0.51
 NDF (D₂R) = 1.52 ± 1.15
 NDF (D₃R) = 0.92 ± 0.14



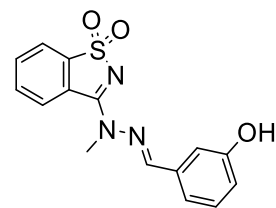
SC342

NDF (D₁R) = 1.24 ± 0.51
NDF (D₂R) = 1.38 ± 0.91
NDF (D₃R) = 1.24 ± 0.29



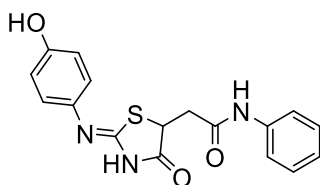
SC343

NDF (D₁R) = 1.36 ± 0.89
NDF (D₂R) = 1.06 ± 0.44
NDF (D₃R) = 1.15 ± 0.32



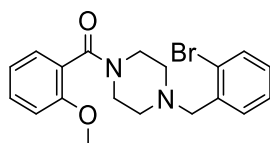
SC344

NDF (D₁R) = 1.09 ± 0.58
NDF (D₂R) = 1.06 ± 0.42
NDF (D₃R) = 0.99 ± 0.13



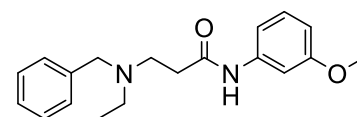
SC345

NDF (D₁R) = 0.93 ± 0.32
NDF (D₂R) = 0.95 ± 0.38
NDF (D₃R) = 1.11 ± 0.36



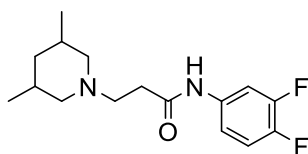
SC346

NDF (D₁R) = 1.04 ± 0.54
NDF (D₂R) = 0.97 ± 0.39
NDF (D₃R) = 1.15 ± 0.37



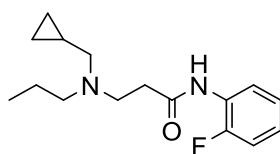
5

NDF (D₁R) = 1.26 ± 0.64
NDF (D₂R) = 15.74 ± 18.15
NDF (D₃R) = 2.65 ± 0.64



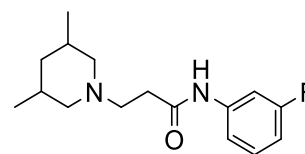
SC347

NDF (D₁R) = 1.12 ± 0.60
NDF (D₂R) = 1.05 ± 0.50
NDF (D₃R) = 1.40 ± 0.33



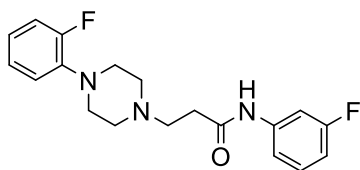
SC348

NDF (D₁R) = 0.89 ± 0.31
NDF (D₂R) = 1.02 ± 0.38
NDF (D₃R) = 1.36 ± 0.36



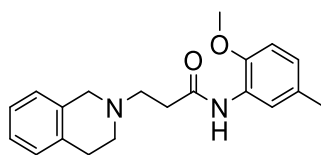
SC349

NDF (D₁R) = 1.12 ± 0.59
NDF (D₂R) = 1.26 ± 0.70
NDF (D₃R) = 1.95 ± 0.47



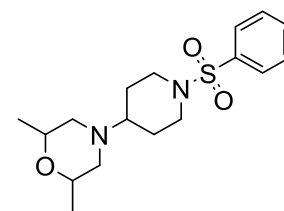
6

NDF (D₁R) = 2.15 ± 1.16
NDF (D₂R) = 8.18 ± 3.62
NDF (D₃R) = 4.21 ± 0.79



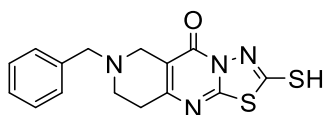
SC350

NDF (D₁R) = 1.19 ± 0.57
NDF (D₂R) = 1.83 ± 0.83
NDF (D₃R) = 3.25 ± 0.47



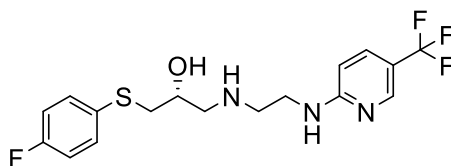
SC351

NDF (D₁R) = 0.93 ± 0.39
NDF (D₂R) = 0.99 ± 0.42
NDF (D₃R) = 1.19 ± 0.45



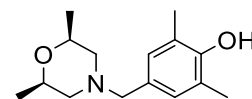
SC352

NDF (D₁R) = 0.91 ± 0.42
NDF (D₂R) = 1.02 ± 0.47
NDF (D₃R) = 1.22 ± 0.38



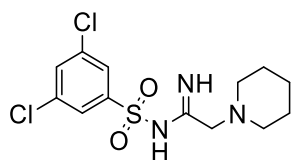
7

NDF (D₁R) = 1.57 ± 0.54
NDF (D₂R) = 10.85 ± 4.93
NDF (D₃R) = 3.79 ± 0.70



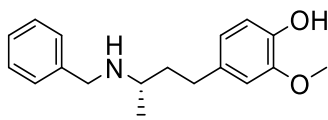
SC353

NDF (D₁R) = 1.08 ± 0.61
NDF (D₂R) = 1.00 ± 0.40
NDF (D₃R) = 1.20 ± 0.43



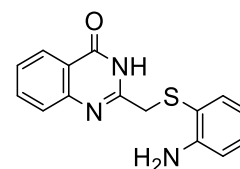
SC354

NDF (D₁R) = 0.98 ± 0.52
NDF (D₂R) = 0.99 ± 0.39
NDF (D₃R) = 1.06 ± 0.24



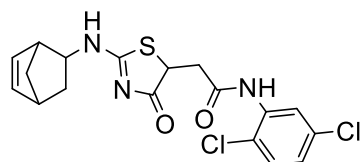
8

NDF (D₁R) = 1.20 ± 0.59
NDF (D₂R) = 1.10 ± 0.52
NDF (D₃R) = 2.63 ± 0.52



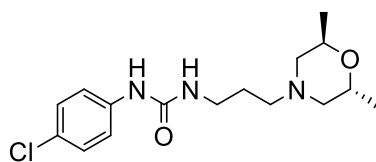
SC355

NDF (D₁R) = 1.34 ± 1.14
NDF (D₂R) = 1.13 ± 0.34
NDF (D₃R) = 1.22 ± 0.35



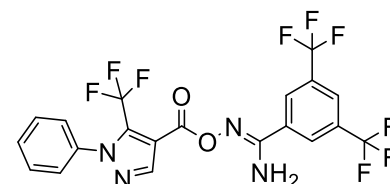
SC356

NDF (D₁R) = 1.28 ± 0.99
NDF (D₂R) = 2.44 ± 1.39
NDF (D₃R) = 1.23 ± 0.29



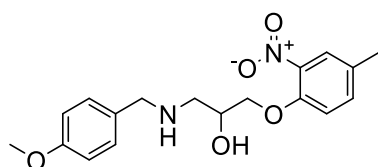
SC357

NDF (D₁R) = 1.05 ± 0.57
NDF (D₂R) = 1.66 ± 1.60
NDF (D₃R) = 2.02 ± 0.83



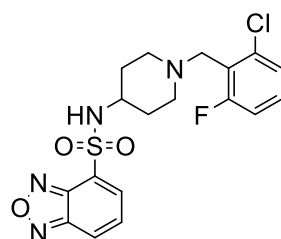
SC358

NDF (D₁R) = 1.33 ± 0.55
NDF (D₂R) = 1.11 ± 0.47
NDF (D₃R) = 1.22 ± 0.53



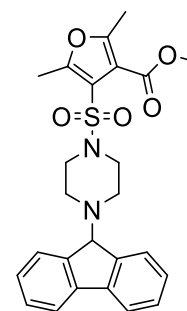
SC359

NDF (D₁R) = 1.28 ± 0.73
NDF (D₂R) = 2.00 ± 0.85
NDF (D₃R) = 2.49 ± 0.74



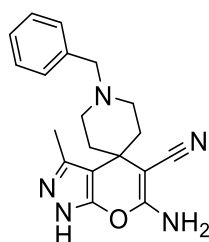
SC360

NDF (D₁R) = 1.05 ± 0.56
NDF (D₂R) = 0.93 ± 0.40
NDF (D₃R) = 1.35 ± 0.33



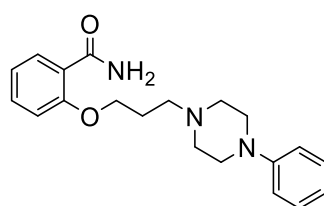
SC361

NDF (D₁R) = 0.96 ± 0.52
NDF (D₂R) = 1.11 ± 0.51
NDF (D₃R) = 1.10 ± 0.24



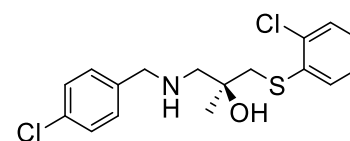
SC362

NDF (D₁R) = 1.05 ± 0.36
NDF (D₂R) = 1.41 ± 0.71
NDF (D₃R) = 2.03 ± 0.45



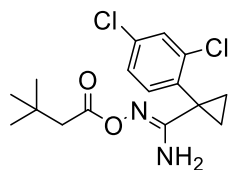
9

NDF (D₁R) = 1.71 ± 0.86
NDF (D₂R) = 22.08 ± 6.62
NDF (D₃R) = 3.97 ± 0.78



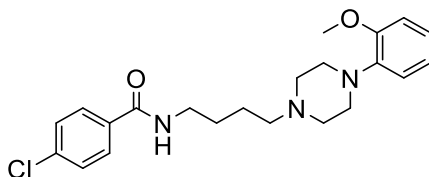
SC363

NDF (D₁R) = 1.11 ± 0.70
NDF (D₂R) = 1.11 ± 0.53
NDF (D₃R) = 1.53 ± 0.21



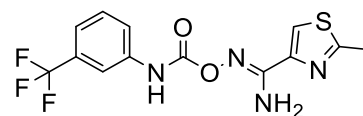
SC364

NDF (D₁R) = 0.92 ± 0.42
 NDF (D₂R) = 1.08 ± 0.56
 NDF (D₃R) = 1.17 ± 0.37



10

NDF (D₁R) = 2.59 ± 1.04
 NDF (D₂R) = 22.89 ± 8.41
 NDF (D₃R) = 4.09 ± 1.07



SC365

NDF (D₁R) = 0.99 ± 0.38
 NDF (D₂R) = 1.04 ± 0.39
 NDF (D₃R) = 1.12 ± 0.14

3. In silico assessment of DR subtype selectivity – a distance-based approach

3.1. ChEMBL validation

Table S6. Summary of the validation of the distance-based in silico approach using the described datasets extracted from ChEMBL. Cons. Gly, conserved glycine residue within the respective SBP. COM, center of mass.

3.2. Retrospective analysis of investigated compounds

Table S7. Summary of the distance-based in silico approach used during the investigation of the binding poses of the selected compounds **1** to **10**. Cpd., compound. Cons. Gly, conserved glycine residue within the respective SBP. COM, center of mass.

Cpd.	Pose rank (out of 30)			Distance (cons. Gly – COM) [Å]			Fold-difference (distance-based)		
	D ₁ R	D ₂ R	D ₃ R	D ₁ R	D ₂ R	D ₃ R	D ₁ R/D ₂ R	D ₁ R/D ₃ R	D ₂ R/D ₃ R
1	4 th	1 st	5 th	19.96	17.70	16.70	1.13	1.19	1.06
2	1 st	4 th	1 st	19.20	14.47	14.96	1.33	1.28	0.97
3	1 st	1 st	1 st	18.16	13.41	14.15	1.35	1.28	0.95
4	1 st	2 nd	1 st	17.70	15.42	14.06	1.15	1.26	1.10
5	1 st	6 th	4 th	18.73	13.95	14.31	1.34	1.31	0.98
6	1 st	1 st	2 nd	17.12	14.77	15.14	1.16	1.13	0.98
7	1 st	2 nd	2 nd	18.32	12.99	14.05	1.41	1.30	0.93
8	1 st	1 st	7 th	19.27	16.45	15.77	1.17	1.22	1.04
9	2 nd	2 nd	2 nd	19.29	12.83	14.07	1.50	1.37	0.91
10	1 st	1 st	1 st	18.11	12.84	13.50	1.41	1.34	0.95

