

Supplementary Table 1. Summary of CB1R agonist-bound (5XRA) and antagonist-bound (5TGZ) amino acid residues interacting with cannabinoid ligands in AutoDock.

HELIX	CP55,940		2-AG		THC		CBD		CBD-DMH		SR141716A		Org27569	
	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ
I		MM ¹⁰³		MM ¹⁰³		MM ¹⁰³						MM ¹⁰³		
											CM ¹⁰⁷			
								MM ¹⁰⁹						
								Q1.31 ¹¹⁵						Q1.31 ¹¹⁵
								Q1.32 ¹¹⁶				Q1.32 ¹¹⁶		Q1.32 ¹¹⁶
								I1.35 ¹¹⁹						
II	F2.57 ¹⁷⁰	F2.57 ¹⁷⁰	F2.57 ¹⁷⁰		F2.57 ¹⁷⁰		F2.57 ¹⁷⁰			F2.57 ¹⁷⁰	F2.57 ¹⁷⁰	F2.57 ¹⁷⁰	F2.57 ¹⁷⁰	
	S2.60 ¹⁷³		S2.60 ¹⁷³											
					F2.61 ¹⁷⁴						F2.61 ¹⁷⁴	F2.61 ¹⁷⁴	F2.61 ¹⁷⁴	F2.61 ¹⁷⁴
	F2.64 ¹⁷⁷		F2.64 ¹⁷⁷		F2.64 ¹⁷⁷		F2.64 ¹⁷⁷		F2.64 ¹⁷⁷		F2.64 ¹⁷⁷		F2.64 ¹⁷⁷	
								H2.65 ¹⁷⁸						H2.65 ¹⁷⁸
III	L3.29 ¹⁹³		L3.29 ¹⁹³		L3.29 ¹⁹³				L3.29 ¹⁹³	L3.29 ¹⁹³	L3.29 ¹⁹³		L3.29 ¹⁹³	
		T3.33 ¹⁹⁷	T3.33 ¹⁹⁷	T3.33 ¹⁹⁷	T3.33 ¹⁹⁷	T3.33 ¹⁹⁷				T3.33 ¹⁹⁷				
IV	F4.77 ²⁶⁸			F4.77 ²⁶⁸		F4.77 ²⁶⁸	F4.77 ²⁶⁸				F4.77 ²⁶⁸			
V	W5.43 ²⁷⁹		W5.43 ²⁷⁹		W5.43 ²⁷⁹				W5.43 ²⁷⁹		W5.43 ²⁷⁹		W5.43 ²⁷⁹	
VI			L6.51 ³⁵⁹				L6.51 ³⁵⁹		L6.51 ³⁵⁹		L6.51 ³⁵⁹			
		M6.55 ³⁶³		M6.55 ³⁶³		M6.55 ³⁶³				M6.55 ³⁶³				
VII	F7.34 ³⁷⁹		F7.34 ³⁷⁹		F7.34 ³⁷⁹		F7.34 ³⁷⁹		F7.34 ³⁷⁹		F7.34 ³⁷⁹	A7.36 ³⁸⁰		A7.36 ³⁸⁰
	S7.39 ³⁸³	S7.39 ³⁸³	S7.39 ³⁸³	S7.39 ³⁸³	S7.39 ³⁸³	S7.39 ³⁸³				S7.39 ³⁸³	S7.39 ³⁸³			

Supplementary Table 2. Summary of modeled CB2R agonist-bound (5XRA) and antagonist-bound (5TGZ) amino acid residues interacting with cannabinoid ligands in AutoDock.

HELIX	CP55,940		2-AG		THC		CBD		CBD-DMH		SR144528	
	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ	5XRA	5TGZ
I		SM ¹⁹						SM ¹⁹				SM ¹⁹
							KM ²³	KM ²³				
										V1.35 ³⁶		
										C1.39 ⁴⁰		
II	F2.57 ⁸⁷		F2.57 ⁸⁷	F2.57 ⁸⁷	F2.57 ⁸⁷	F2.57 ⁸⁷						
	S2.60 ⁹⁰				S2.60 ⁹⁰				S2.60 ⁹⁰		S2.60 ⁹⁰	
	F2.61 ⁹¹						F2.61 ⁹¹		F2.61 ⁹¹			
							LECL2 ¹⁸²	LECL2 ¹⁸²				
			FECL2 ¹⁸³		FECL2 ¹⁸³		FECL2 ¹⁸³	FECL2 ¹⁸³			FECL2 ¹⁸³	
III	K3.28 ¹⁰⁹											
	T3.33 ¹¹⁴	T3.33 ¹¹⁴	T3.33 ¹¹⁴	T3.33 ¹¹⁴	T3.33 ¹¹⁴	T3.33 ¹¹⁴		T3.33 ¹¹⁴	T3.33 ¹¹⁴		T3.33 ¹¹⁴	T3.33 ¹¹⁴
IV												
V		W5.43 ¹⁹⁴		W5.43 ¹⁹⁴		W5.43 ¹⁹⁴						W5.43 ¹⁹⁴
VI		W6.48 ²⁵⁸		W6.48 ²⁵⁸		W6.48 ²⁵⁸						W6.48 ²⁵⁸
VII			K7.31 ²⁷⁸						K7.31 ²⁷⁸		K7.31 ²⁷⁸	
			A7.36 ²⁸²		A7.36 ²⁸²		A7.36 ²⁸²		A7.36 ²⁸²	A7.36 ²⁸²	A7.36 ²⁸²	
										A7.39 ²⁸⁵		
										M7.39 ²⁸⁶		
		L7.43 ²⁸⁹		L7.43 ²⁸⁹		L7.43 ²⁸⁹						L7.43 ²⁸⁹