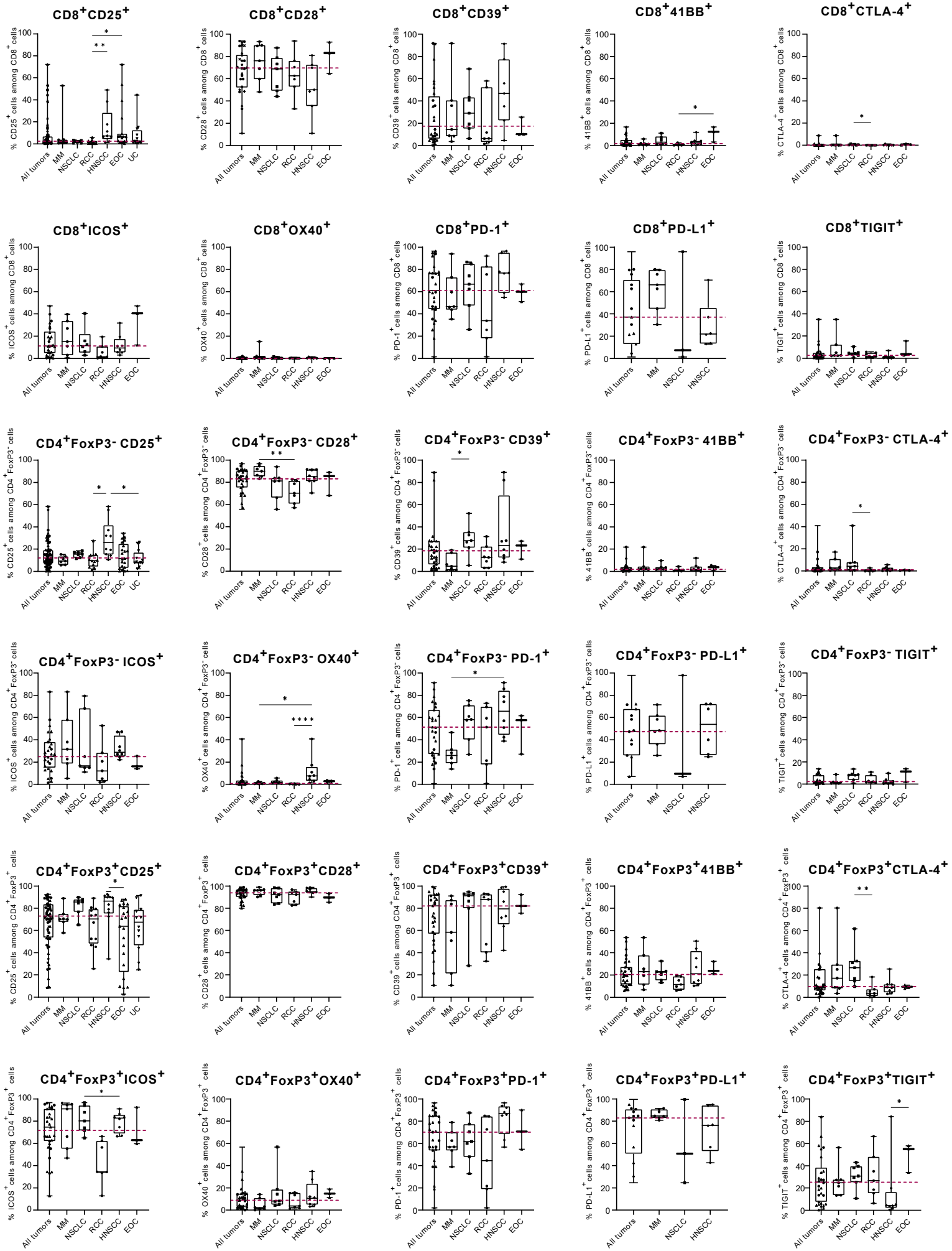Supplementary Data 6