

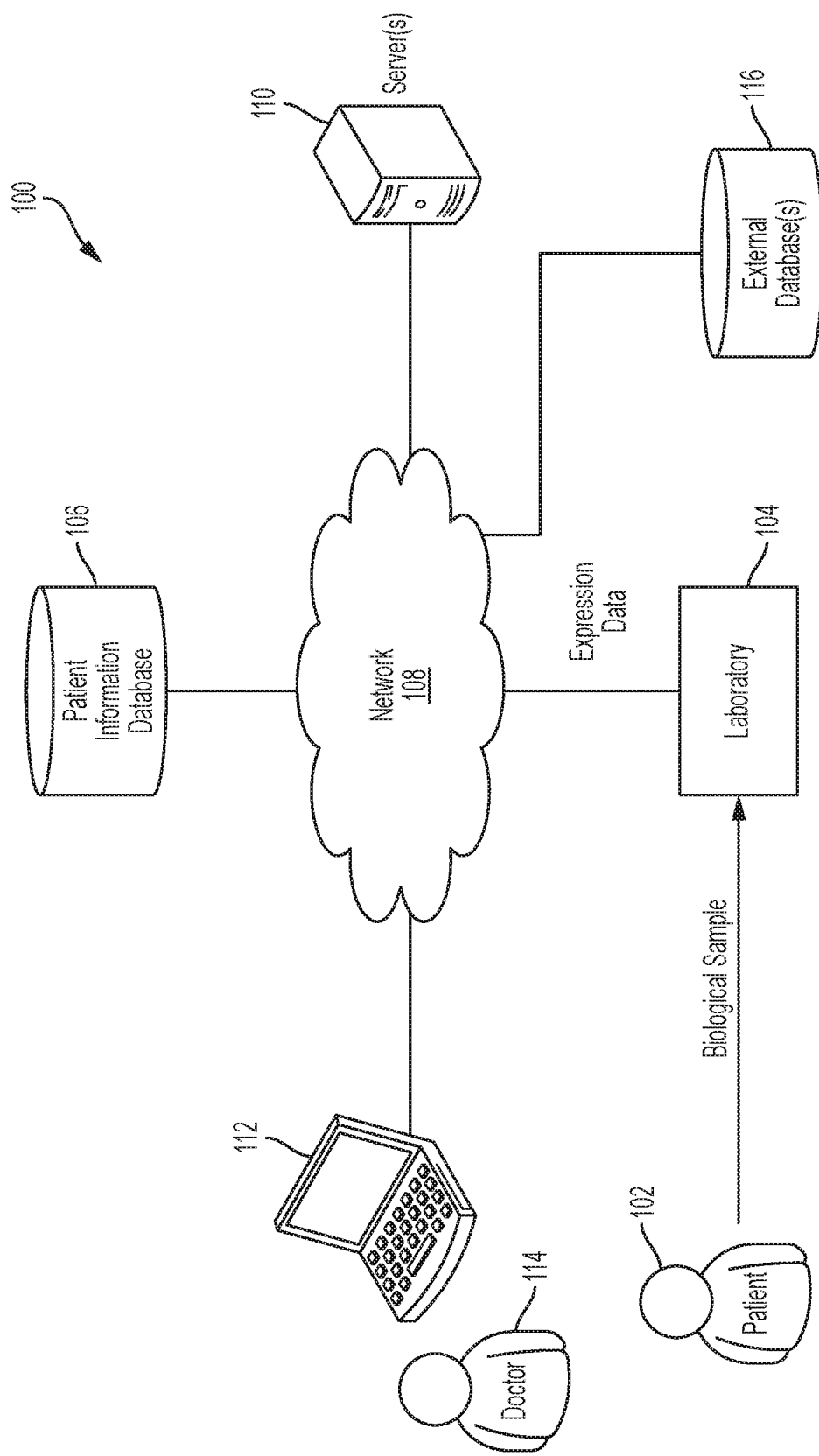


FIG. 1A

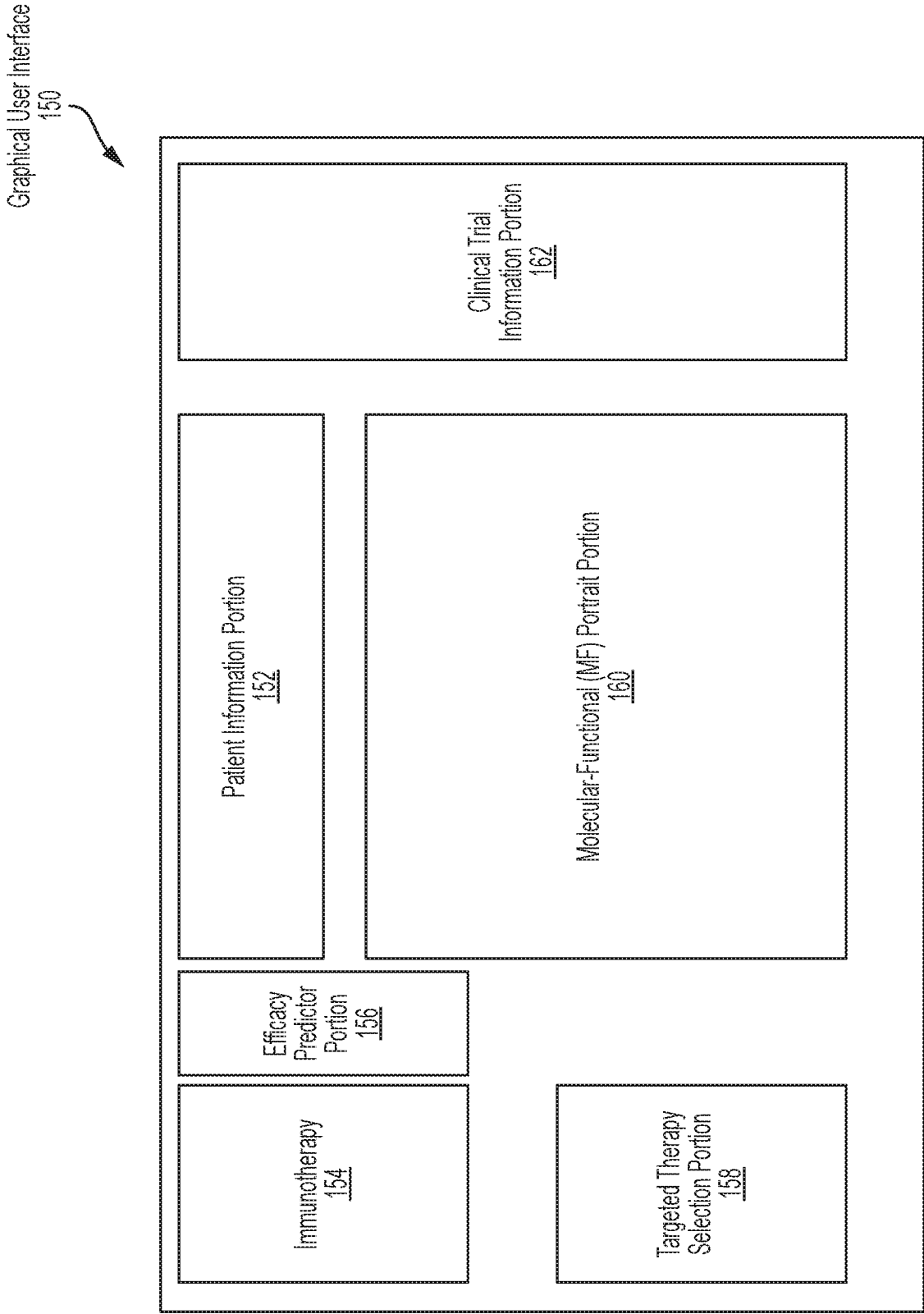
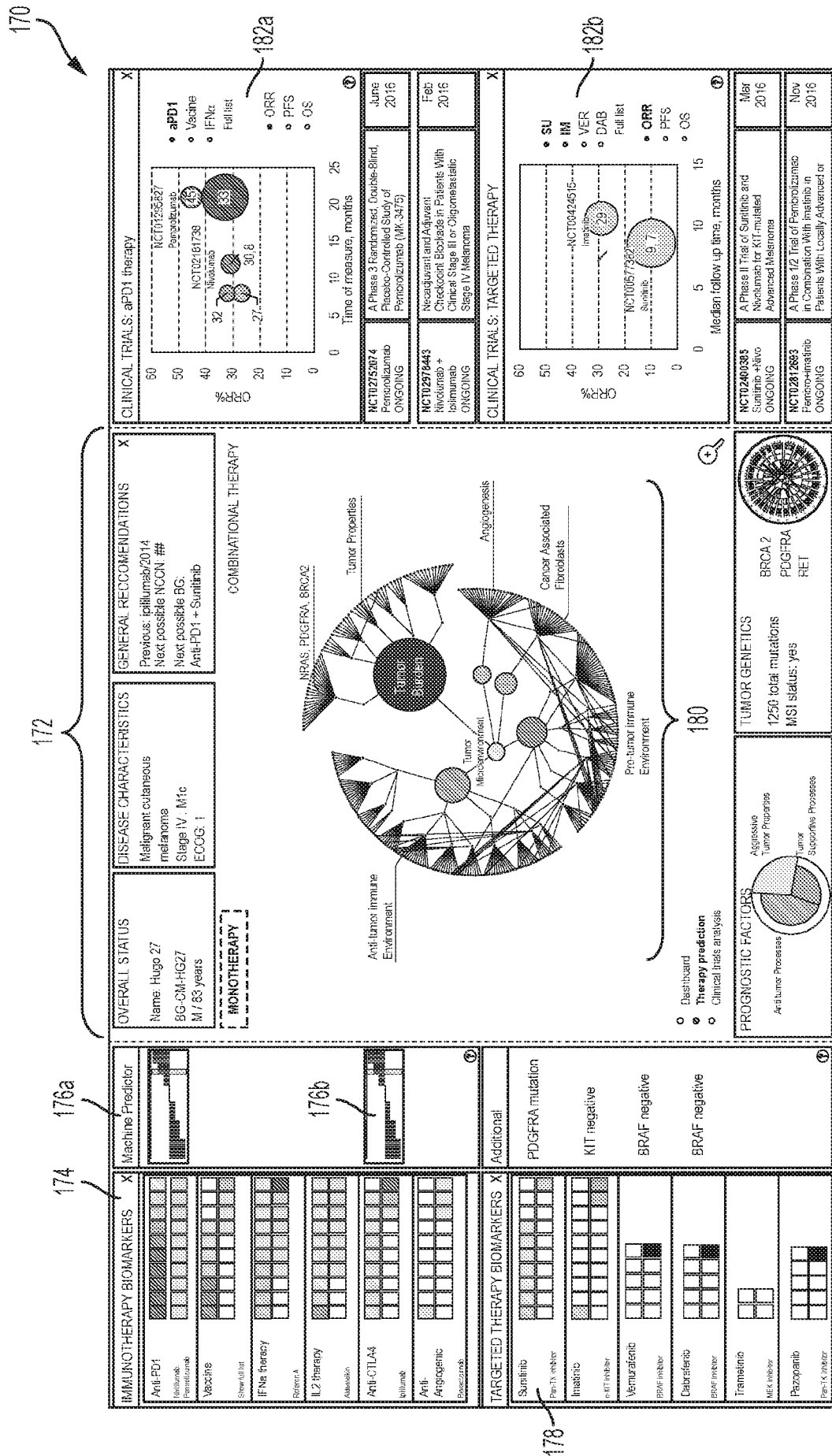


