OPPOSITE EFFECTS OF ESTROGEN RECEPTOR ON *PHLDA1* AND *PAWR* GENE EXPRESSION IN BREAST CANCER CELLS

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The estrogen receptors (ER), when activated by estrogens or by diverse signaling pathways, act as transcription factors binding directly to the estrogen response elements (ERE) or to other transcription factors. The pro-apoptotic genes, PHDLA1 and PAWR, show potential consensus sequences, in their promoter regions, that can be recognized by the ER. Therefore, our goal was to investigate the ER effects on the expression of these genes in the breast cancer cells MCF-7. The cells were grown in striped serum and treated with 10nM 17β estradiol (E2) or 1µM fulvestrant (ICI-182.780) for 2, 6 and 24 hours, and the relative expression of the target genes was analyzed by real time RT-PCR. The cells maintained in striped serum did not show altered PHLDA1 expression, but showed 2-fold increase in PAWR expression. The cells treated with E2 showed 2.5-fold increase in PHLDA1 expression after 6 hours, and 2-fold decrease in PAWR expression after 24 hours, compared to the control cells, However, after 2 and 24 hours of ICI treatment, respectively, the cells showed 2-fold decrease in PHLDA1 expression and 2.5-fold increase in PAWR expression, compared to the control ones. These results indicate that ER activation, both directly and indirectly, induces PHLDA1 and inhibits PAWR expression. Supported by FAPESP.