

Supplementary Table 1. Modified cell lines together with their corresponding wild-type were screened for expression of aromatase mRNA and aromatase activity.

Supplementary Figure 1. Chemical structure of A. letrozole and B. PTK/ZK

Supplementary Figure 2. PTK/ZK inhibits the proliferation and ER/ERE mediated transcription of MCF7 AROM 2A cells. **A.** Proliferation. MCF7 AROM 2A [AROM+/ER+] cells were treated with vehicle (0.01% (v/v) ethanol) or androstenedione (10 nM) in the presence or absence of Log₁₀M increasing concentrations of PTK/ZK. After 6 days of treatment cell number was established using a coulter counter. **B.** ER/ERE mediated transcription. MCF7 AROM 2A cells co-transfected with EREIItkLuc and pCH110 were treated with vehicle (0.01% (v/v) ethanol), androstenedione (10 nM) or estradiol (1 nM) and increasing concentrations of PTK/ZK. Luciferase activity was normalised by β-galactosidase from co-transfected pCH110. Data is expressed as fold change compared to the vehicle treated control. Bars represent ± SEM. Effects were confirmed in three independent experiments. * p<0.01 compared with controls.

Supplementary Figure 3. PTK/ZK does not affect E2 driven proliferation or ER/ERE mediated transcription in MCF7 AROM 2A cells.

A. Proliferation. MCF7 AROM 2A [AROM+/ER+] cells were treated with vehicle (0.01% (v/v) ethanol) or estradiol (1 nM) in the presence or absence of Log₁₀M increasing concentrations of PTK/ZK. After 6 days of treatment, cell number was established using a Coulter counter. Data is expressed as fold change compared to the vehicle treated control. **B.** ER/ERE mediated transcription. MCF7 AROM 2A cells

co-transfected with EREIItkLuc and pCH110 were treated with vehicle (0.01% (v/v) ethanol), androstenedione (10 nM) or estradiol (1 nM) and increasing concentrations of PTK/ZK. Luciferase activity was normalised by β -galactosidase from co-transfected pCH110. Normalized luciferase activity from triplicate wells was expressed relative to the vehicle treated control. Bars represent \pm SEM. Effects were confirmed in three independent experiments.

Supplementary Figure 4. Comparison of the IC₅₀ for PTK/ZK versus letrozole and anastrozole.

Microsomes were isolated from MCF7 AROM [AROM+/ER+] cells. Microsomes were treated in triplicate with androst-4-ene-3,17-dione, [1 β ³H(N)] (0.125 μ M) in the presence of vehicle (ethanol 0.01% (v/v)), increasing concentrations of **A.** letrozole (LET), **B.** anastrozole or **C.** PTK/ZK. Aromatase activity was measured by the conversion of androstenedione to tritiated water and expressed relative to the vehicle control.