4. In vitro careening activity of selected compounds

4.1. Summary of in vitro activity (based on NDF values) of control compounds and selected virtual hits

Screening results in vitro activity screened at 10 μ M

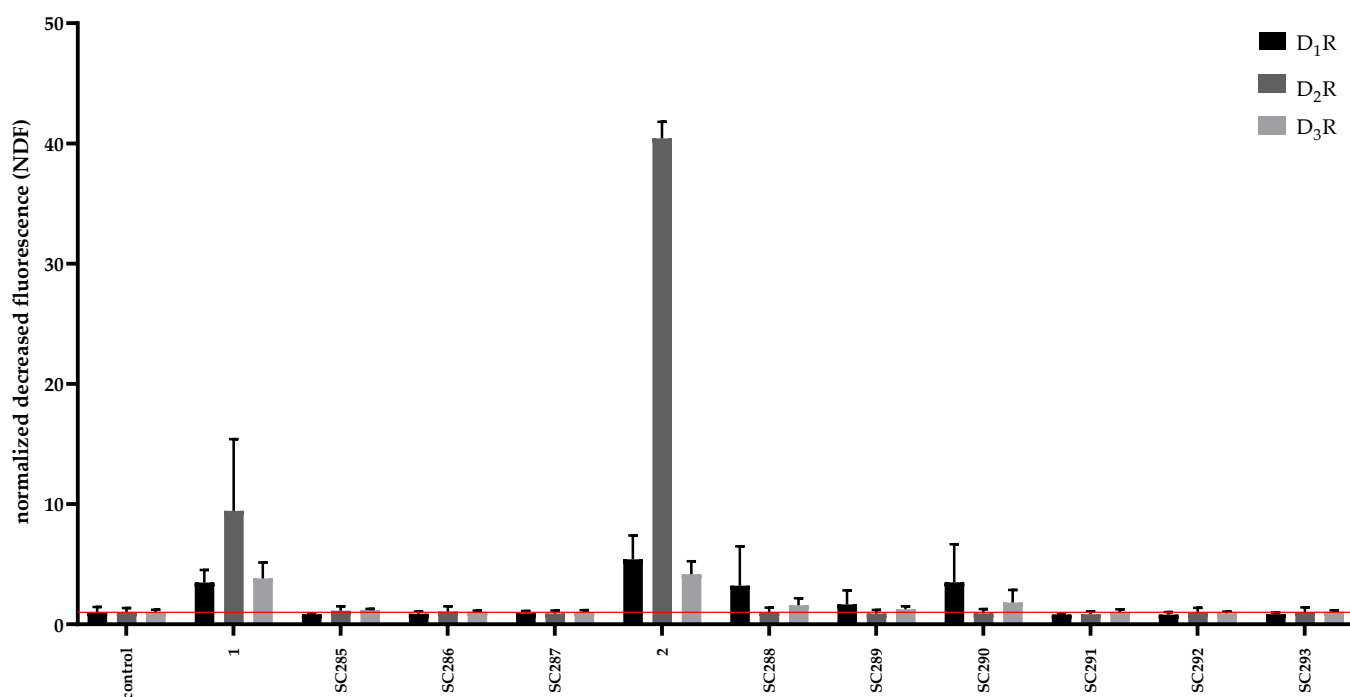


Figure S1. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

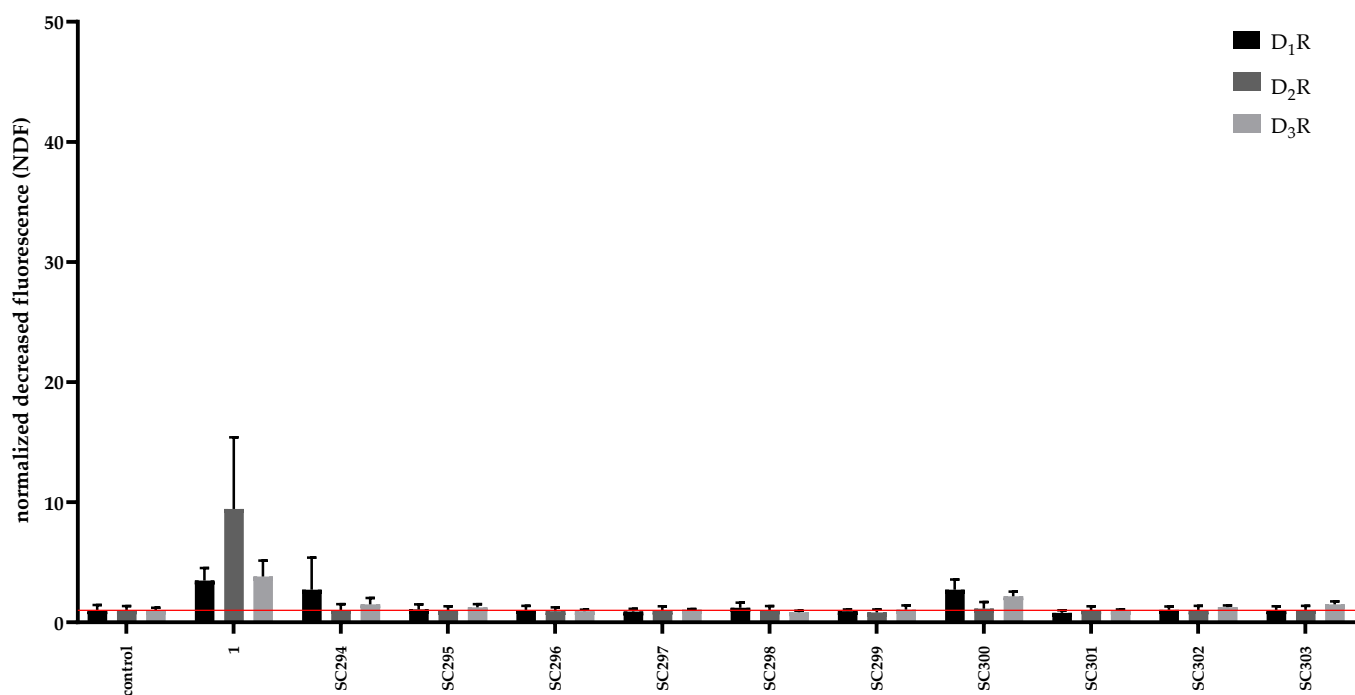


Figure S2. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

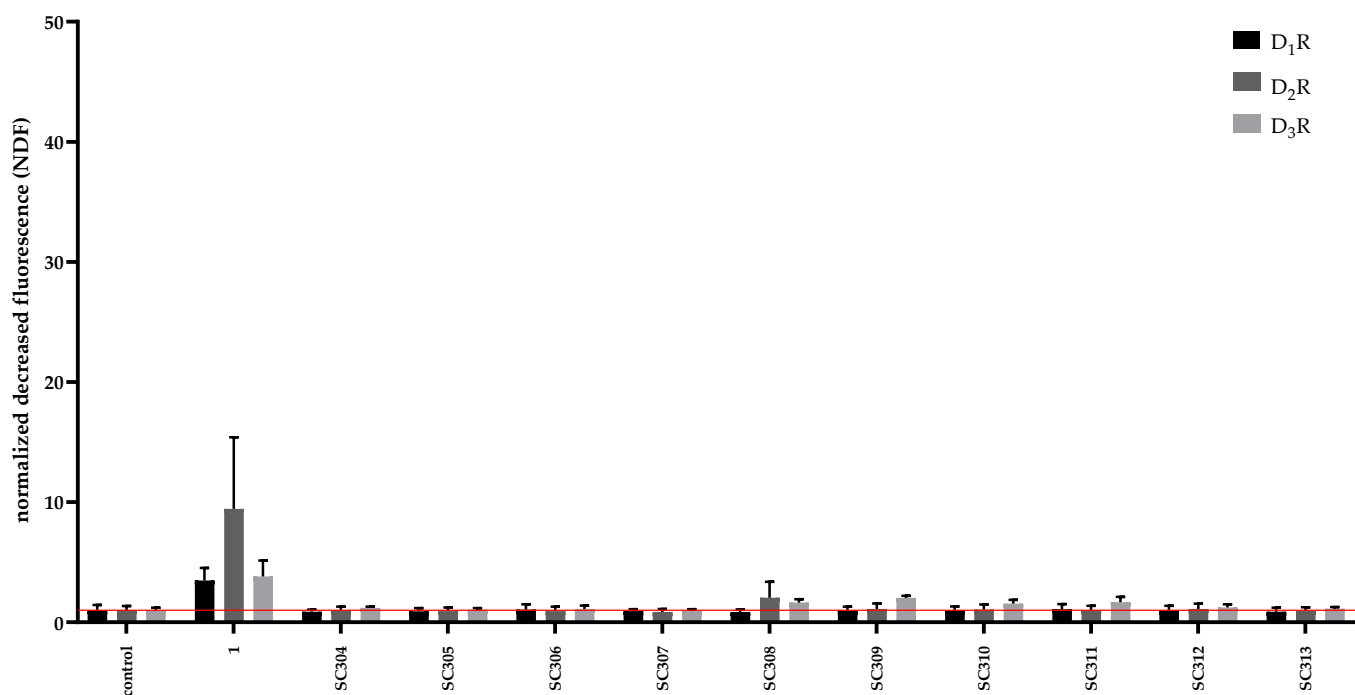


Figure S3. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

