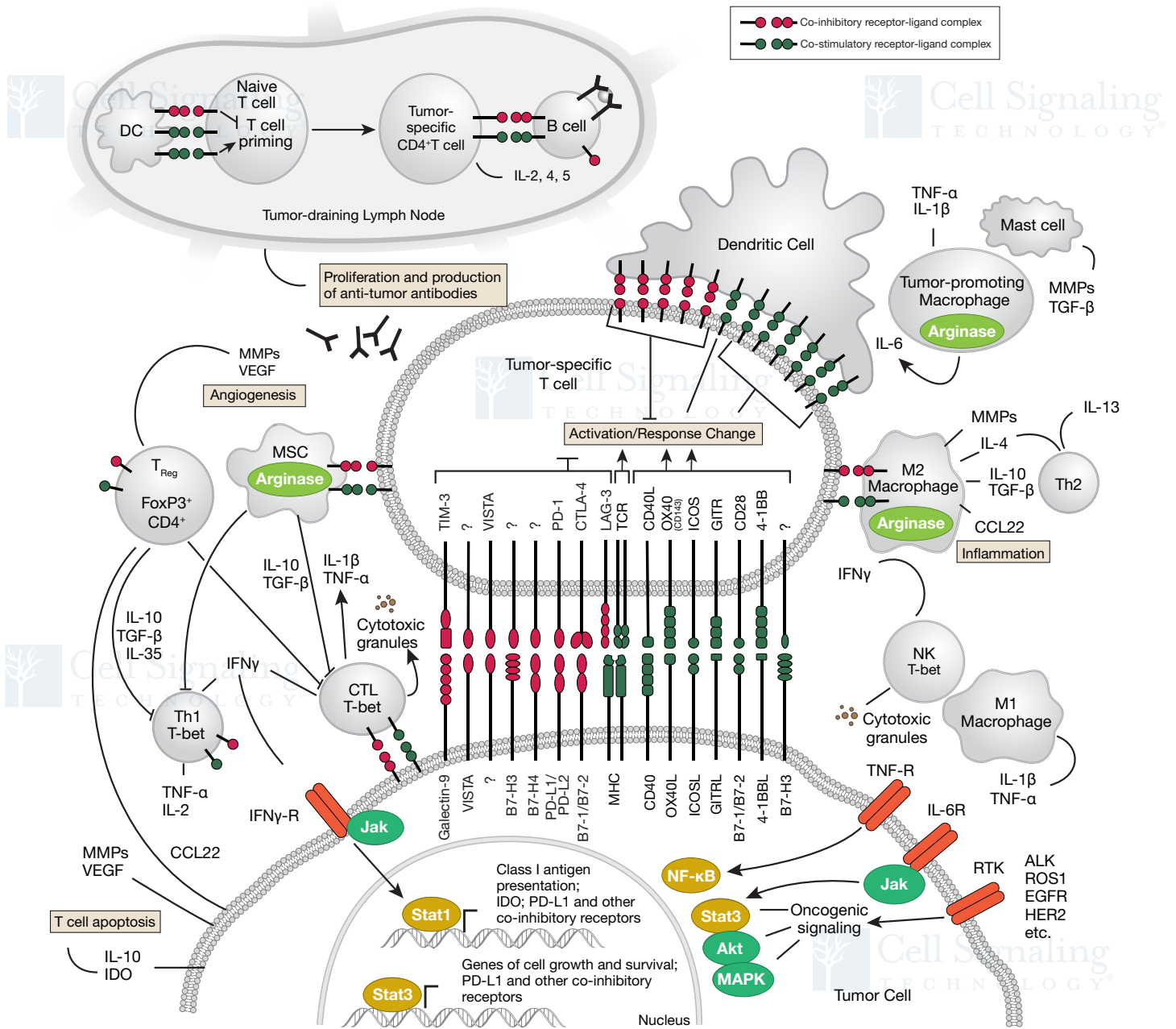


# Immune Checkpoint Signaling in the TME



## Pathway Diagram Key

- Direct Stimulatory Modification
  - Multistep Stimulatory Modification
  - Tentative Stimulatory Modification
  - Transcriptional Inhibitory Modification
  - Direct Inhibitory Modification
  - Multistep Inhibitory Modification
  - Tentative Inhibitory Modification
- |           |                                |   |                |                                |                 |  |   |
|-----------|--------------------------------|---|----------------|--------------------------------|-----------------|--|---|
| Acetylase | Apoptosis/ Autophagy Regulator | Deacetylase or Cytoskeletal Protein         | GTPase/GAP/GEF | Metabolic Enzyme               | Phosphatase     | Receptor                                   | Ubiquitin/SUMO Ligase or Deubiquitinase |
| Adaptor   | Cell Cycle Regulator           | Growth Factor/Cytokine/ Development Protein | Kinase         | Methyltransferase or G-protein | Protein Complex | Transcription Factor or Translation Factor | Other                                   |