FIG. 1B



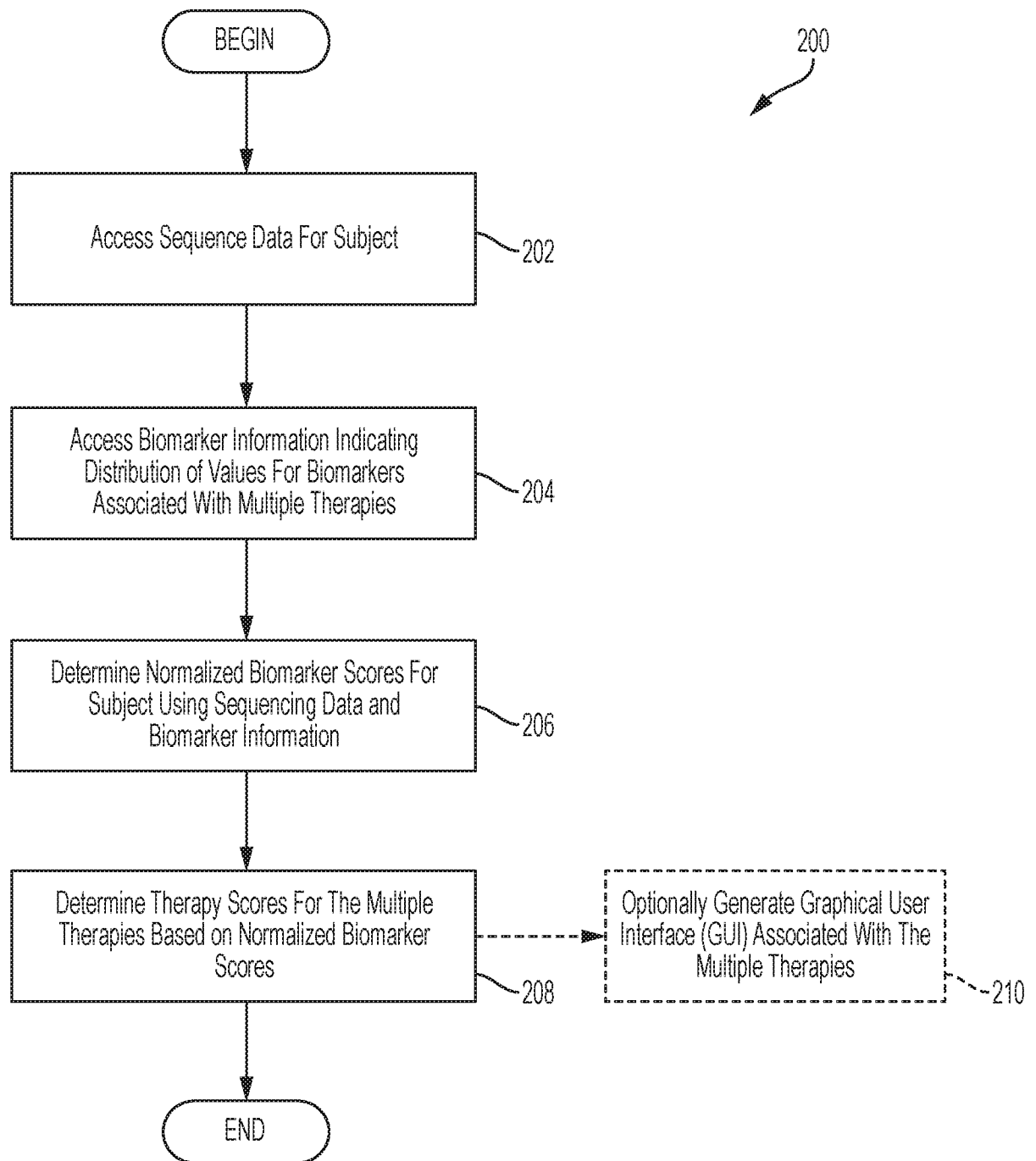


FIG. 2A

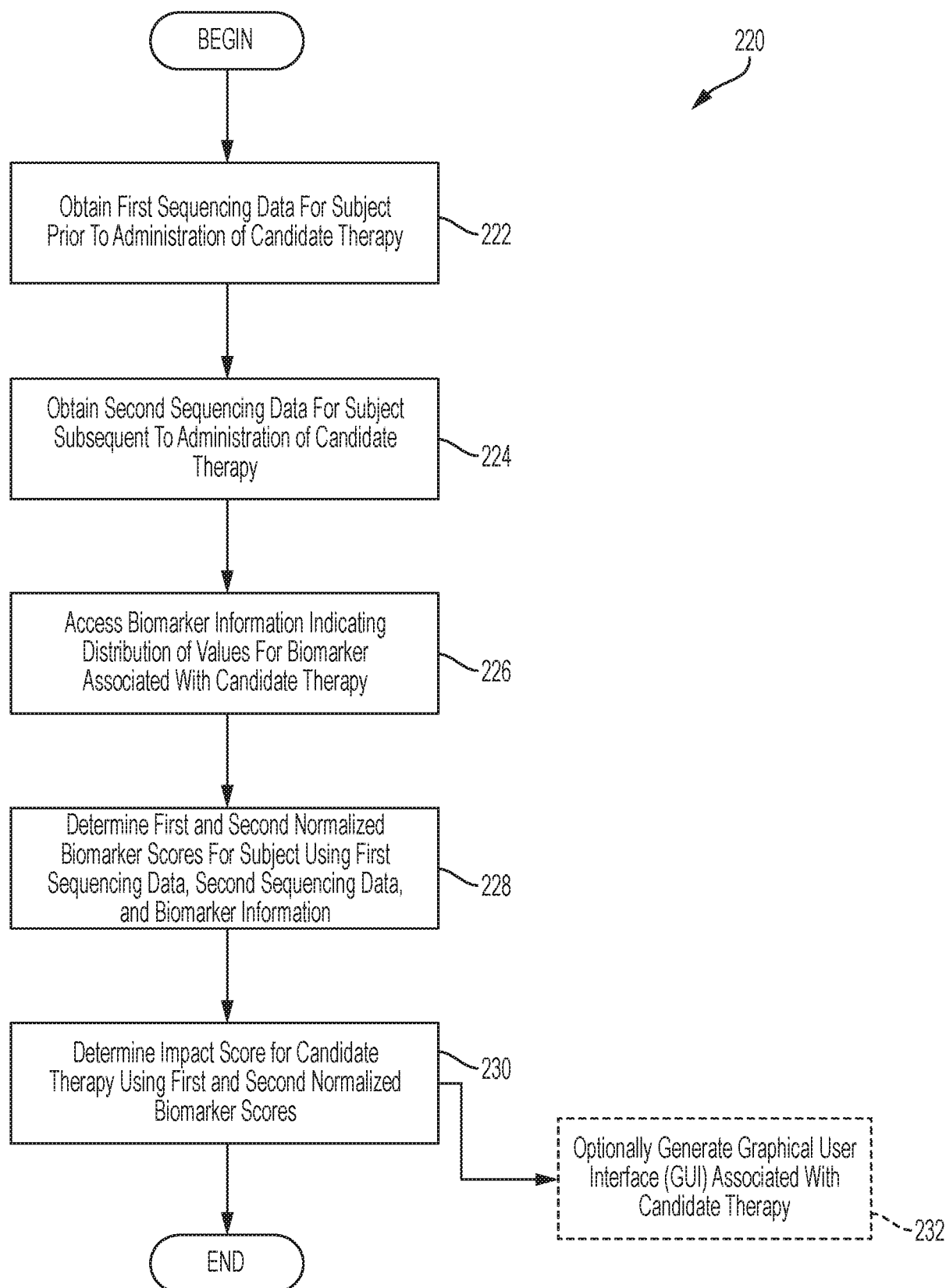


FIG. 2B

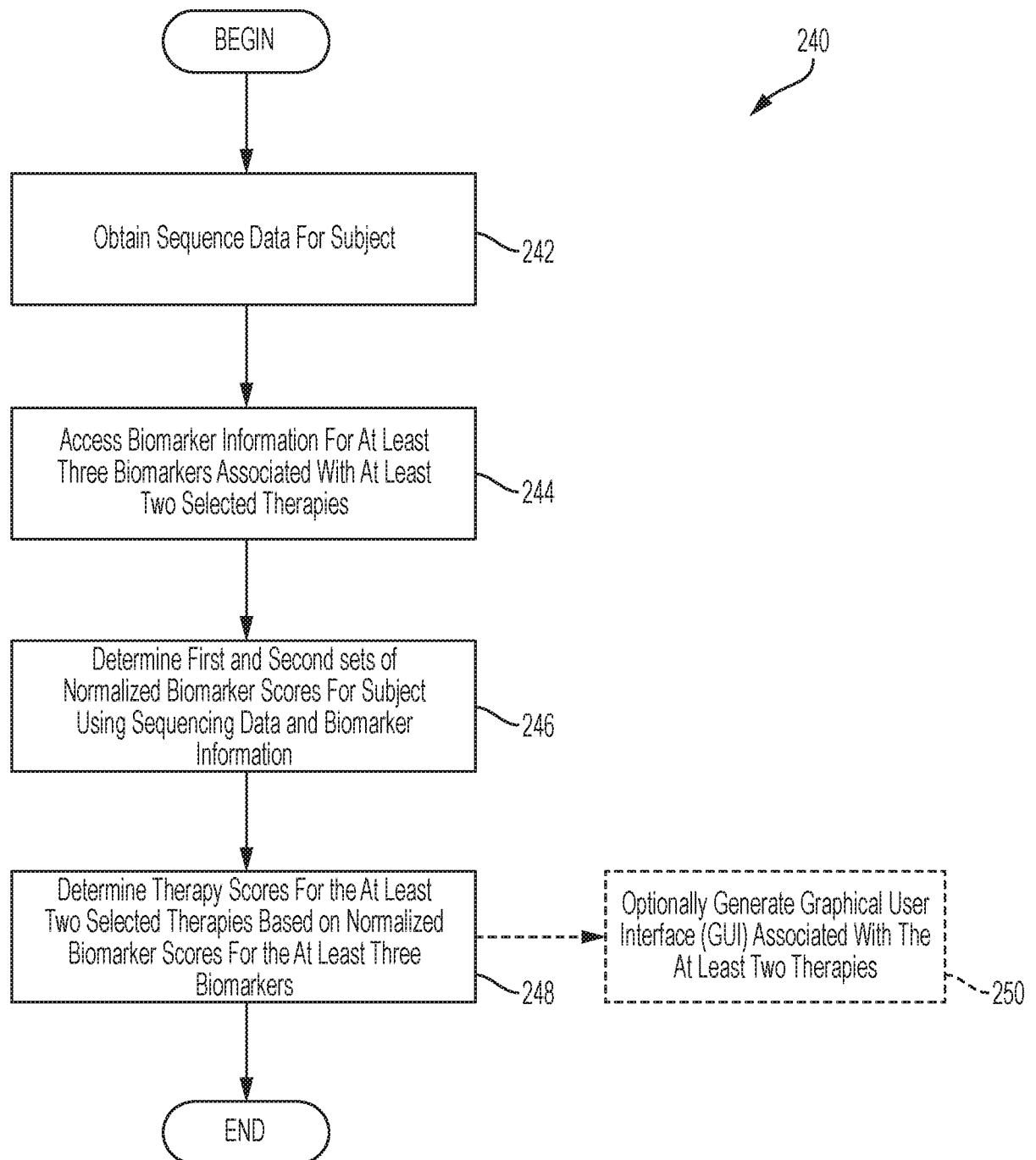


FIG. 2C

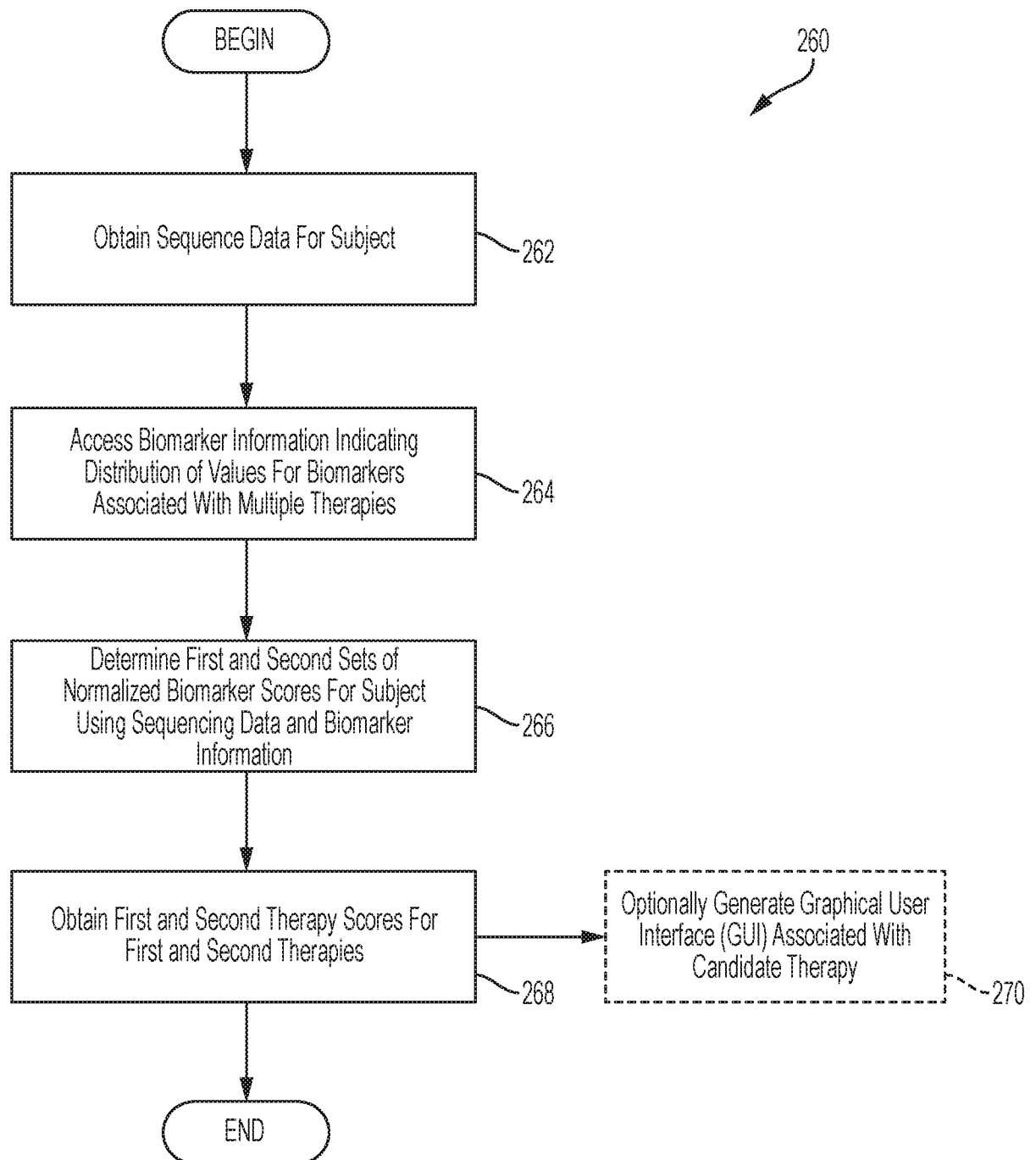


FIG. 2D

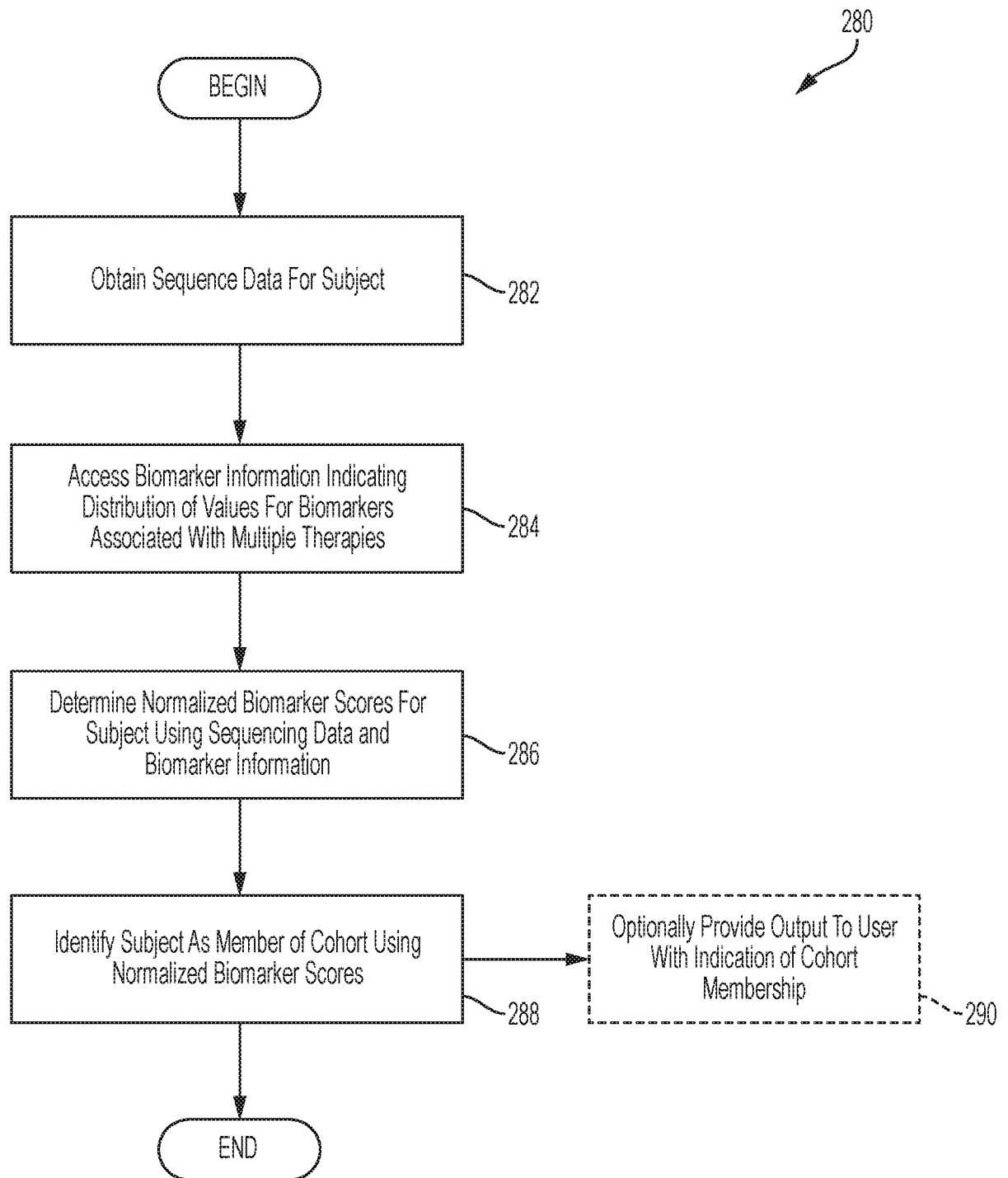


FIG. 2E

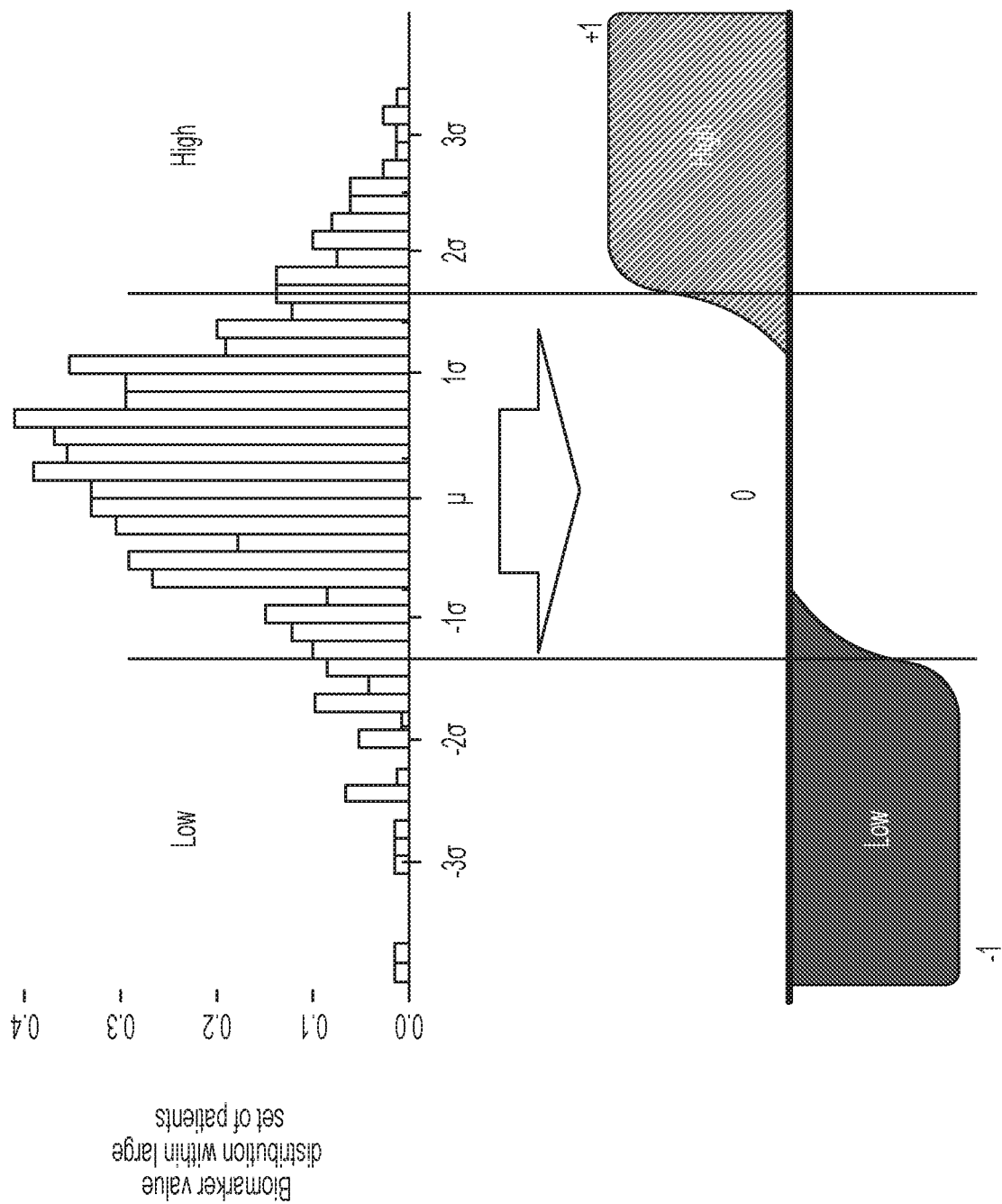


FIG. 3

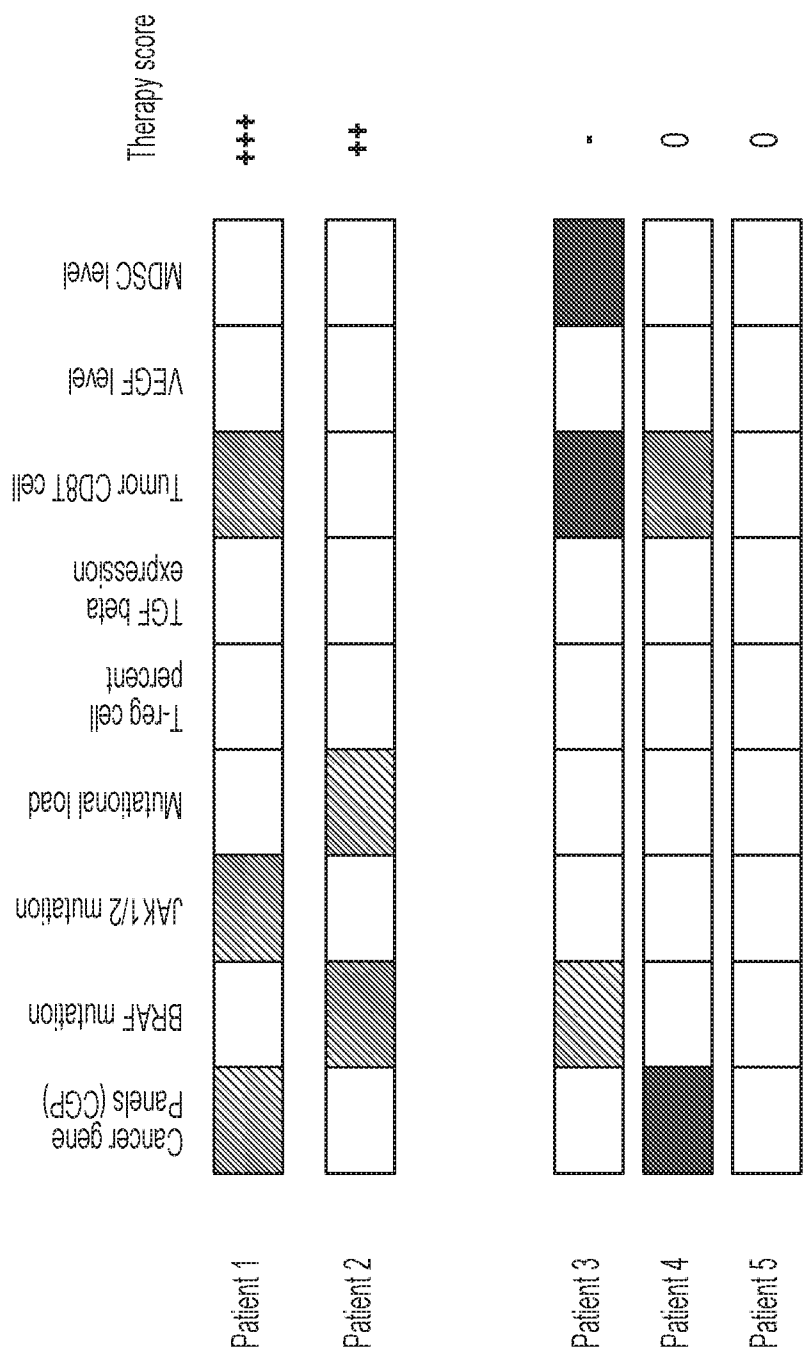


FIG. 4

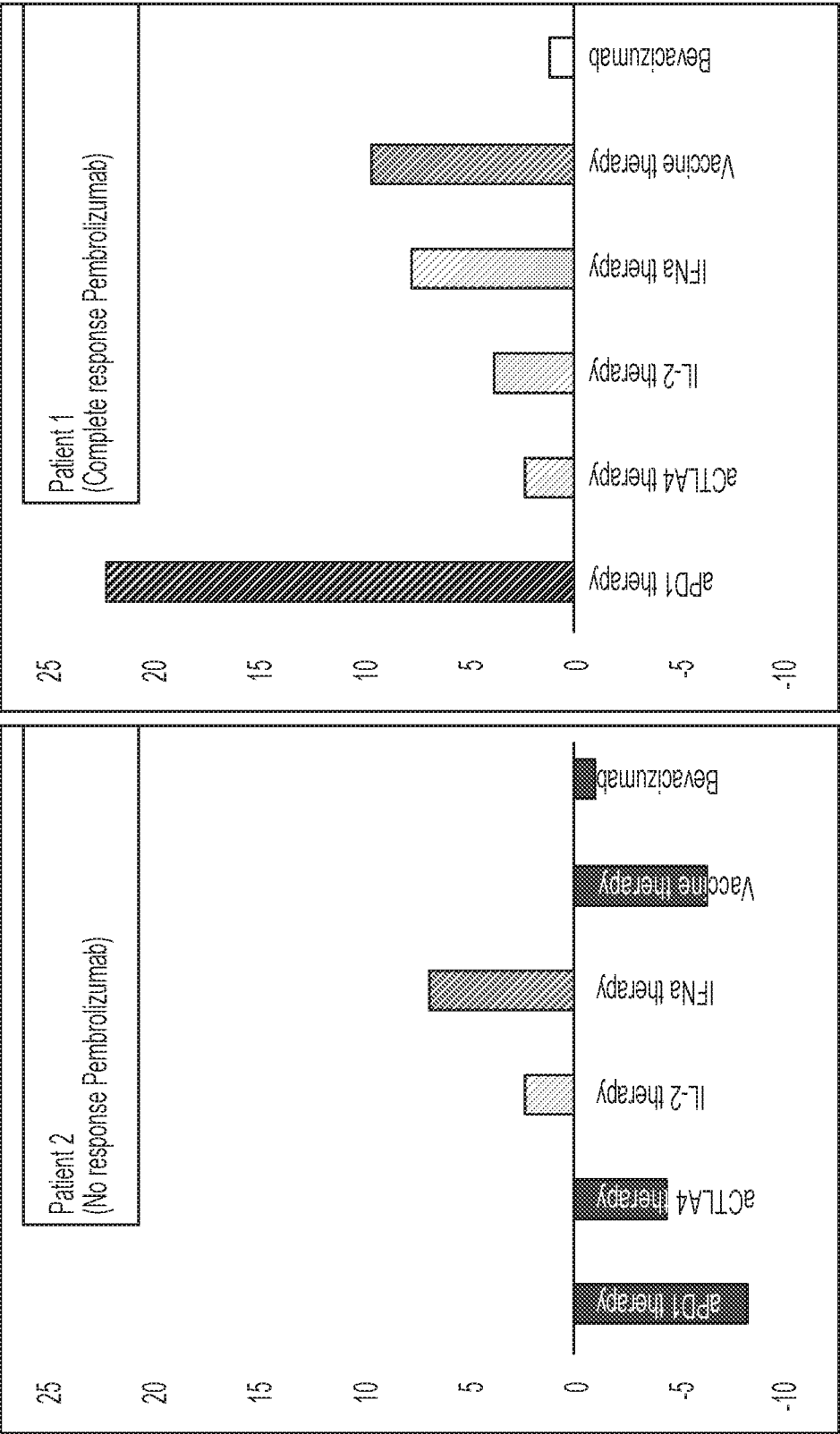


FIG. 5

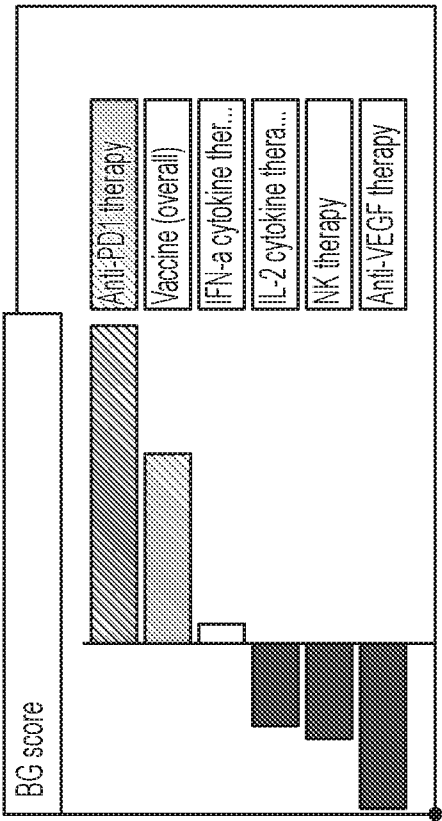


FIG. 6B

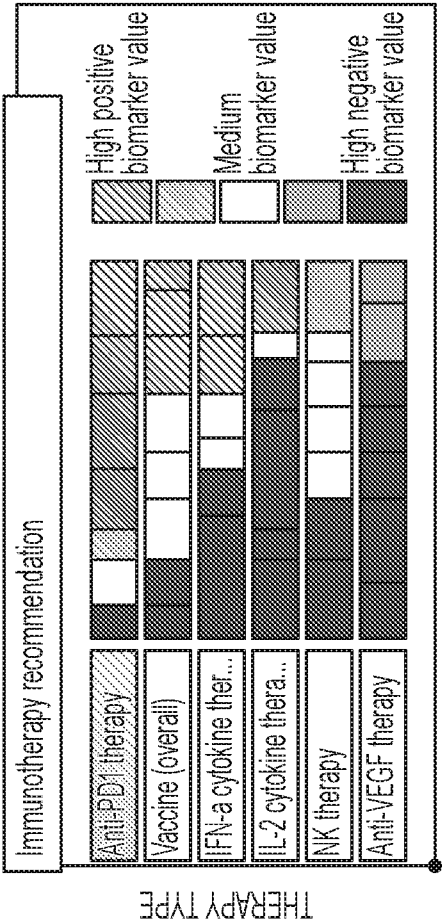


FIG. 6A

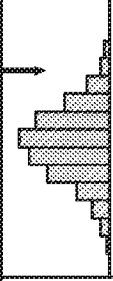
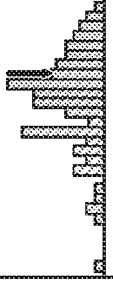
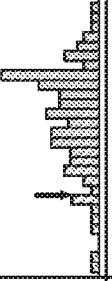
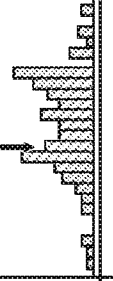
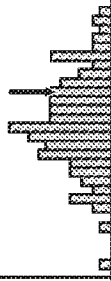
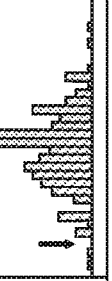
Biomarkers for Anti-PD1 therapy			