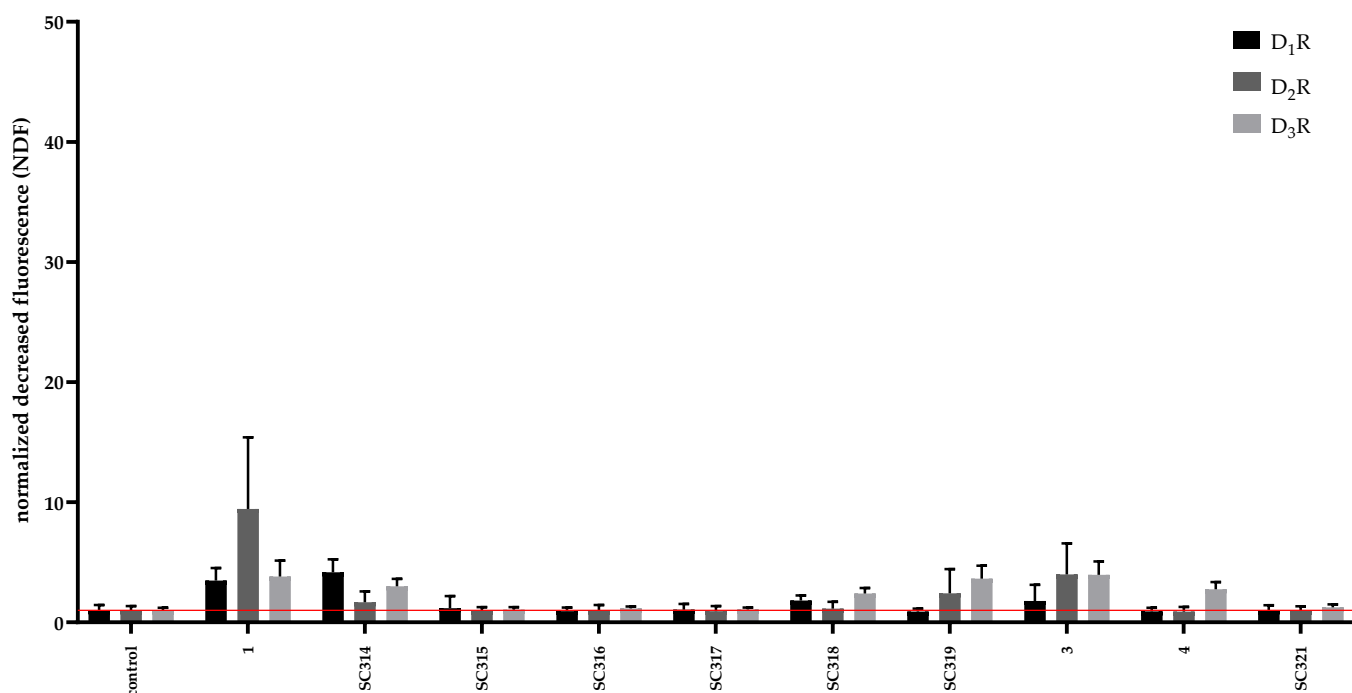


Figure S4. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

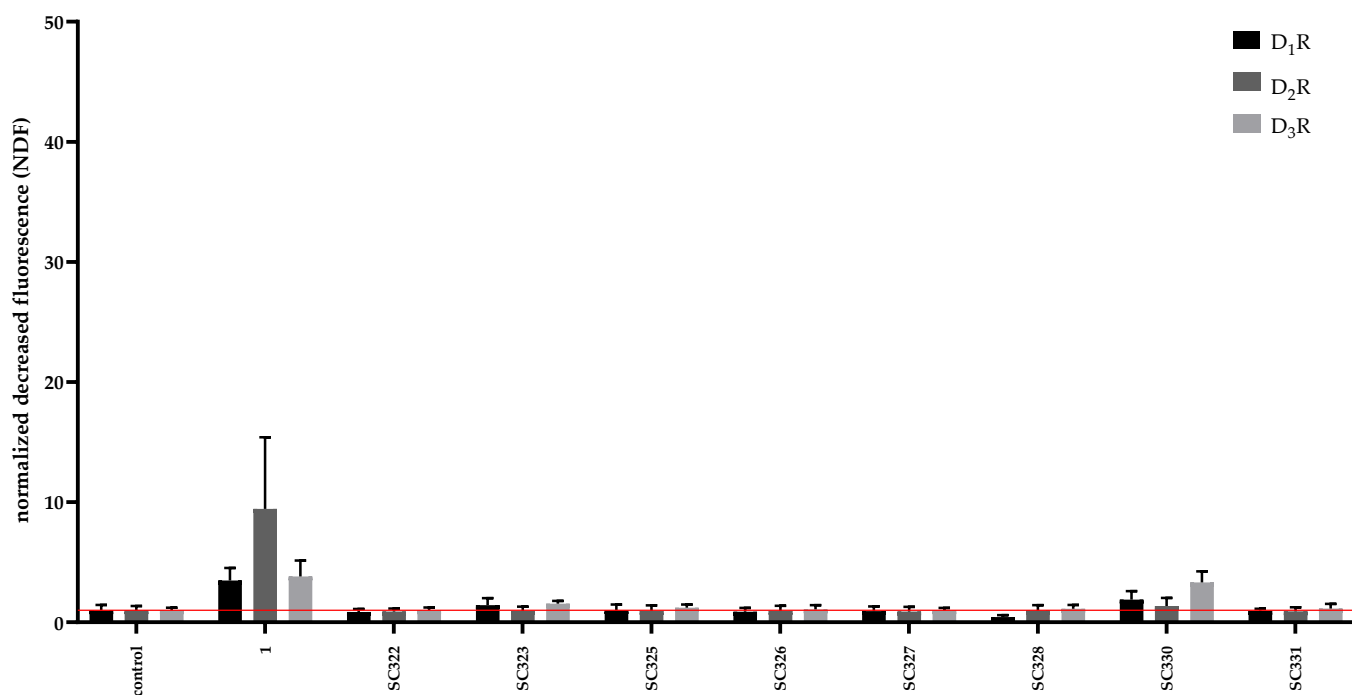


Figure S5. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

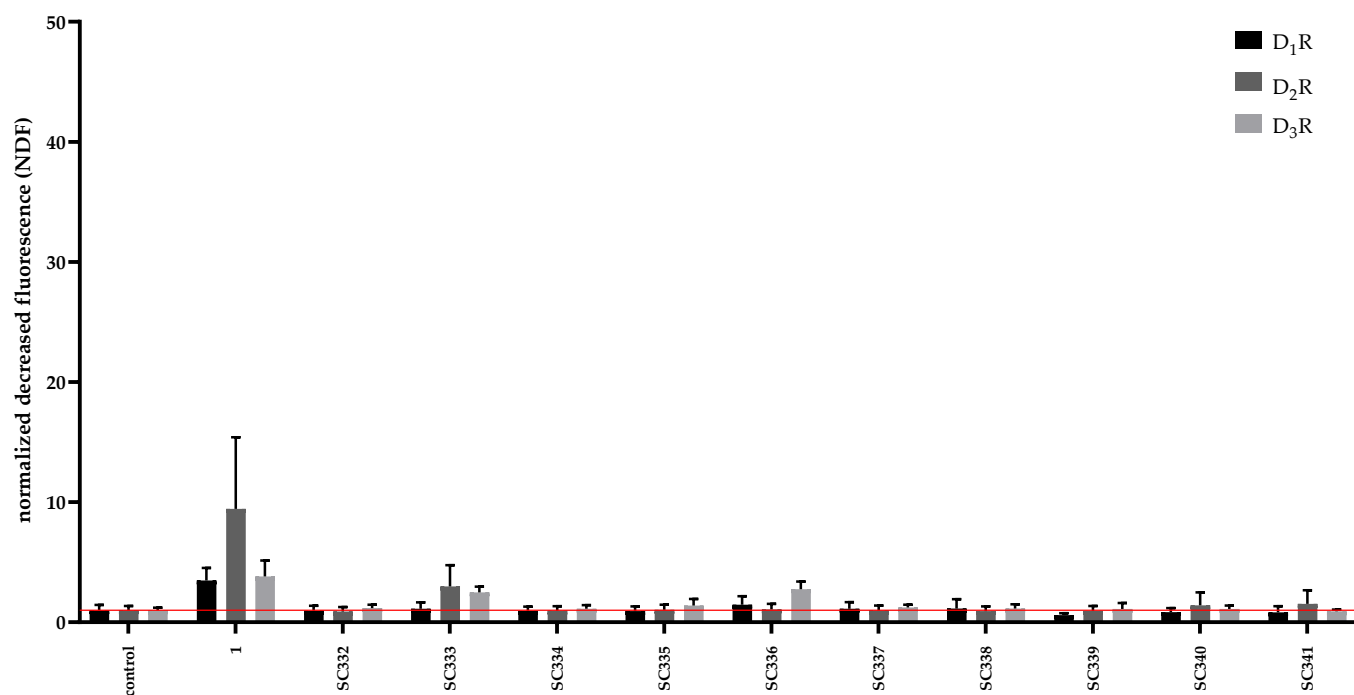


Figure S6. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

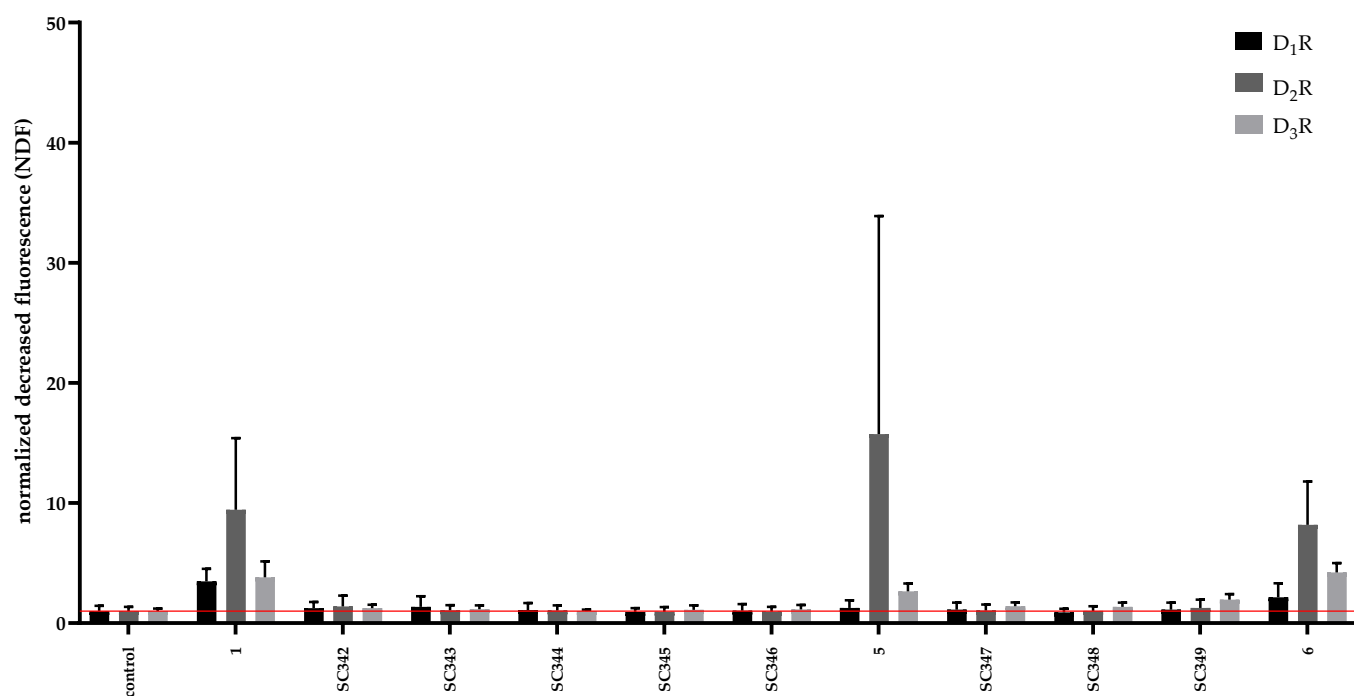


Figure S7. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

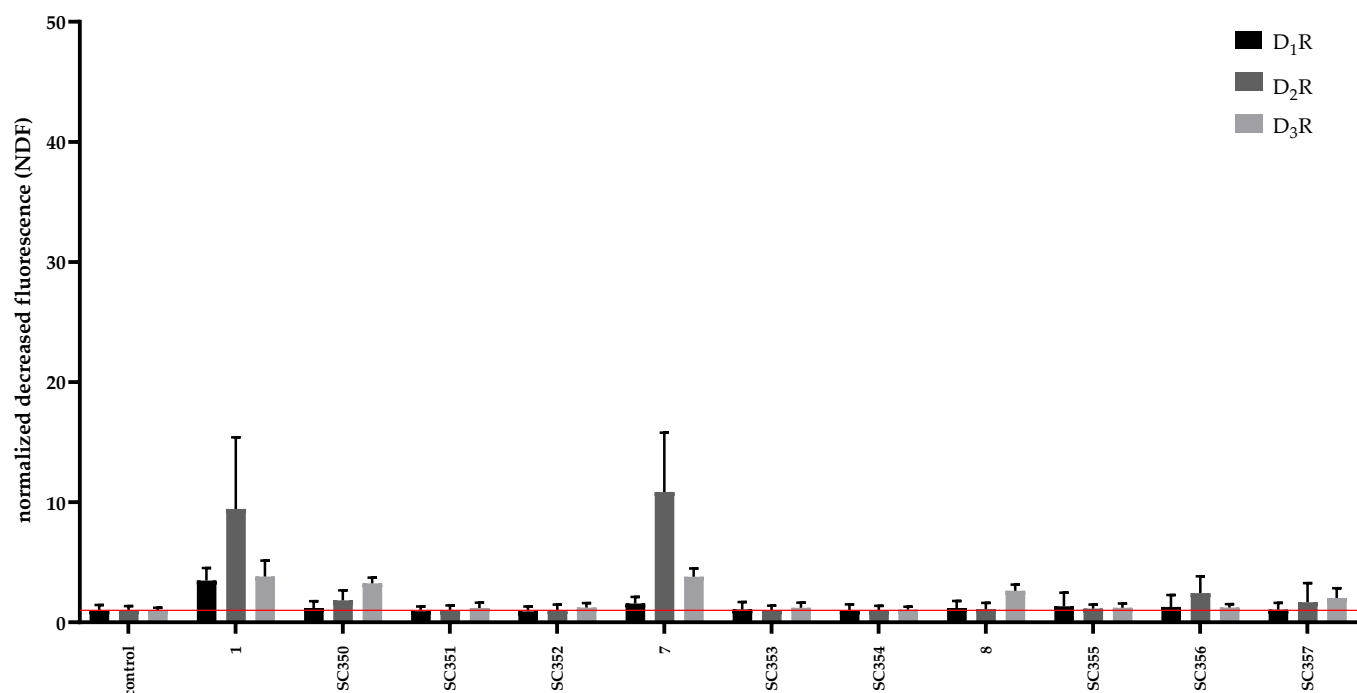


