Table S1. qHTS protocol

Step	Parameter	Value	Description
1	Reagent	2 μl	Dispense HEK293-ACE2 cells at 2000 cells/well in media
2	Time	16 - 24 hr	Incubate at standard cell culture conditions
3a	Reagent	23 nl	Dispense library compounds to columns 5-48
3b	Reagent	23 nl	Dispense DMSO control to columns 1-4
4	Time	1 hr	Incubate at standard cell culture conditions
5a	Reagent	2 μl	Dispense bald PP in columns 1-2
5b	Reagent	2 μl	Dispense SARS-CoV-2-S PP in columns 3-48
6	Centrifugation	45 min	Spinoculation at 453 xg
7	Time	48 hr	Incubate at standard cell culture conditions
8	Aspiration	-4 μl	Supernatant removal via centrifugation
9	Reagent	4 μl	Dispense luciferase detection reagent
10	Time	5 min	Incubate at room temperature
11	Detector		Luminescence read

Step	Protocol details
1	Dispense with Multidrop. Growth media: DMEM 10% FBS, 1 mg/ml puromycin
2	Overnight incubation at 37 oC, 5% CO2
3	Dispense via pintool transfer. Compounds in DMSO.
4	Incubate at 37 oC, 5% CO2
5	Dispense with Multidrop.
6	Spinoculation at 1500 rpm (453 xg) at room temperature.
7	Incubate at 37 oC, 5% CO2
8	Blue Washer (BlueCat Bio) gentle spin setting.
9	Dispense Bright-Glo Luciferase detection reagent (Promega) with BioRapr
10	Incubate at room temperature.
11	PHERAStar plate reader (BMG Labtech) luminescence setting.

Table S2. Public datasets. Primary screen and follow up assay data can be found at https://pubchem.ncbi.nlm.nih.gov/ under the following assay IDs (AIDs).

AID	# of Compounds	Concentration response format	Assay
1645846	5,158	4-pt, 1:5	SARS-CoV-2-S PP entry
1645845	5,158	4-pt, 1:5	Cytotoxicity
1645844	382	11-pt, 1:3	SARS-CoV-2-S PP entry
1645847	382	11-pt, 1:3	VSV-G PP entry