Nonsynonymous mutational burden Neointens load	Mutations, genetic rearrangements, insertions and deletions sometimes lead neointens - antigens which uniquely expressed in cancer cell but not in normal cells. Cytotoxic T-cells ...		HIGH
CD8 T cell number T cell density in tumor	One mechanism by which cancer tissues limit the host immune response is via upregulation of PD-1 ligand (PD-L1) and its ligation to PD-1 on antigen-specific CD8 T cells ...		HIGH
Mismatch-repair deficiency status	Mismatch repair-deficient tumors are more responsive to PD-1 blockade than are mismatch repair-proficient tumors ...	MLH1, MSH2 Mutations	YES
MDSC number	Myeloid-derived suppressor cells (MDSCs) are a heterogeneous population of cells that expand during cancer, inflammation and infection, and that have a remarkable ability to		LOW
T regulatory number	regulatory T (Treg) cells, crucial for the maintenance of immunological self-tolerance, are abundant in tumors. Most of them are chemo-attracted to tumor tissues, expanding locally and differentiating ...		LOW
VEGF expression VEGF concentration	VEGF-A a proangiogenic molecule produced by the tumors, plays a key role in the development of an immunosuppressive microenvironment. We report in the present work that VEGF-A produced in the tumor microenvironment enhances expression of PD-1 and other inhibitory checkpoints		Neutral
PD-L1 expression PD-L1 protein expression	The results of subgroup analyses suggest that the greatest benefit with the nivolumab alone may occur in the context of high PD-L1 tumor expression		LOW