Figure S8. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

Screening results
in vitro activity screened at 10 μ M

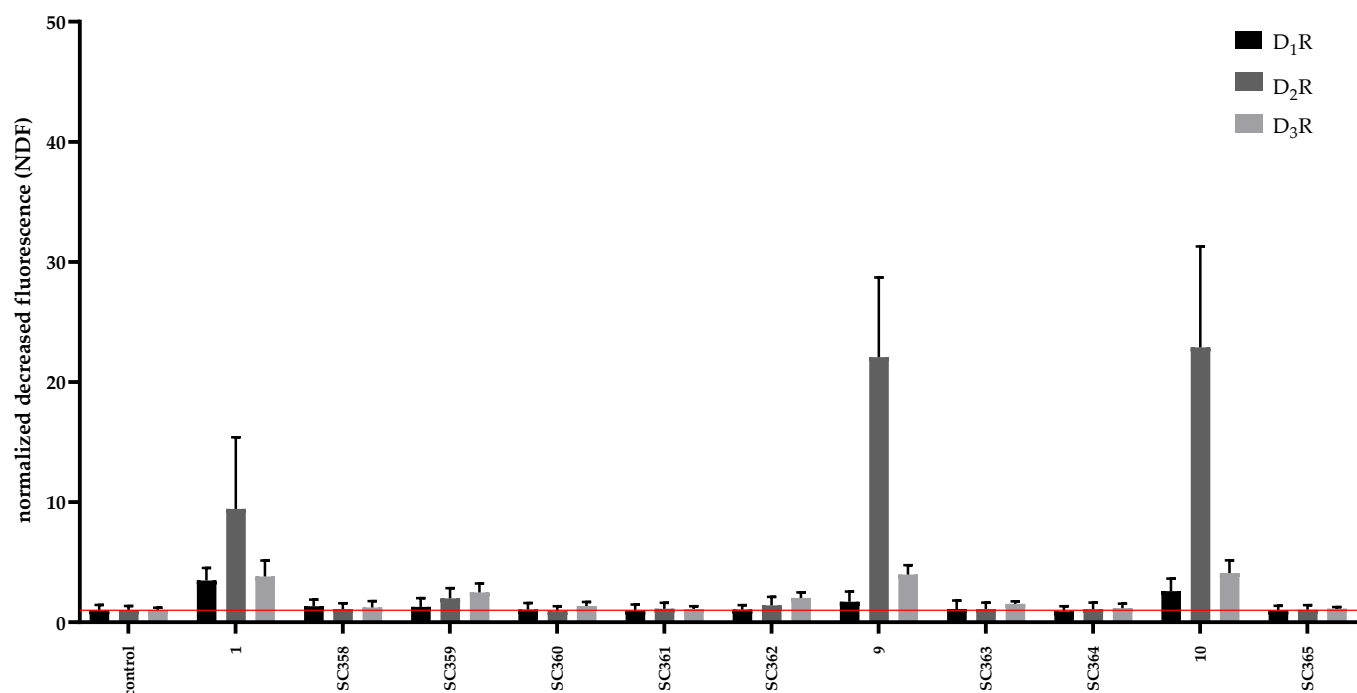


Figure S9. Comparison of biological activity of selected compounds at DR subtypes D₁R (black), D₂R (dark grey) and D₃R (light grey). Compound binding is shown as NDF in comparison to the control. Activities were measured in biological duplicates (two technical replicates each). Error bars show standard deviations (SD). The red line indicates a NDF of 1, thus, an inactive ligand at 10 μ M screening concentration. NDF, normalized decreased fluorescence.

