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Magn. Reson. Discuss., referee comment RC1
https://doi.org/10.5194/mr-2021-7-RC1, 2021

Comment on mr-2021-7

Walter Chazin (Referee)

Referee comment on "High-affinity tamoxifen analogues retain extensive positional disorder when bound to calmodulin" by Lilia Milanesi et al., Magn. Reson. Discuss., https://doi.org/10.5194/mr-2021-7-RC1, 2021

This manuscript presents an investigation of the interaction of calmodulin with the small molecule Tamoxifen using solution NMR. Interestingly, Tamoxifen was found to bind with high affinity without occupying specific hydrophobic pockets. This was clearly evident from the inability to satisfy inter-molecular NOEs by a single structure of the complex. The experimental approach is sound and the experiments are well designed, including several well thought out controls. This includes an important titration carried out to control for chemical shift perturbations that arise from CD_3OD. The data are properly interpreted. The manuscript is well written and clear. The manuscript is suitable for publication requiring only a few typographical/grammar adjustments.

- Line 184 – “were allowed”
- Line 266 – Remove “from:
- Line 395 – “have been”
- Line 420 – “was”

This review was performed primarily by postdoctoral fellow Randika (Randy) Perera, Ph.D.