FIG. 6C

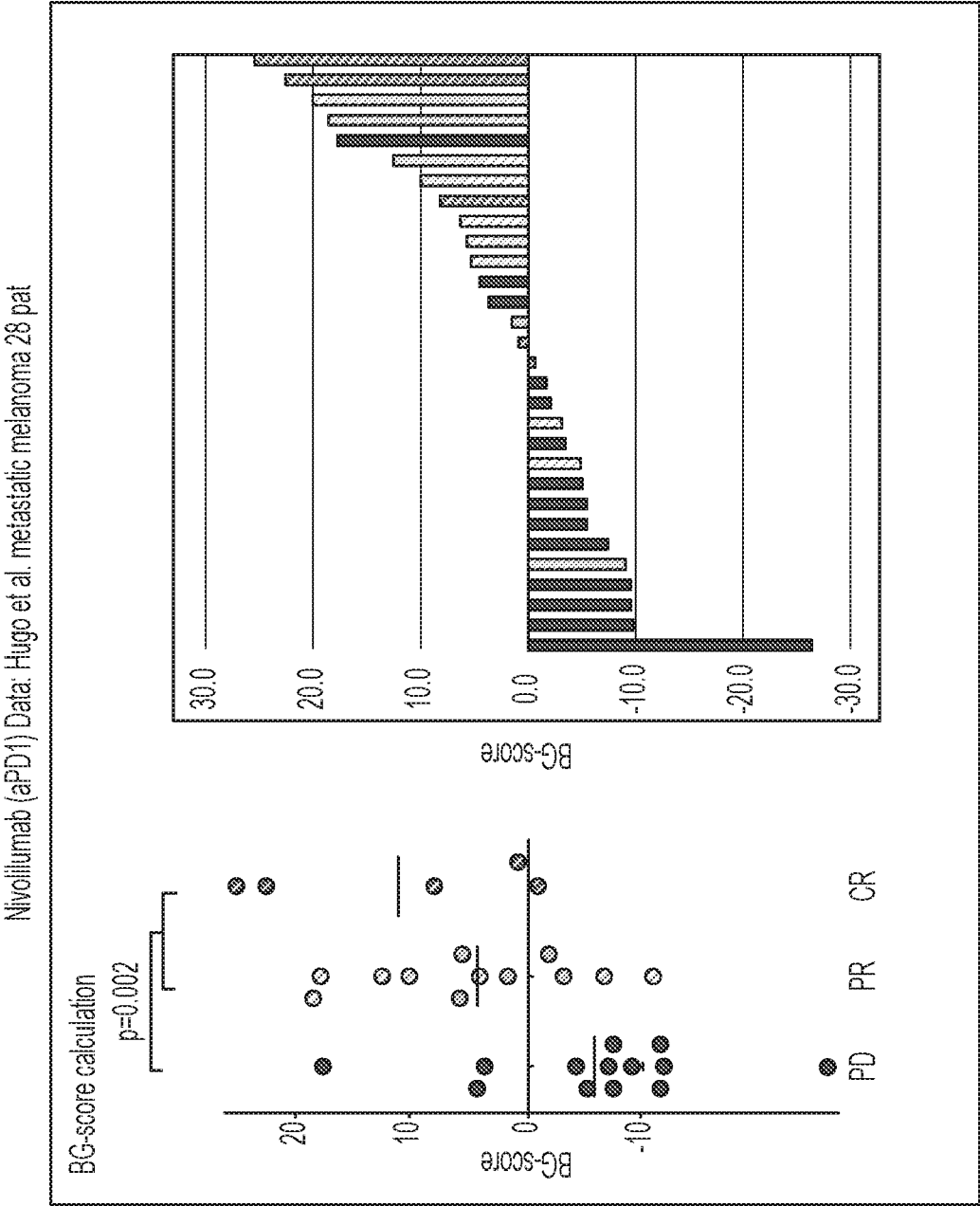
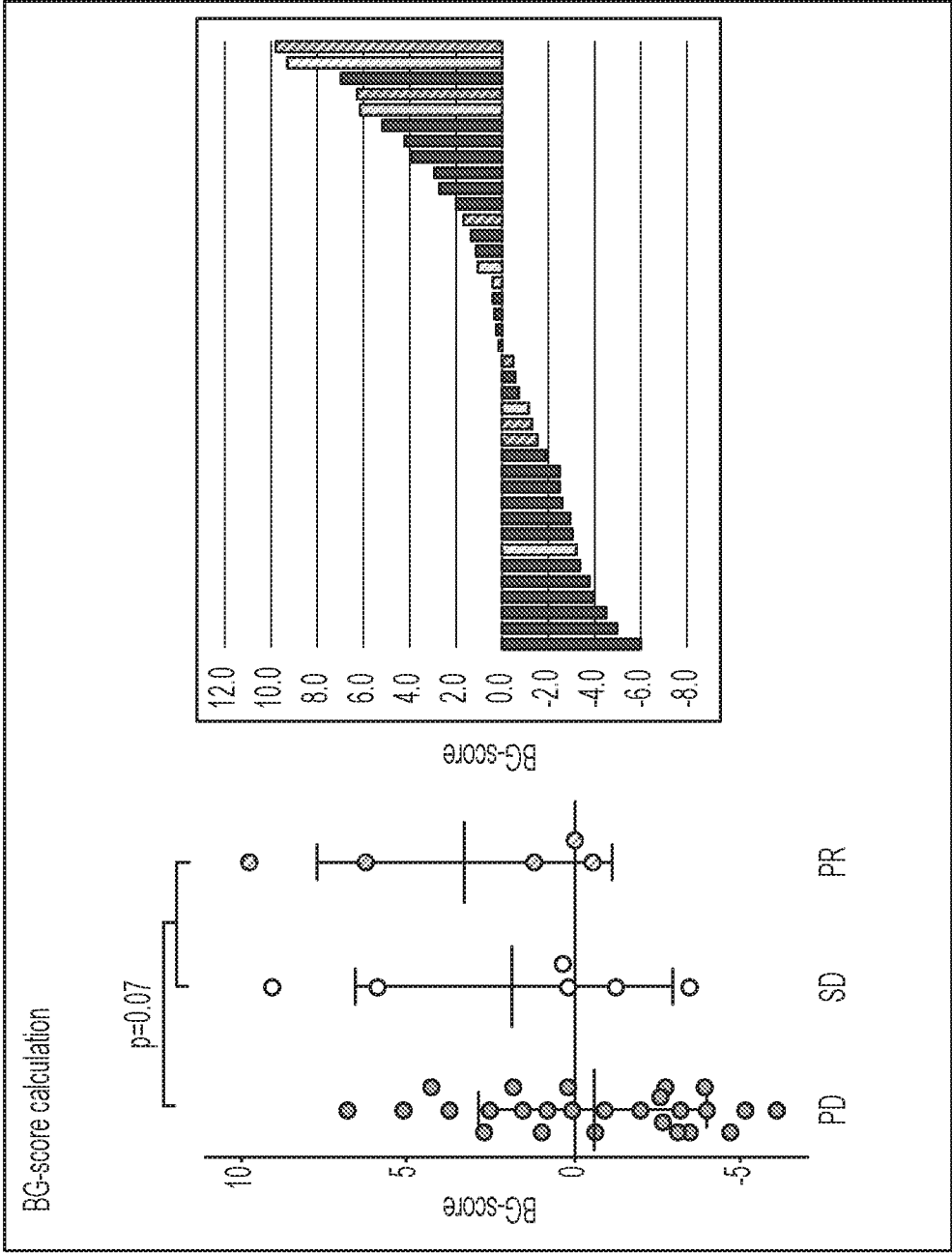


