

# Accepted Manuscript

Design and synthesis of indoline thiohydantoin derivatives based on enzalutamide as antiproliferative agents against prostate cancer

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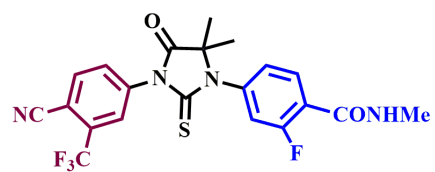
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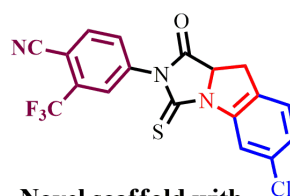
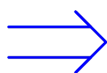


**Enzalutamide**

$IC_{50} = 12.5 \mu\text{M}$  (LNCaP cell)

$IC_{50} = 46.1 \mu\text{M}$  (DU145 cell)

Selectivity ratio = 3.7-fold

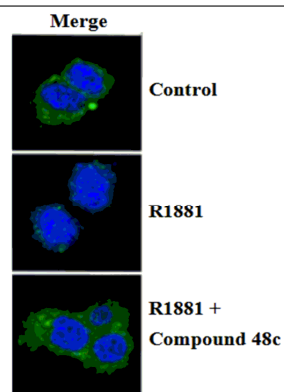


**Novel scaffold with rigid structure (48c)**

$IC_{50} = 27.9 \mu\text{M}$  (LNCaP cell)

$IC_{50} > 200 \mu\text{M}$  (DU145 cell)

Selectivity ratio  $> 7.2$ -fold



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