



REVIEW

# Clinical Features and Management of Dermatologic Immune-Related Adverse Events for the Inpatient Dermatologist

Christopher J. Fay<sup>1</sup> · Christopher Iriarte<sup>1</sup> · Blair S. Allais<sup>1</sup> · Catherine E. Pisano<sup>1</sup> · Abdulhadi Jfri<sup>2,3,4</sup> · Cecilia A. Larocca<sup>1</sup> · Nicole R. LeBoeuf<sup>1</sup>

Accepted: 26 June 2023 / Published online: 17 July 2023

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## Abstract

**Purpose of Review** Immune checkpoint inhibitors (ICIs) have become a mainstay of oncologic treatment over the past decade. There are currently 11 ICIs approved by the U.S. Food and Drug Administration in the treatment of over a dozen malignancies. ICIs are associated with immune-related adverse events that can affect virtually any organ system. In some instances, these toxicities may lead to inpatient hospitalization and/or discontinuation of the ICI. This review highlights the clinical features and management of cutaneous toxicities most relevant to the inpatient dermatologist, including morbilliform eruptions, severe mucocutaneous reactions including lichenoid eruptions and Stevens-Johnson syndrome (SJS)/toxic epidermal necrolysis (TEN)-like reactions, drug reaction with eosinophilia and systemic symptoms, acute generalized exanthematous pustulosis, and autoimmune bullous disease. We also review other systemic toxicities that present with skin findings that are relevant to the inpatient dermatologist.

**Recent Findings** A distinct ICI-induced mucocutaneous eruption mimicking SJS/TEN but with a delayed presentation and more favorable prognosis has been reported. Biologic therapy for ICI-induced bullous pemphigoid is effective and does not impair survival compared with other systemic regimens. Early exposure to and higher doses of systemic corticosteroids have been shown to result in reduced progression-free and overall survival among patients on immunotherapy.

**Summary** Given the increasing number of patients receiving ICIs alone and in combination with other anti-cancer regimens, as well as the complicated and varied clinical appearance of immune related adverse events, it is vital to promptly diagnose and treat these toxicities to provide optimal patient care while minimizing interruptions in oncologic treatment.

**Keywords** Immune checkpoint inhibitors · Immunotherapy · Immune-related adverse events (irAE) · Skin toxicity · Inpatient dermatology · Stevens-Johnson syndrome

## Introduction

Immune checkpoint inhibitors (ICIs) are derived from monoclonal antibodies and target inhibitory receptors and ligands expressed on T lymphocytes, antigen presenting cells, and

tumor cells [1]. By targeting inhibitory receptors such as programmed cell death protein-1 (PD-1) and cytotoxic T-lymphocyte associated protein-4 (CTLA-4) as well as cell ligands such as programmed cell death ligand-1 (PD-L1), ICIs elicit an immune response that overrides the tumor-mediated immune evasion [2, 3].

Currently, there are 11 United States Food and Drug Administration (FDA) approved ICIs including anti-PD-1 agents (nivolumab, pembrolizumab, cemiplimab, dostarlimab, retifanlimab), anti-CTLA-4 agents (ipilimumab, tremelimumab), anti-PD-L1 agents (atezolizumab, avlumab, and durvabumab), and a fixed dose combination anti-PD-1 and anti-LAG-3 agent (nivolumab/relatlimab-rmbw), used in treatment of over a dozen malignancies [4–7]. While ICIs have been groundbreaking for oncologic treatment, they have also been associated with immune-related adverse events (irAEs) that can affect virtually

✉ Nicole R. LeBoeuf  
nleboeuf@bwh.harvard.edu

<sup>1</sup> Department of Dermatology, Brigham and Women's Hospital and the Center for Cutaneous Oncology, Dana-Farber/Brigham and Women's Cancer Center, Harvard Medical School, 450 Brookline Avenue, Boston, MA 02115, USA

<sup>2</sup> College of Medicine, King Saud Bin Abdulaziz University for Health Sciences, