

Figure 1—figure supplement 1

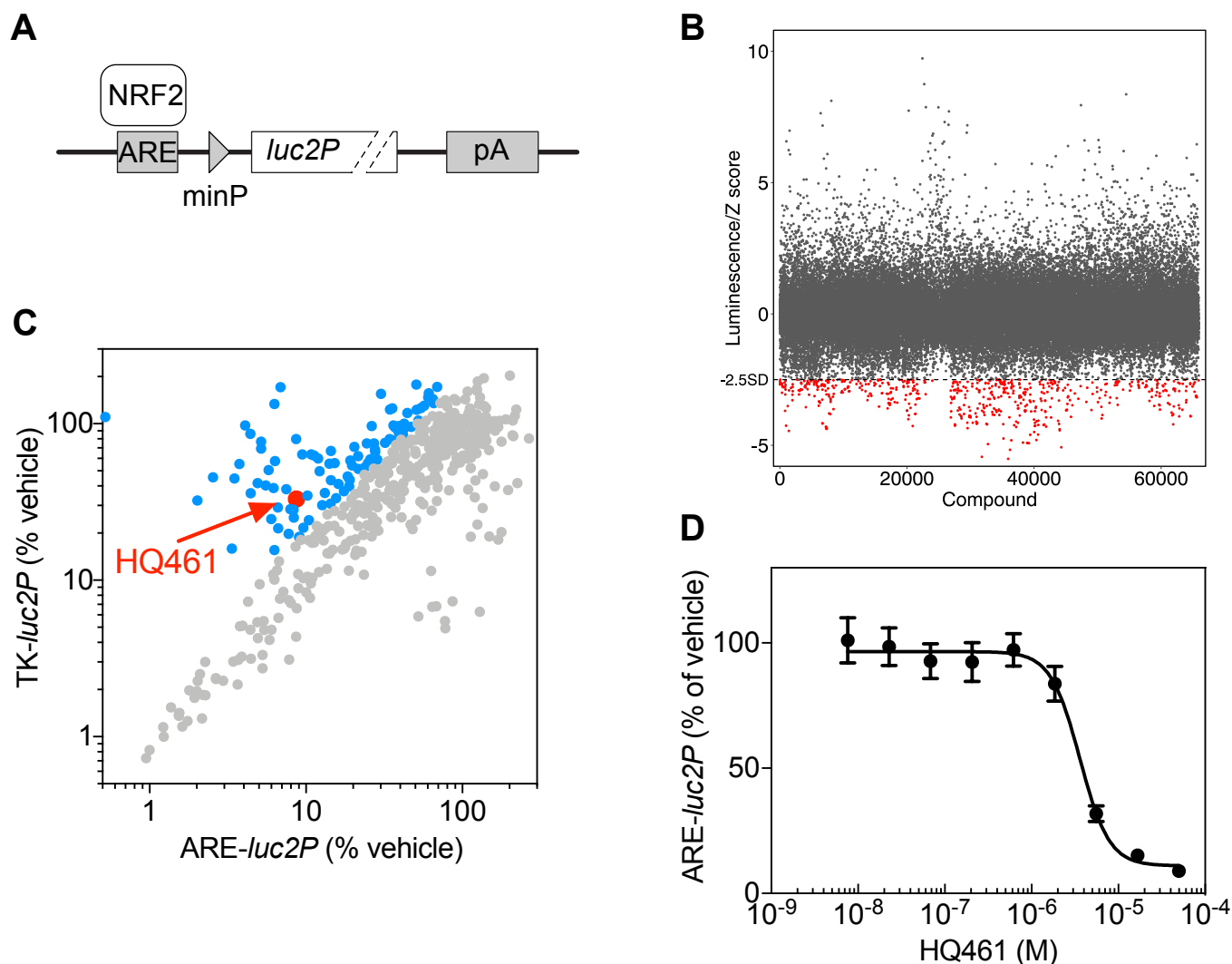


Figure 1—figure supplement 1. High-throughput chemical screening of NRF2 inhibitors.

(A) ARE-*luc2P* reporter used for high-throughput screening. NRF2 binds to anti-oxidant response elements (ARE) to drive luciferase (*luc2P*) expression.

(B) Primary screen results. The ARE-*luc2P* reporter cell line was treated for 12 hours with library compounds at 10 μ M. Each screening plate was individually Z-score normalized, and all the Z-scores were aggregated. The dotted line represents the cutoff (Z-score < -2.5). Red dots mark hits from the primary screen.

(C) Counter screen results. Hits from the primary screen were tested in both ARE-*luc2P* and TK-*luc2P* reporter cell lines at 10 μ M for 12 hours. Blue dots represent compounds selectively inhibiting ARE-*luc2P* relative to TK-*luc2P*.

(D) Measurement of the HQ461 IC_{50} on ARE-*luc2P* activity (IC_{50} =3.6 μ M, 95% CI: 2.9 μ M-4.5 μ M). Error bars represent standard errors of the mean (SEM) from three biological replicates.