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I am reading the updated background review documents (May 4) and have some comments for the committee on ER binding.

1. Why use 1 nM hot E2 when the Kd is an order of magnitude lower than this. This will increase the difficulty in detecting weaker estrogens. These assays generally should be conducted using conc of the radiolabelled ligand close to the Kd.
2. ER binding data on unknowns often result in bizarre shaped curves or only partial inhibition at high concentrations. For such data one must measure the Ki value experimentally rather than estimate it (as done by Kelce et al., 1994 for the antiandrogenic ligands M1 and M2) to assure that one is looking at competitive inhibition of E2 binding and not destabilization of the assay.

Questionable curves seen include 1) U-shaped, 2) declines from 100% to 0 is less than one log unit, 3) partial inhibition not reaching the IC50, 4) partial inhibition, leveling off near the IC50, 5) incomplete inhibition with values not attaining a value lower than 20% hot.

Good statistics section.