FIG. 7A



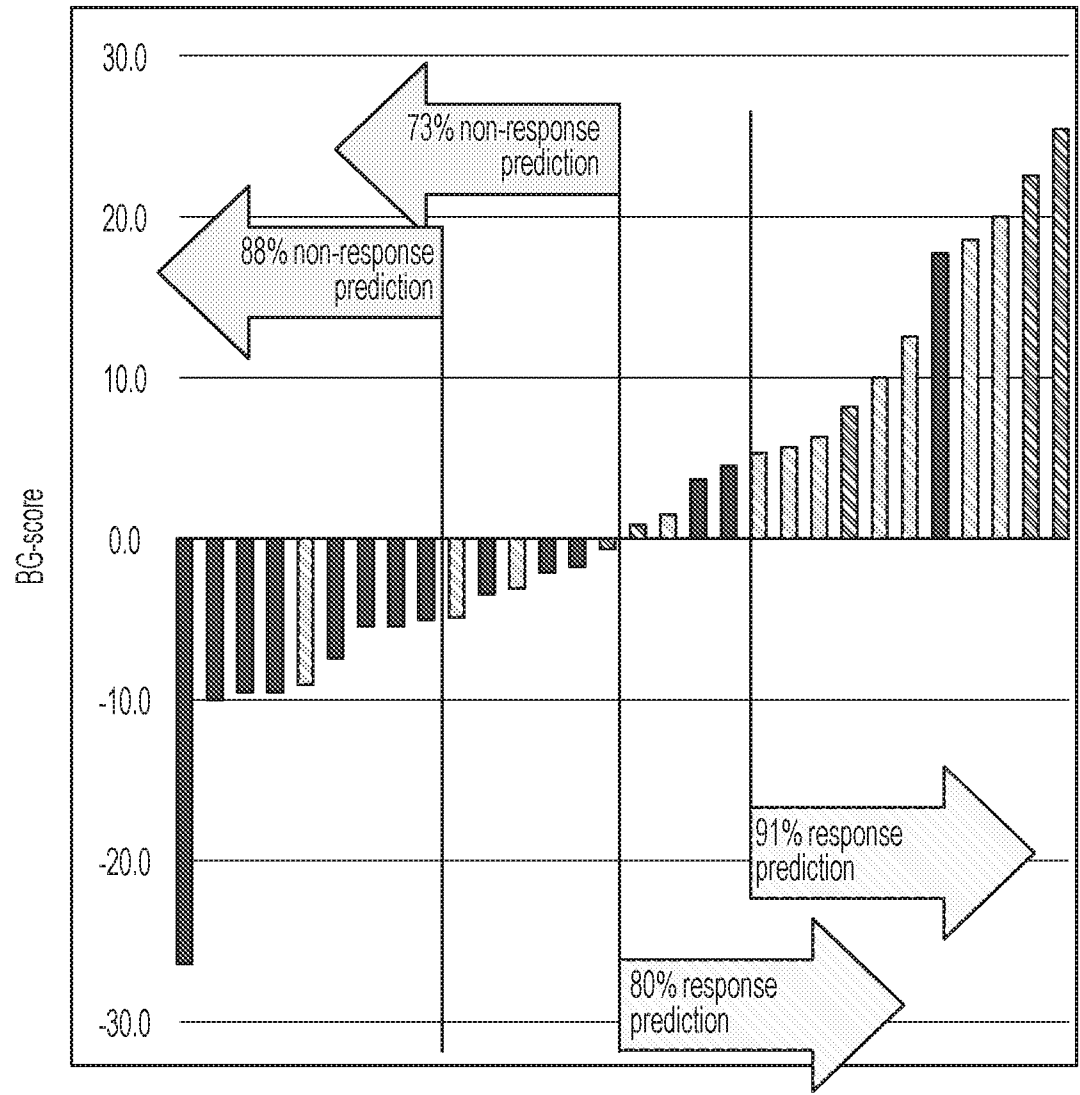


FIG. 7C

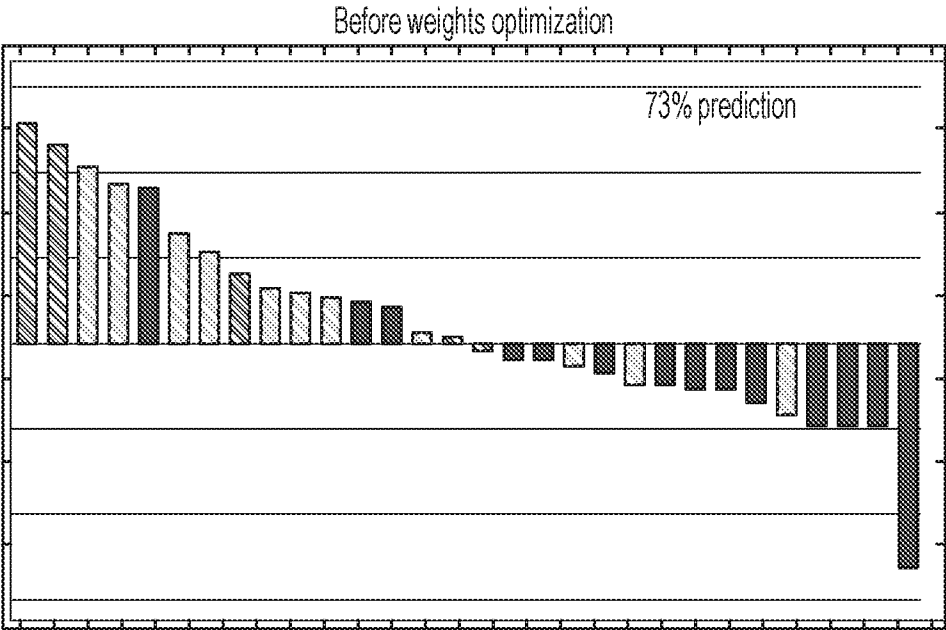


FIG. 8A

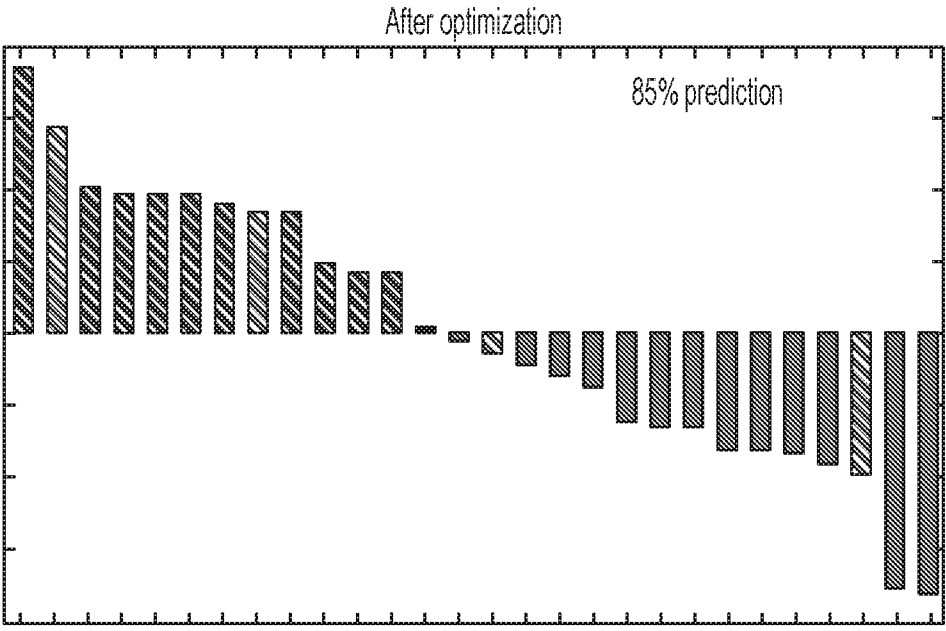


FIG. 8B

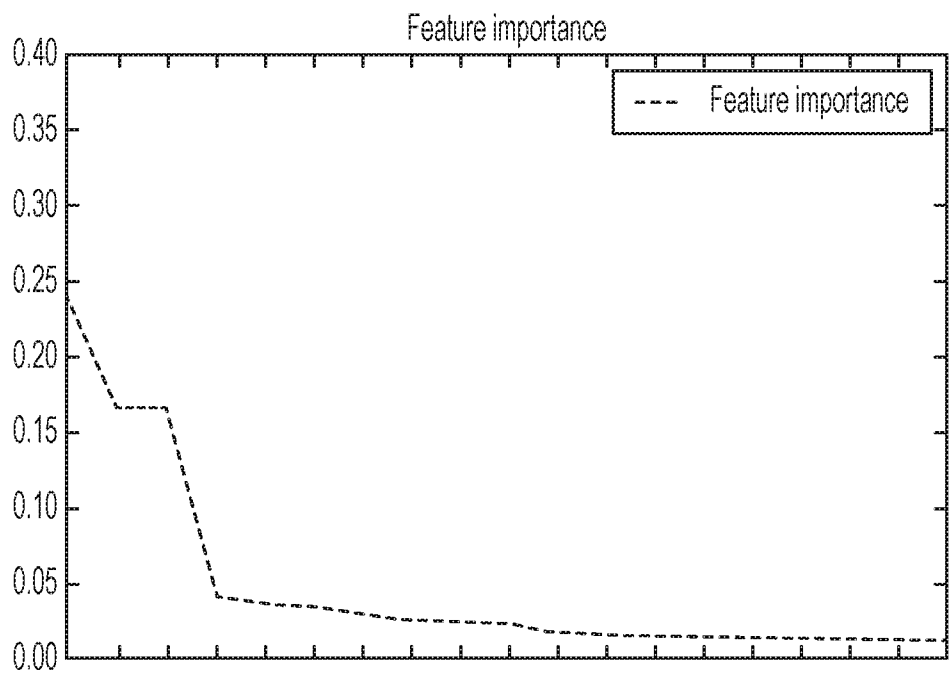


