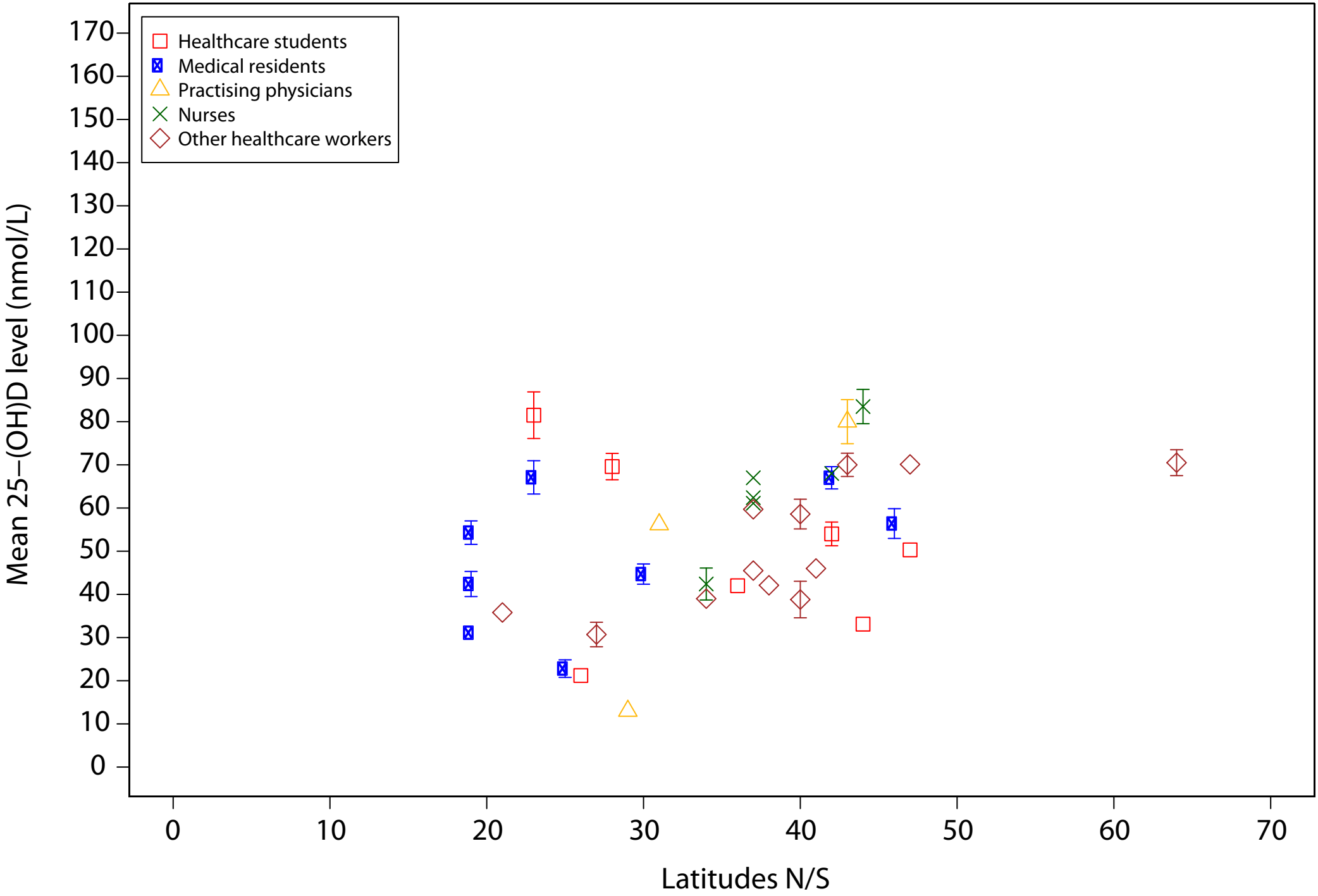Supplementary Fig. 4C