



Cancer Immunotherapy: Chapter 11. Immune Escape: Immunosuppressive Networks

Sandra Demaria

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Tumors that become clinically apparent have invariably developed ways to escape immune control. The mechanisms of escape are multiple and diverse and do not necessarily require the loss by tumor cells of expression of antigens recognized by the immune system. More commonly, tumor-specific T cells are rendered dysfunctional by a number of soluble and cell surface-bound molecules that are produced or modified by the action of cancer cells or suppressive and regulatory immune cells recruited to the tumor. Overall, the obstacles that prevent tumor rejection are active at several levels, affecting T-cell priming, functional differentiation, recruitment to the tumor, survival and effectors' function inside the tumor. The multifaceted nature of these immunosuppressive networks represents a formidable obstacle to the success of immunotherapy. Nevertheless, the recent achievements of therapeutics targeting key checkpoint receptors highlight the great potential of strategies that are based on selective disruption of immunosuppressive networks.

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