FIG. 8C

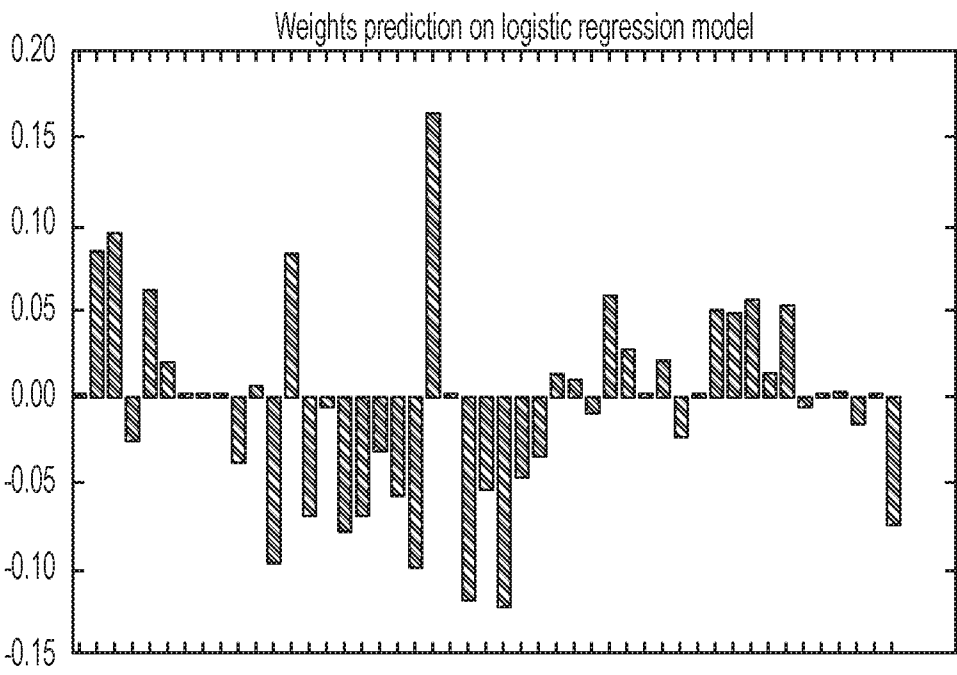


FIG. 8D

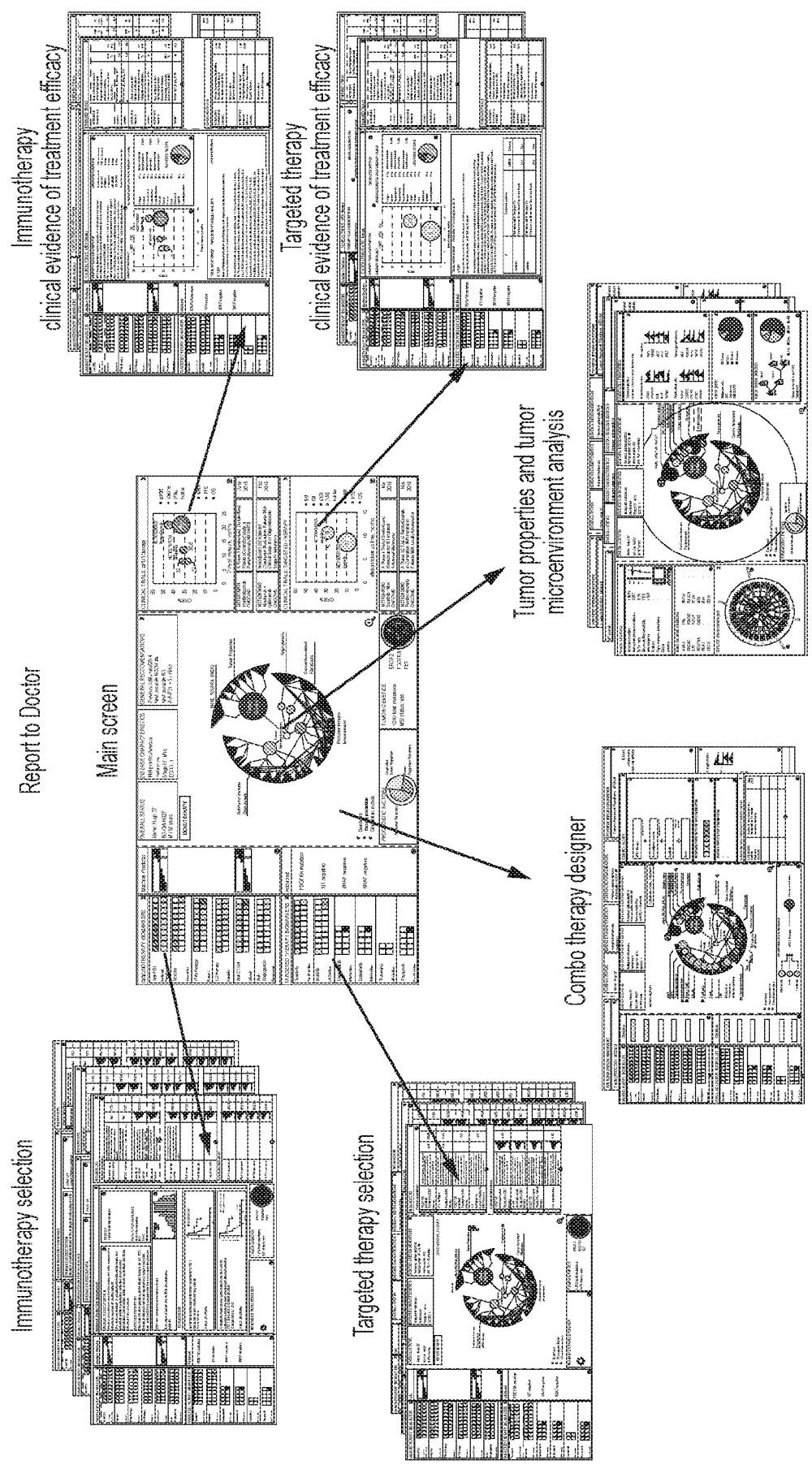
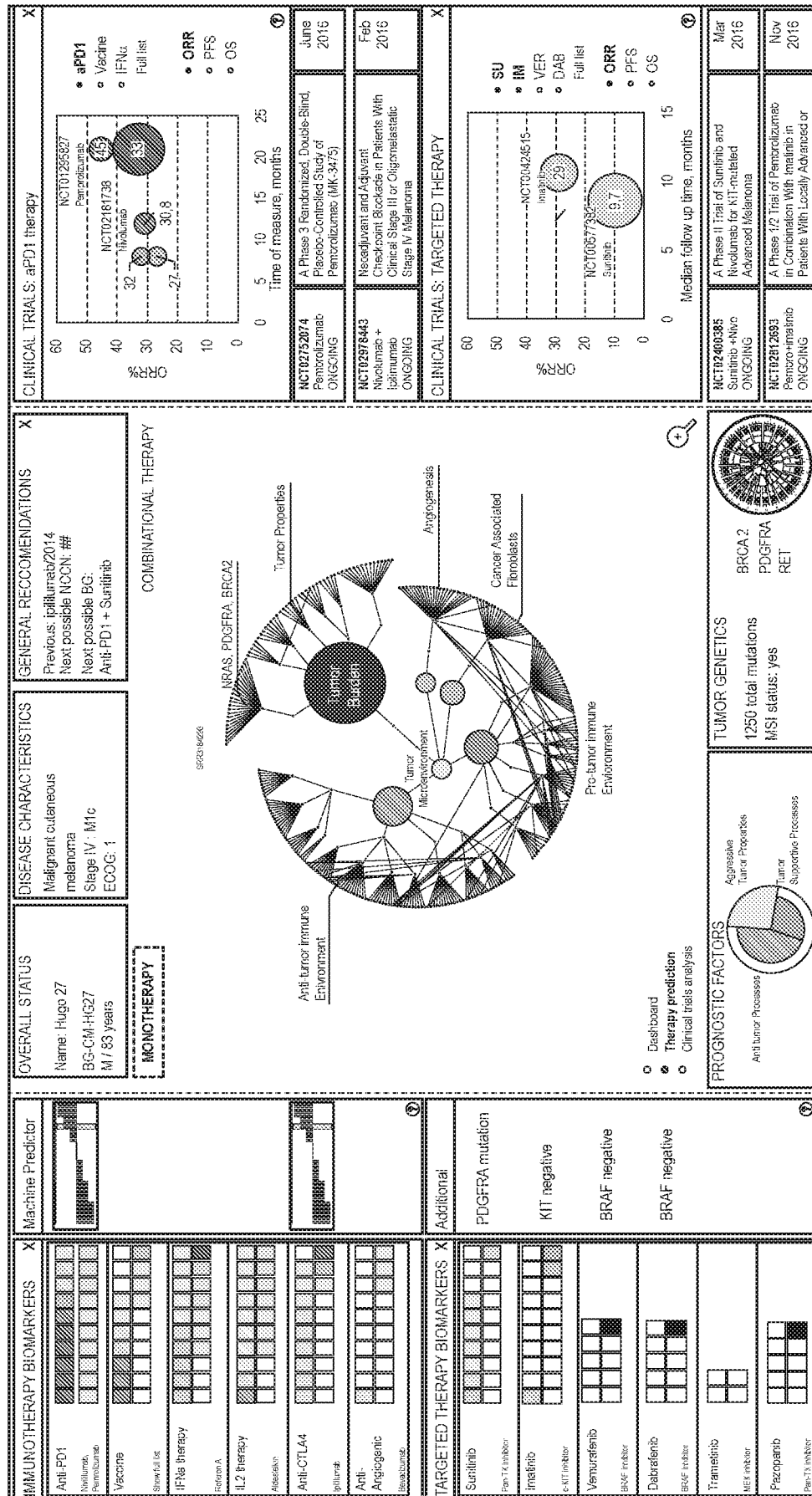
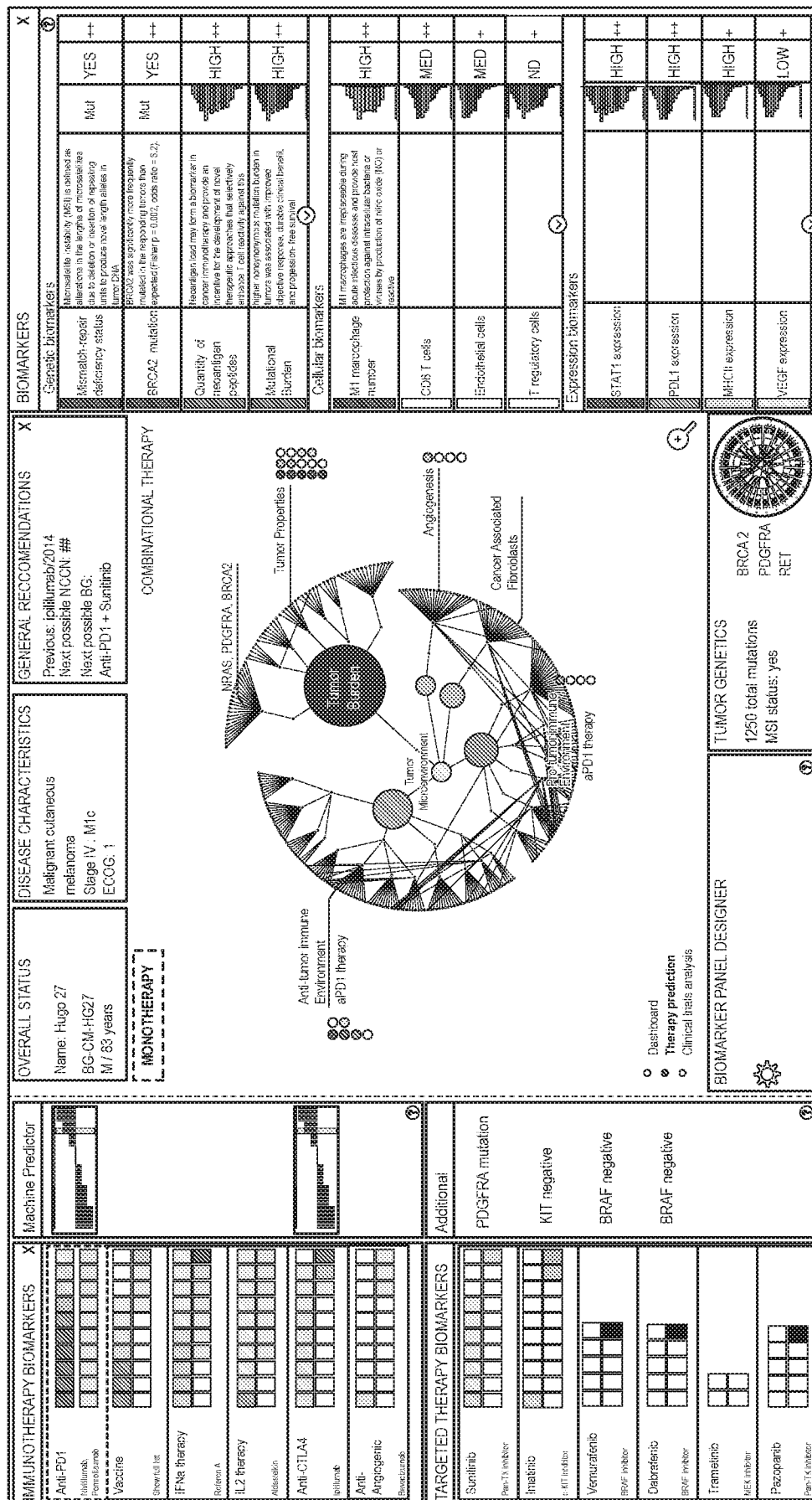


