

Figure Legends for supplementary data

Supplementary Figure S1. Growth inhibitory activity of WZ4002 and erlotinib on NSCLC cell lines harboring EGFR mutations. PC-9/Vec (A), PC-9/HGF#4 (B) and PC-9/HGF#5 (C), HCC827 (D), HCC827ER (E) and H1975 (F) cells (2×10^3 cells/well) were incubated with various concentrations of WZ4002 or erlotinib for 72 h. Cell growth was determined by the MTT assay.

Supplementary Figure S2. E7050 inhibits tumor cell motility induced by HGF overexpression. PC-9/Vec and PC-9/HGF#5 (4×10^5 cells) were seeded in the upper wells of 8 μ m Transwell chambers. Medium containing 50 μ g/mL fibronectin were added to the lower wells, and the cells were cultured for 24 h with or without WZ4002 0.3 μ M or E7050 0.3 μ M. Cells migrating to the lower side of the membrane were stained and calculated. Mean \pm SD *P<0.05.

Supplementary Figure S3. Sustained inhibition of Akt and Erk phosphorylation by combining WZ4002 and E7050. PC-9/HGF#5 cells were incubated with WZ4002 (0.3 μ M) and/or E7050 (0.3 μ M) for the indicated times. The cell lysates were harvested and

the phosphorylation of the indicated proteins was determined by western blotting.

Supplementary Figure S4. Cytostatic effects of WZ4002 and E7050 treatment.

PC-9/Vec (A), PC-9/HGF#5 (B) and HCC827ER (C) cells were incubated with WZ4002 (0.3 μ M) and/or E7050 (0.3 μ M), and the numbers of cells were counted at each time point. (D) PC-9/HGF#5 cells were incubated with WZ4002 (0.3 μ M) and/or E7050 (0.3 μ M) for 72 h. The cell lysates were harvested and the phosphorylation of indicated proteins was determined by western blotting. As a positive control, cells were treated with paclitaxel 0.1 μ M to induce cleavage of PARP and caspase-3

Supplementary Figure S5. Relative body weight of mice during treatment with

WZ4002 and/or E7050. SCID mice bearing PC-9/Vec (A), PC-9/HGF#5 (B), H1975 (C), or HCC827ER (D) tumors were administered WZ4002 and/or E7050, 25mg/kg, once daily for 14-21 days, and their body weights were measured every 3 days.

Supplementary Figure S6. Treatment with WZ4002 and/or E7050 does not induce

apoptosis *in vivo*. Representative TUNEL assay images of PC-9/HGF#5 and HCC827ER tumors treated with WZ4002 and/or E7050. The red arrow, pointing to

green fluorescence, indicates apoptotic cells. Bar, 100 μm . Mean \pm SD

Supplementary Table. S1 IC₅₀ values of each compound in MTT assay