

## SUPPLEMENTARY INFORMATION

### Chemical space, diversity and activity landscape analysis of estrogen receptor binders

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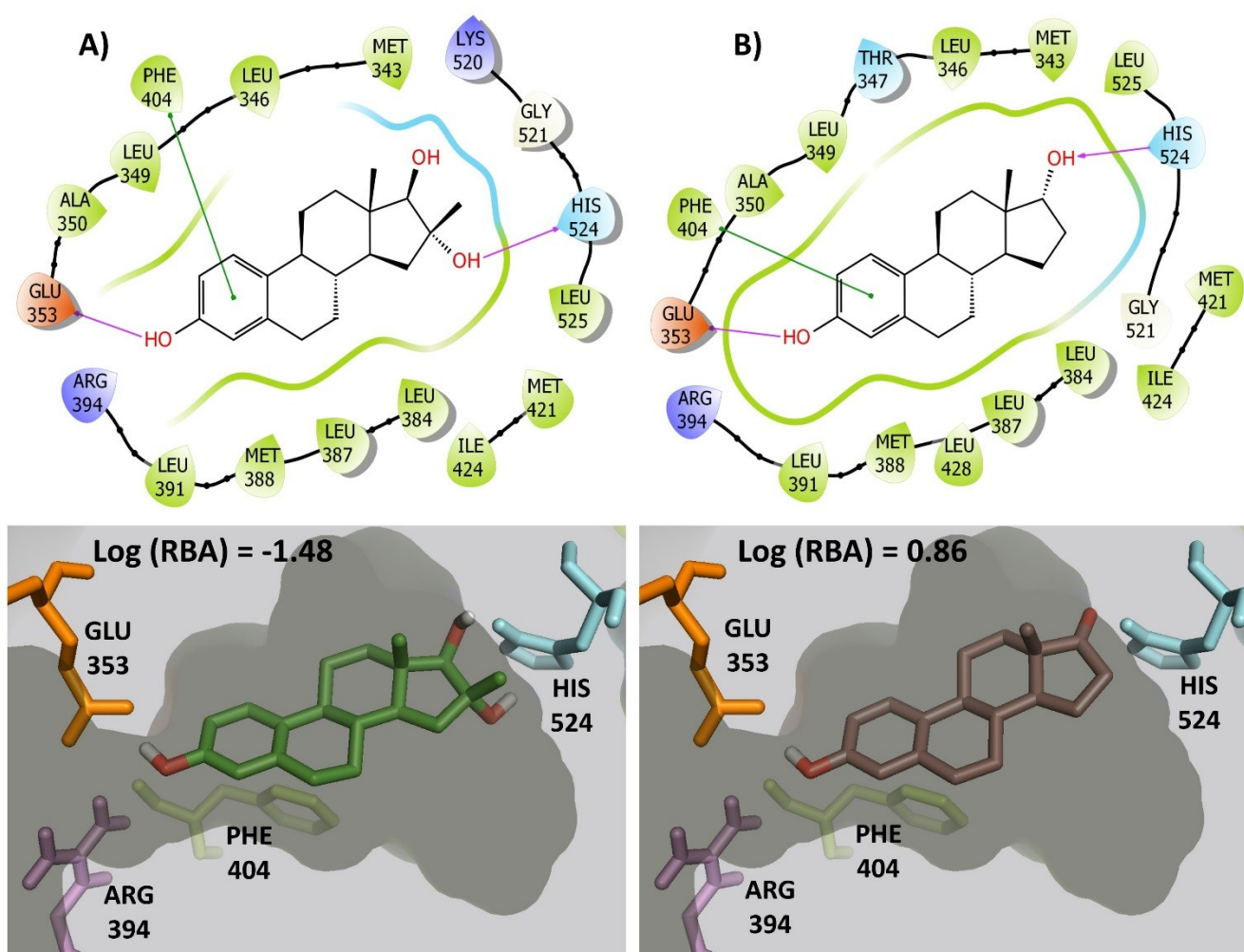
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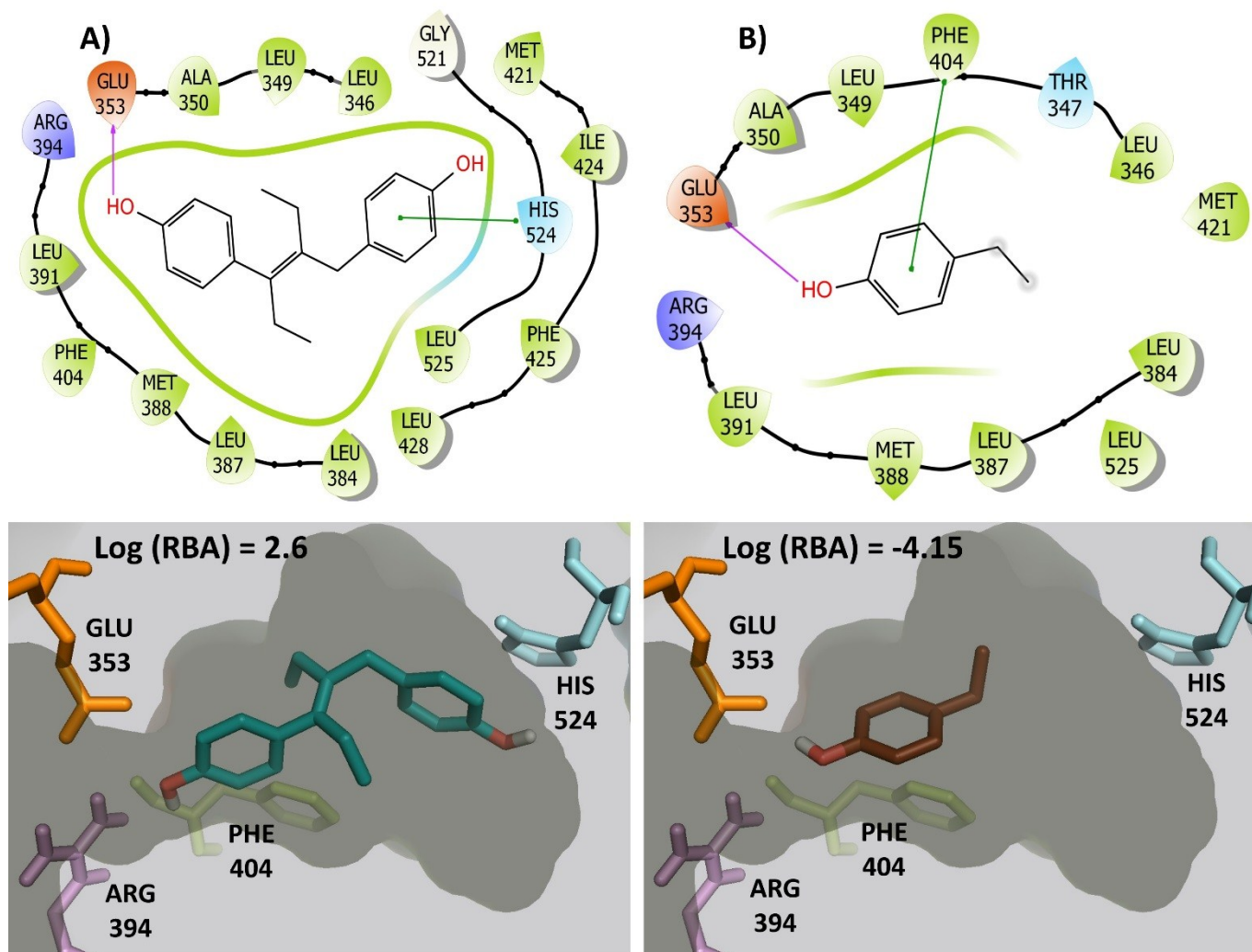
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**Figure S1.** 2D and 3D representation of representative activity cliff generators and selected pairs of compounds with greater difference in activity. **A)** 16beta-ol-16alfa-methyl-3-methyl-estradiol and **B)** estrone. The figure includes the value of the relative binding affinity (RBA) as reported by.<sup>14</sup>



**Figure S2.** 2D and 3D representation of representative activity cliff generators and selected pairs of compounds with greater difference in activity. **A)** diethylstilbestrol and **B)** 4-ethylphenol. The figure includes the value of the relative binding affinity (RBA) as reported by.<sup>14</sup>