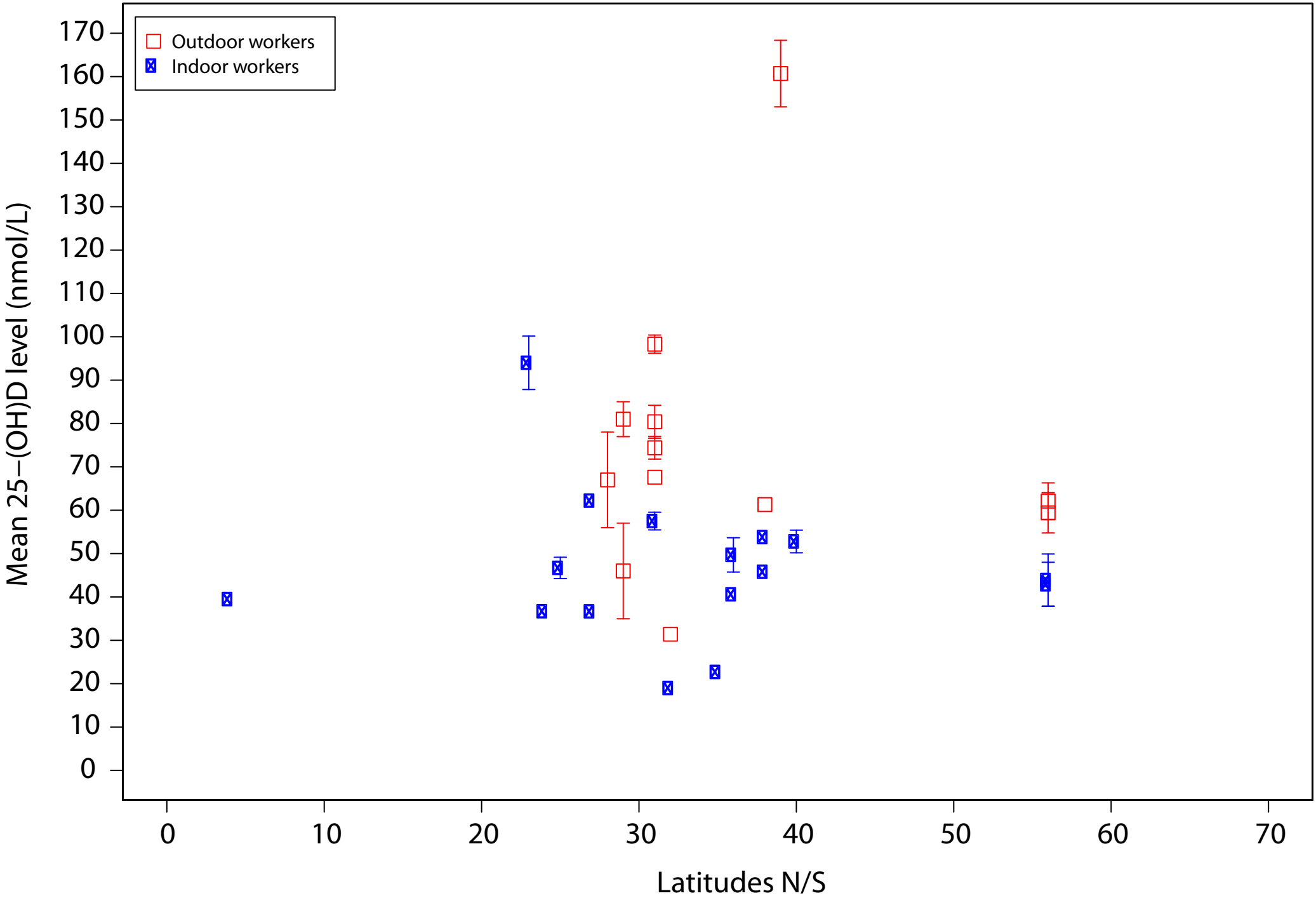Supplementary Fig. 4B