4.2. Summary of K_1 values determined *in vitro* – overview of DR subtype selectivity

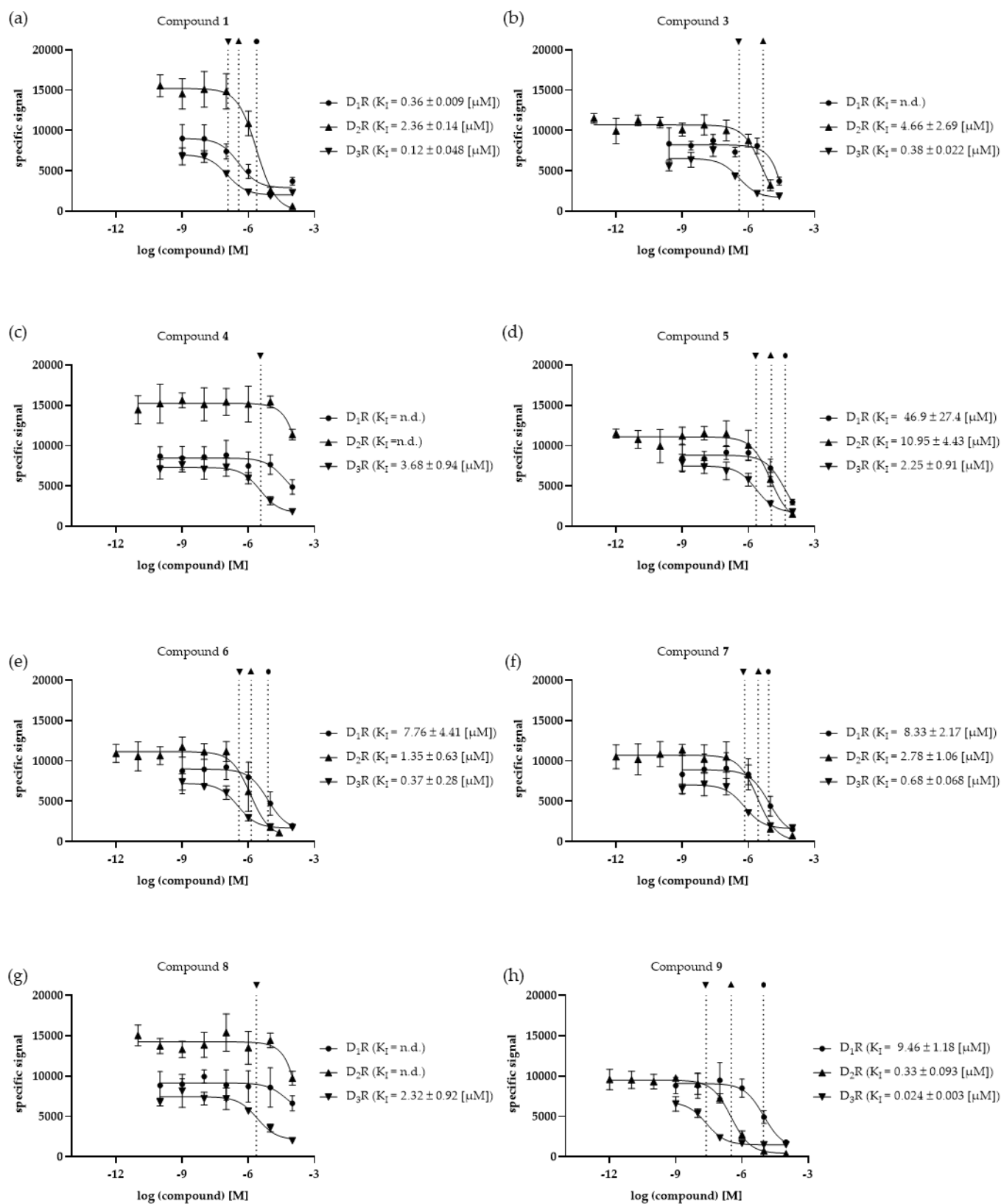


Figure S10. Comparison of the K_1 values of compounds (a) 1, (b) 3, (c) 4, (d) 5, (e) 6, (f) 7, (g) 8 and (h) 9 determined at the investigated DR subtypes D₁R, D₂R and D₃R. Vertical, dotted lines indicate the respective K_1 values at the different DR subtypes. K_1 values [μM] ± SD were determined with n = 6.

