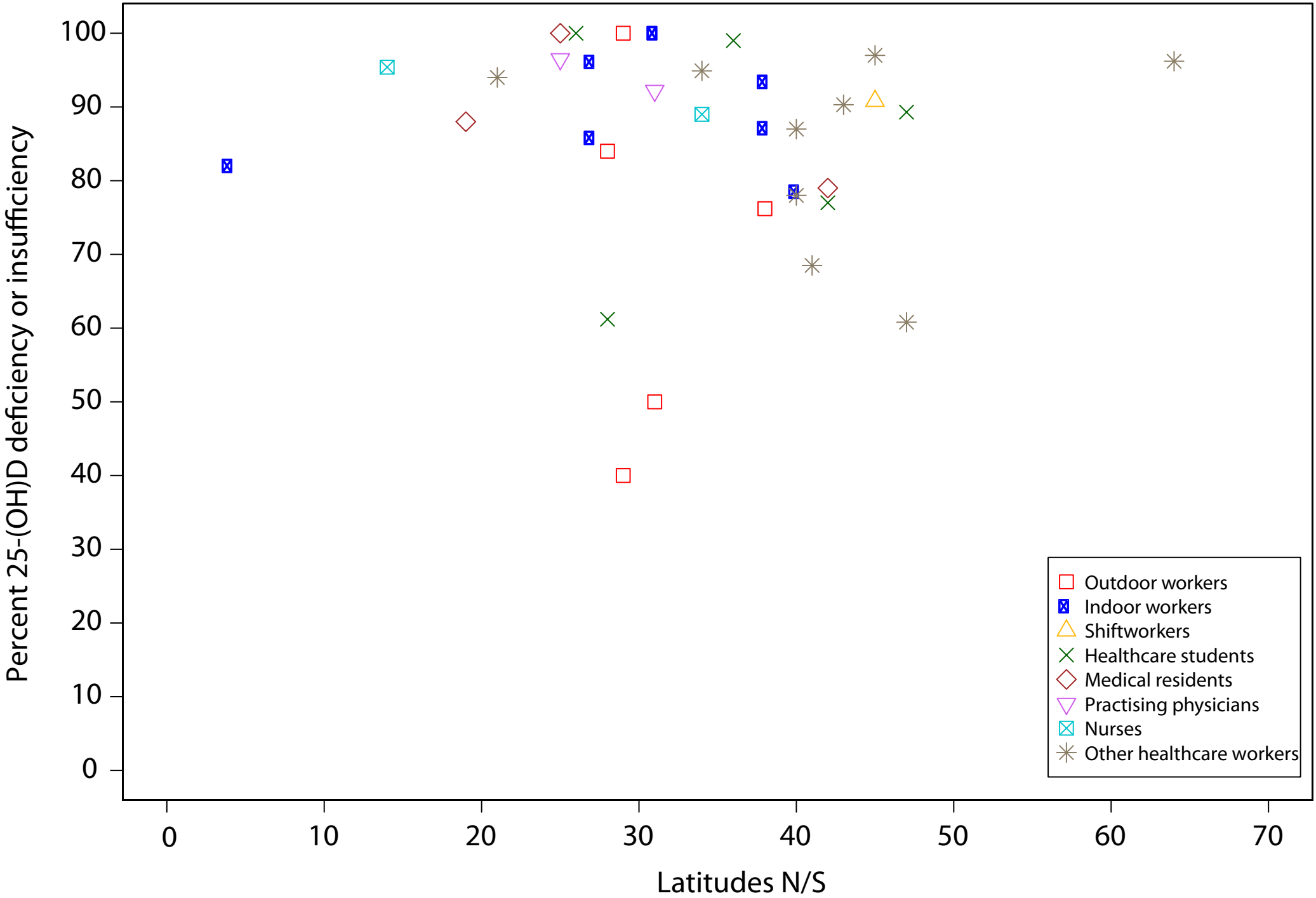Supplementary Fig. 6A