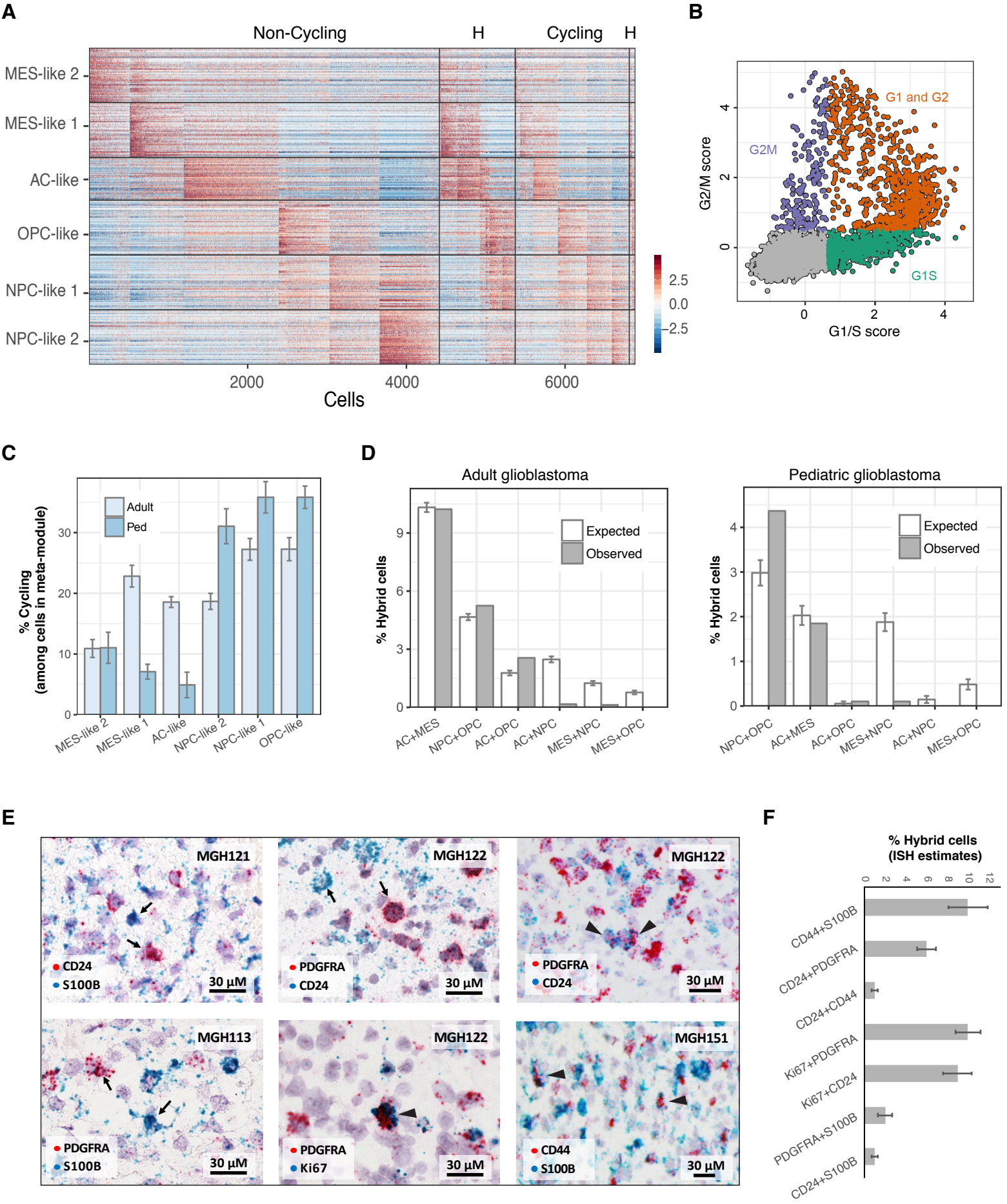Suppl. Figure 3