5. In silico insights into the DR binding pockets

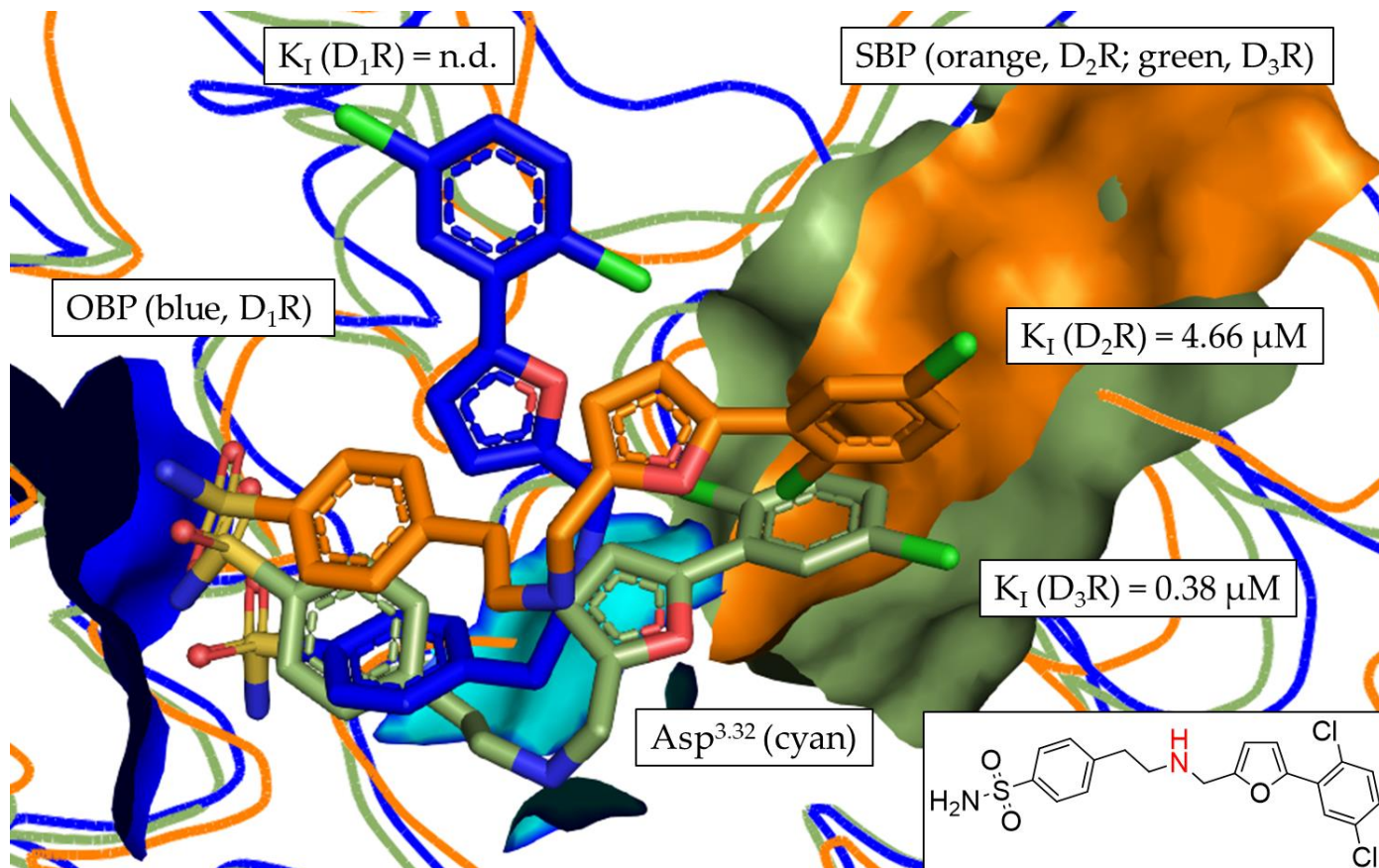


Figure S11. Alignment of the most frequent poses of compound **3** docked into D_1R , D_2R and D_3R . The surface of the highly conserved OBP is shown in blue (based on D_1R) consisting of Asp^{3.32} (highlighted in cyan) and Ser^{5.42/5.43/5.46}. The conserved SBP-surface is displayed in orange (D_2R) and green (D_3R) consisting of Val^{2.61}, Leu^{2.64}, Gly^{ECL1}, Phe^{3.28} and Cys^{ECL2} (individual amino acid labels shown in Table 3), respectively. K_I values determined in vitro are shown for each DR subtype. 2D structure of compound **3** is shown. Amine functional group involved in formation of the salt-bridge is highlighted in red.

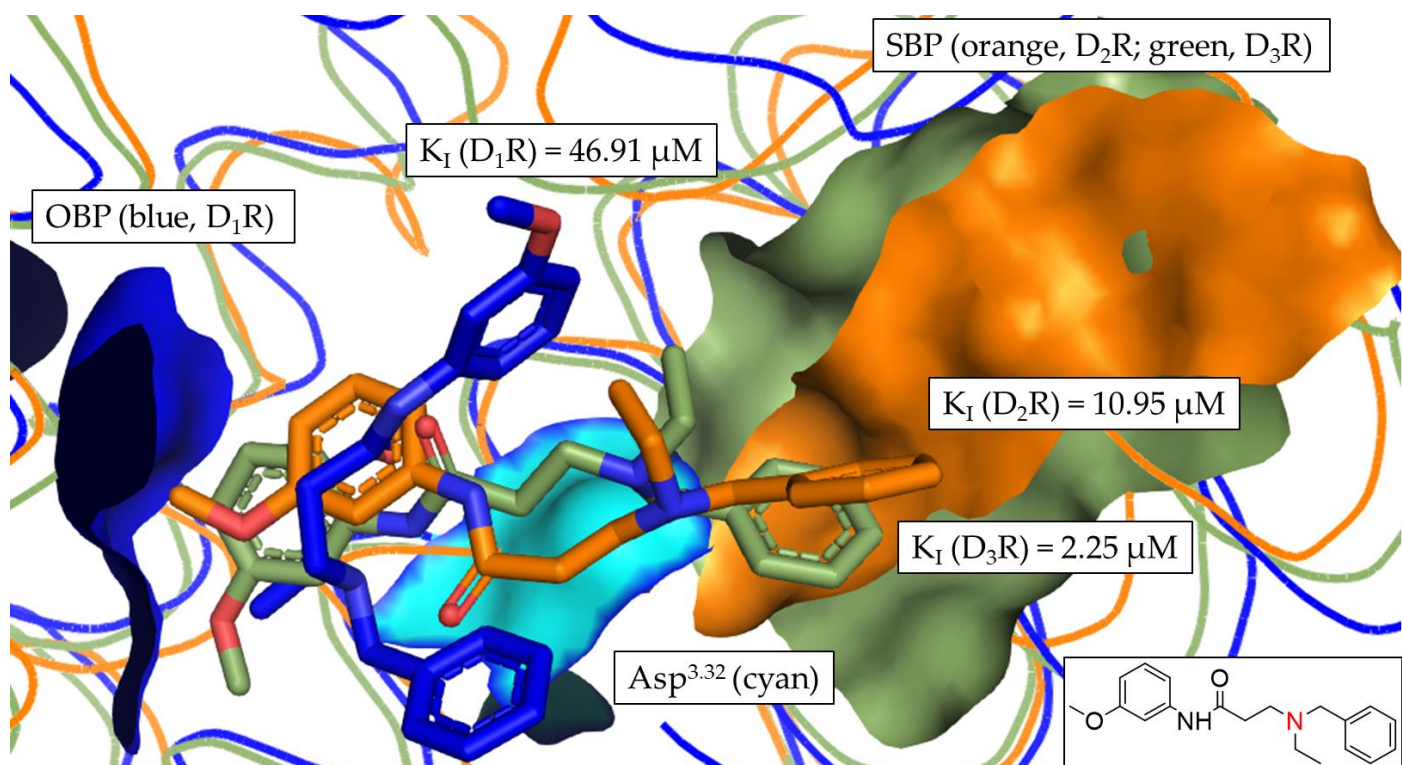


Figure S12. Alignment of the most frequent poses of compound 5 docked into D₁R, D₂R and D₃R. The surface of the highly conserved OBP is shown in blue (based on D₁R) consisting of Asp^{3.32} (highlighted in cyan) and Ser^{5.42/5.43/5.46}. The conserved SBP-surface is displayed in orange (D₂R) and green (D₃R) consisting of Val^{2.61}, Leu^{2.64}, Gly^{ECL1}, Phe^{3.28} and Cys^{ECL2} (individual amino acid labels shown in Table 3), respectively. K_I values determined in vitro are shown for each DR subtype. 2D structure of compound 5 is shown. Amine functional group involved in formation of the salt-bridge is highlighted in red.