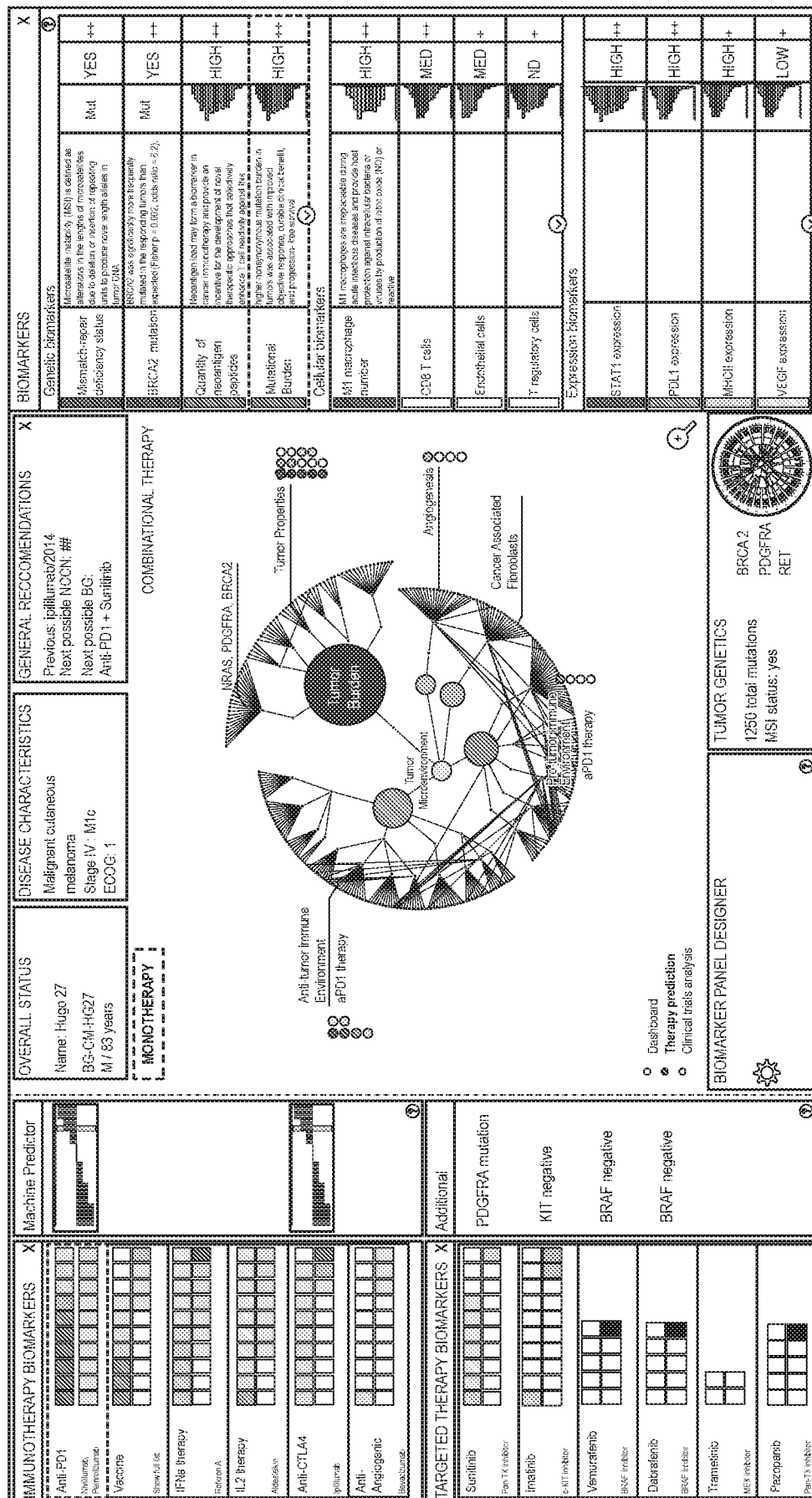
FIG. 9



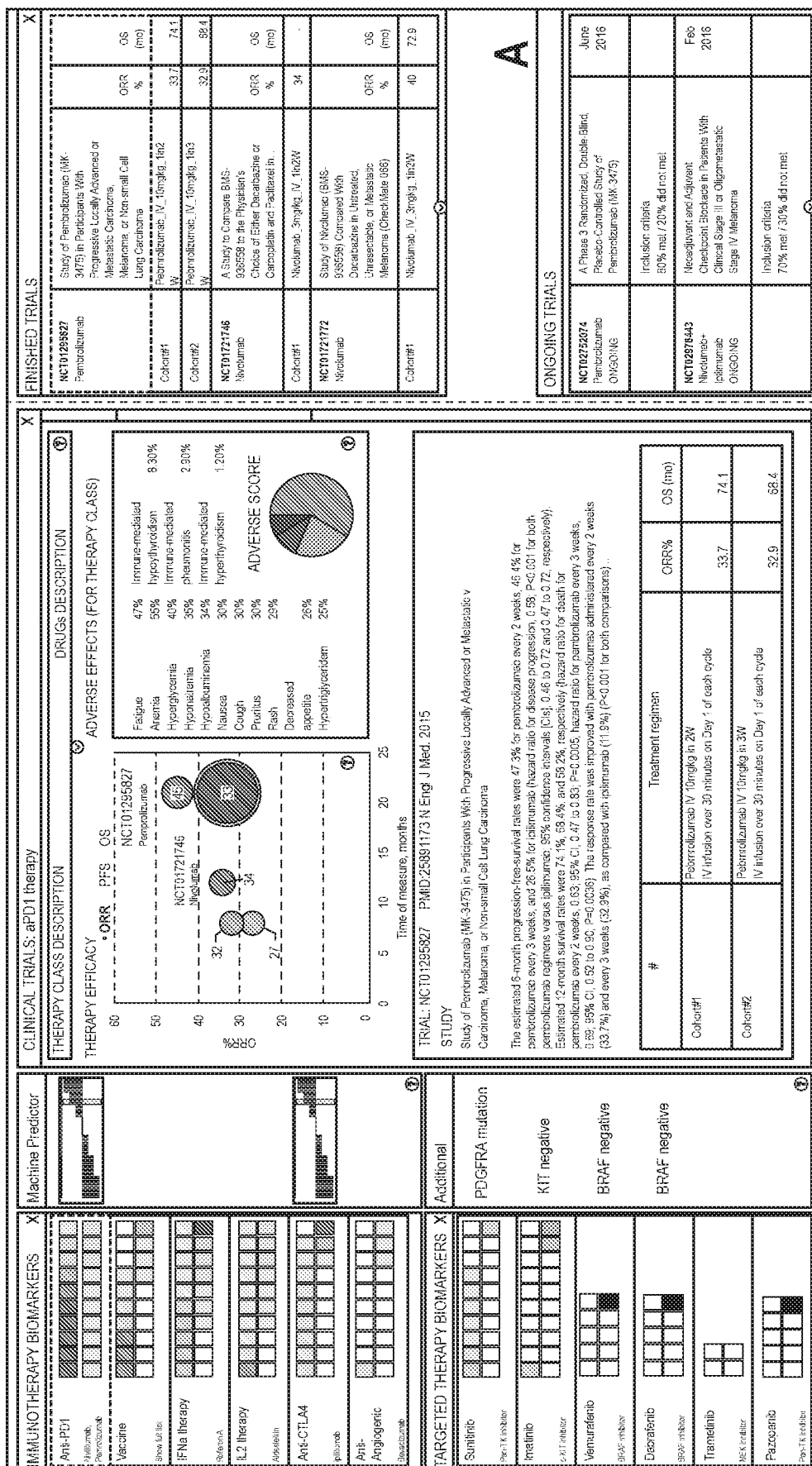


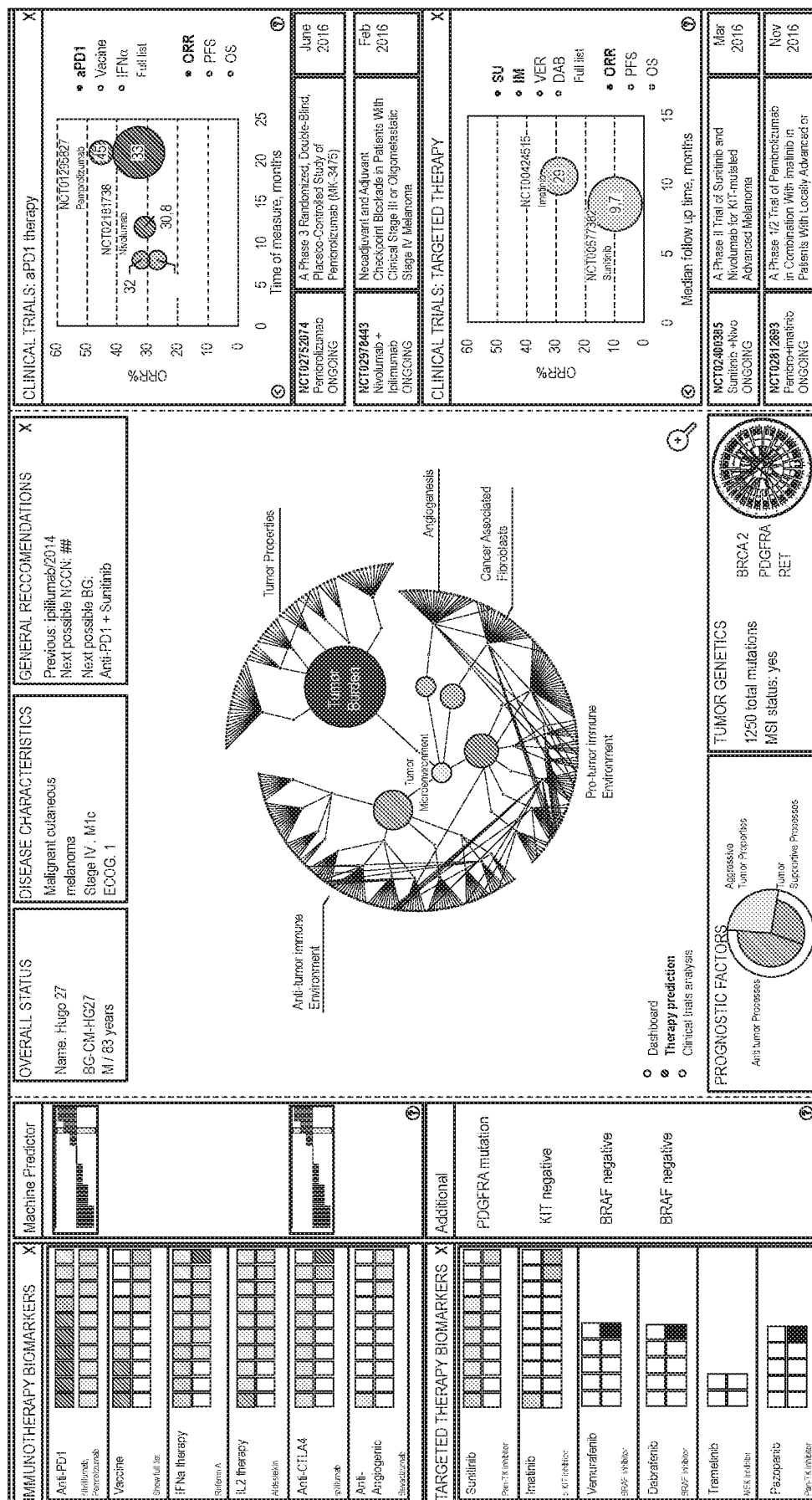


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216





415

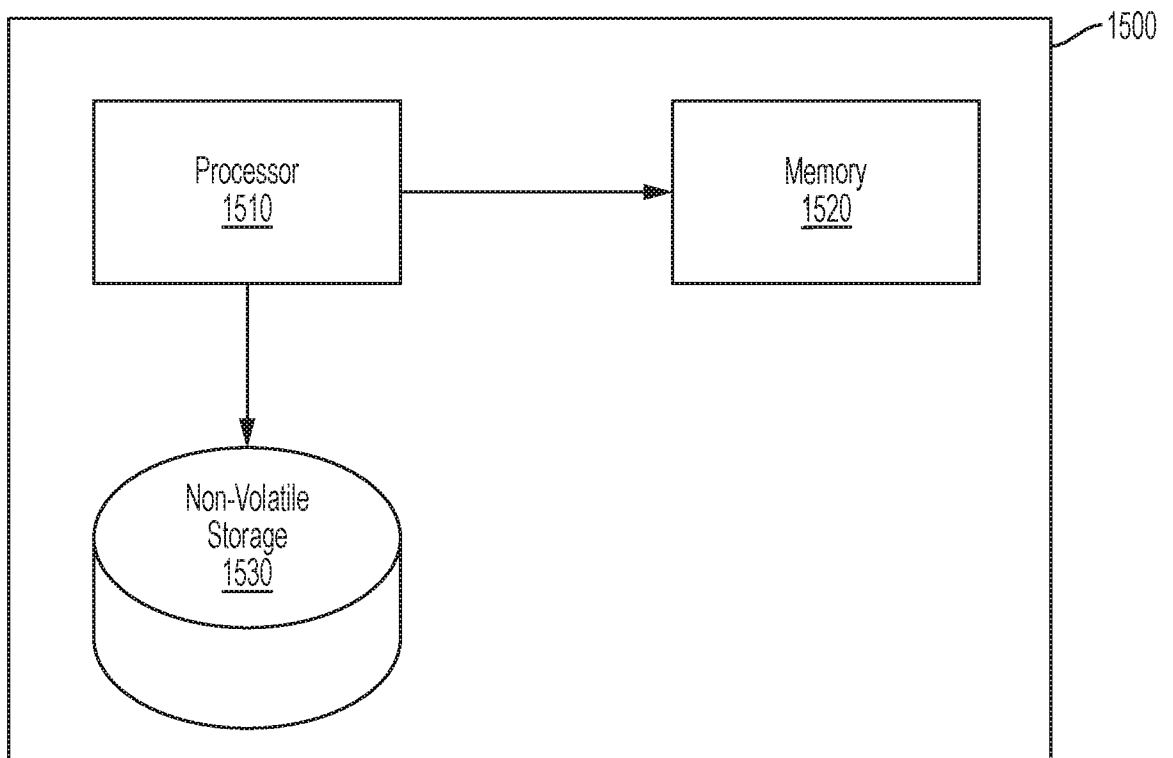


FIG. 15