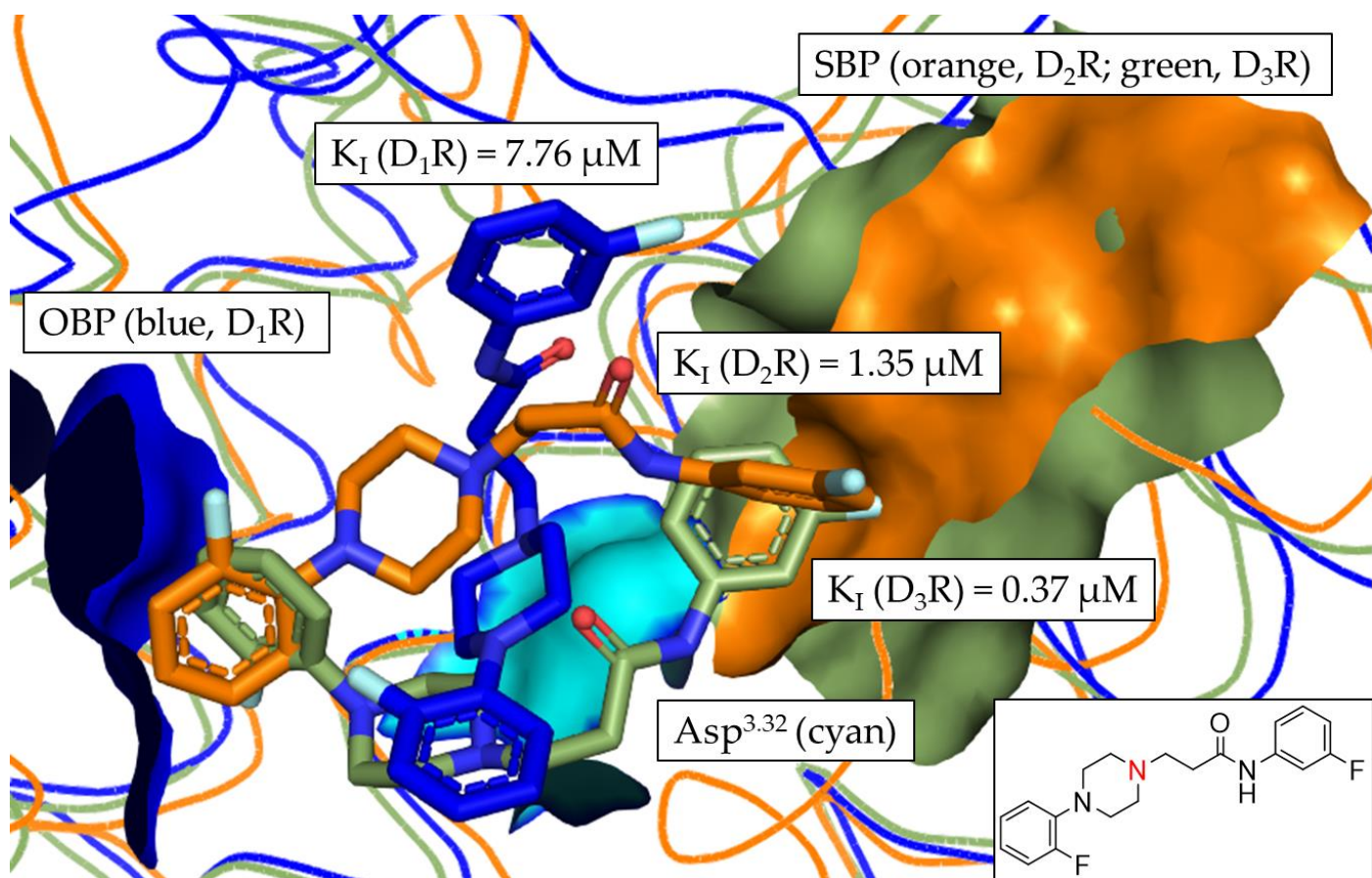


Figure S13. Alignment of the most frequent poses of compound 6 docked into D₁R, D₂R and D₃R. The surface of the highly conserved OBP is shown in blue (based on D₁R) consisting of Asp^{3.32} (highlighted in cyan) and Ser^{5.42/5.43/5.46}. The conserved SBP-surface is displayed in orange (D₂R) and green (D₃R) consisting of Val^{2.61}, Leu^{2.64}, Gly^{ECL1}, Phe^{3.28} and Cys^{ECL2} (individual amino acid labels shown in Table 3), respectively. K_I values determined in vitro are shown for each DR subtype. 2D structure of compound 6 is shown. Amine functional group involved in formation of the salt-bridge is highlighted in red.

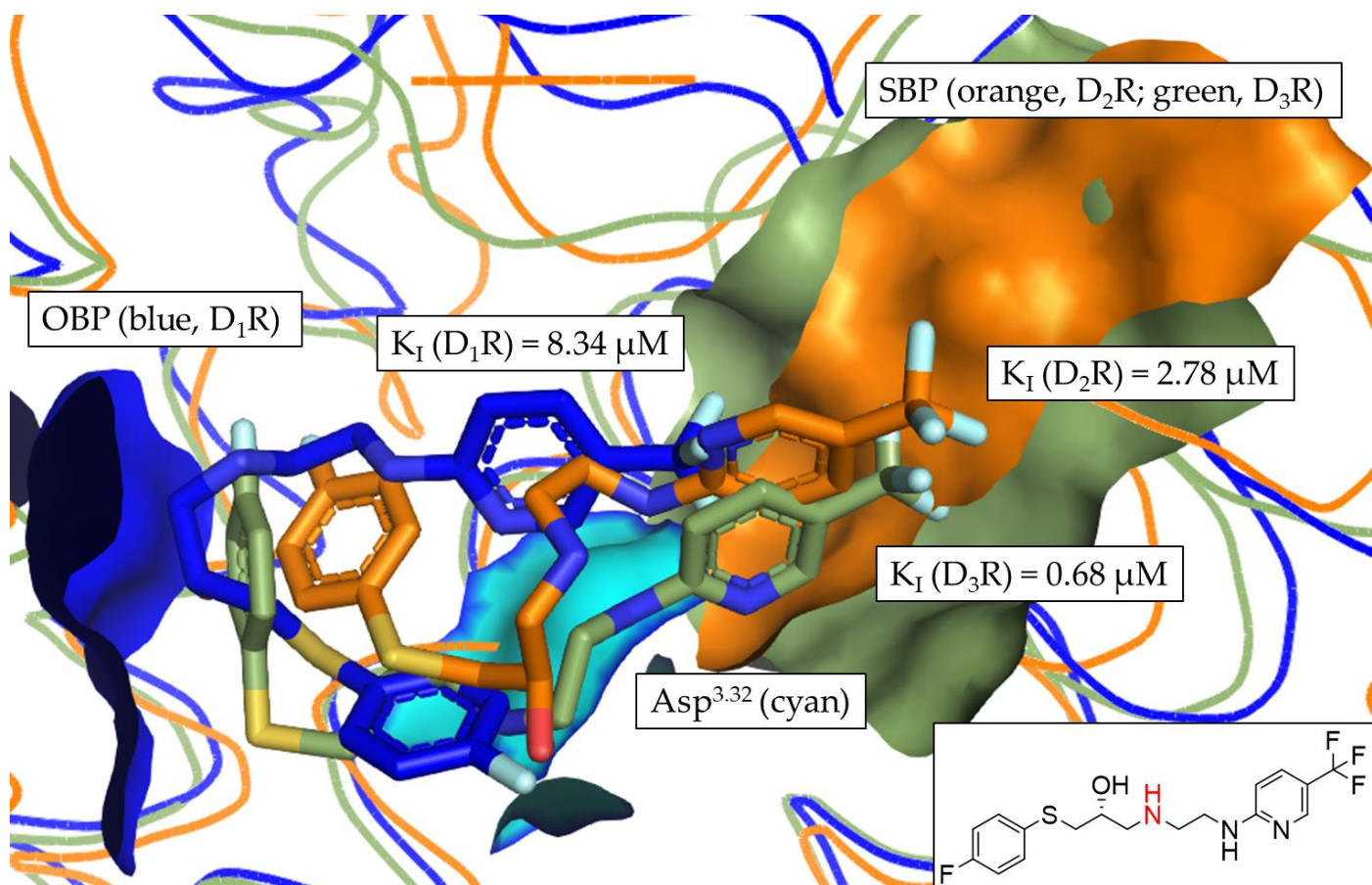


Figure S14. Alignment of the most frequent poses of compound **7** docked into D₁R, D₂R and D₃R. The surface of the highly conserved OBP is shown in blue (based on D₁R) consisting of Asp^{3.32} (highlighted in cyan) and Ser^{5.42/5.43/5.46}. The conserved SBP-surface is displayed in orange (D₂R) and green (D₃R) consisting of Val^{2.61}, Leu^{2.64}, Gly^{ECL1}, Phe^{3.28} and Cys^{ECL2} (individual amino acid labels shown in Table 3), respectively. K_I values determined in vitro are shown for each DR subtype. 2D structure of compound **7** is shown. Amine functional group involved in formation of the salt-bridge is highlighted in red.

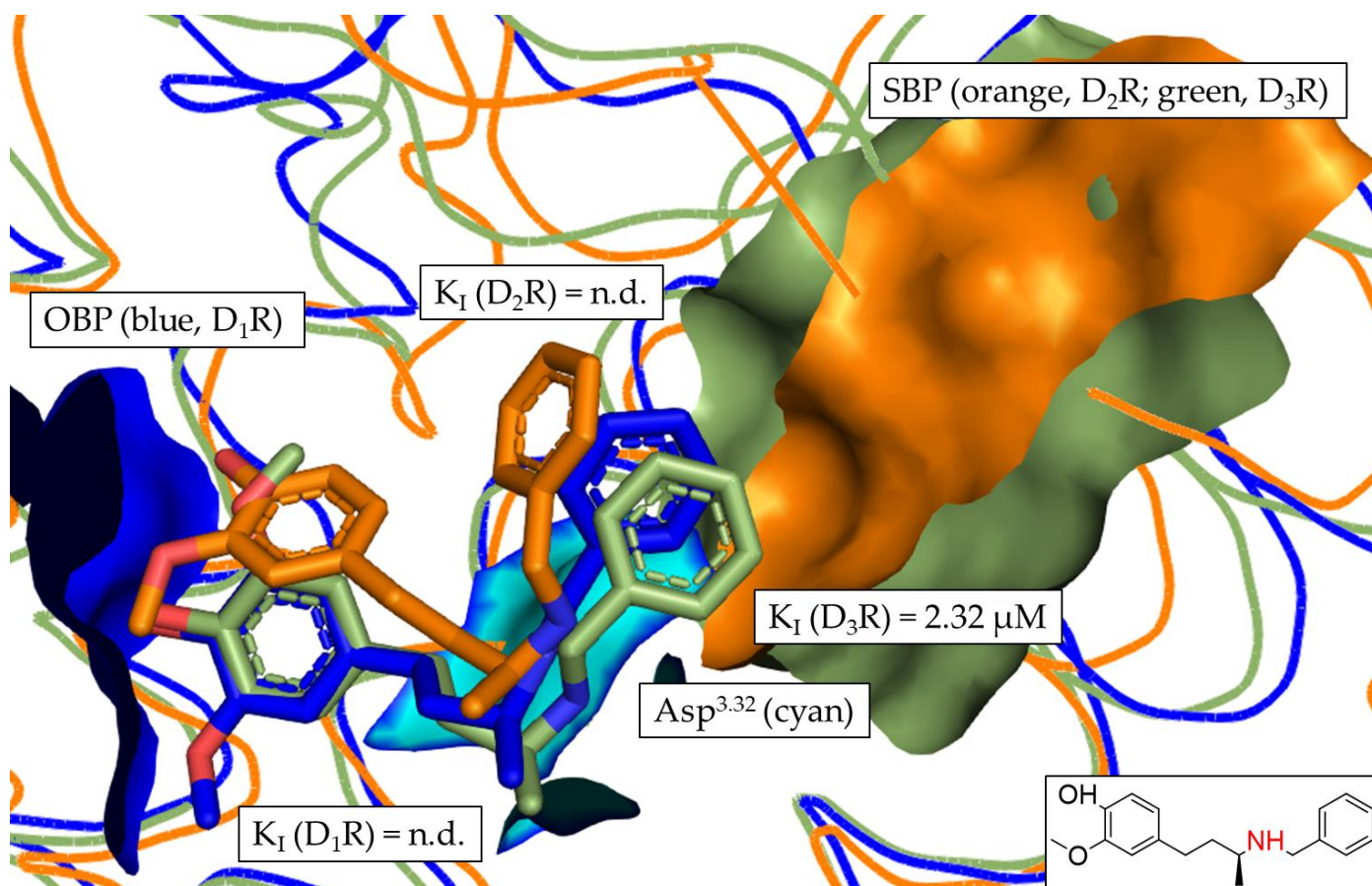


Figure S15. Alignment of the most frequent poses of compound **8** docked into D₁R, D₂R and D₃R. The surface of the highly conserved OBP is shown in blue (based on D₁R) consisting of Asp^{3.32} (highlighted in cyan) and Ser^{5.42/5.43/5.46}. The conserved SBP-surface is displayed in orange (D₂R) and green (D₃R) consisting of Val^{2.61}, Leu^{2.64}, Gly^{ECL1}, Phe^{3.28} and Cys^{ECL2} (individual amino acid labels shown in Table 3), respectively. K_I values determined in vitro are shown for each DR subtype. 2D structure of compound **8** is shown. Amine functional group involved in formation of the salt-bridge is highlighted in red.

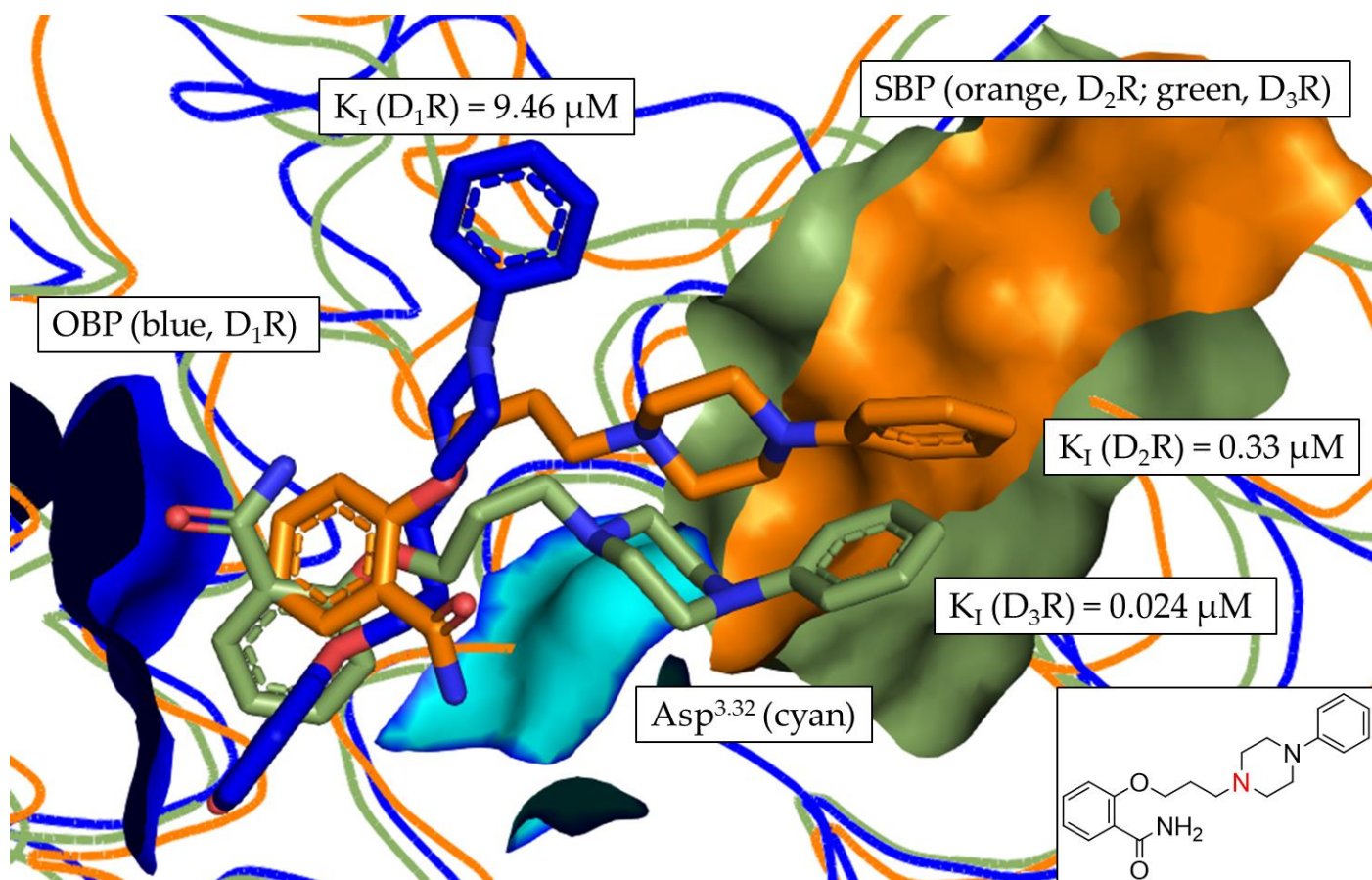


Figure S16. Alignment of the most frequent poses of compound **9** docked into D_1R , D_2R and D_3R . The surface of the highly conserved OBP is shown in blue (based on D_1R) consisting of Asp^{3.32} (highlighted in cyan) and Ser^{5.42/5.43/5.46}. The conserved SBP-surface is displayed in orange (D_2R) and green (D_3R) consisting of Val^{2.61}, Leu^{2.64}, Gly^{ECL1}, Phe^{3.28} and Cys^{ECL2} (individual amino acid labels shown in Table 3), respectively. K_I values determined in vitro are shown for each DR subtype. 2D structure of compound **9** is shown. Amine functional group involved in formation of the salt-bridge is highlighted in red.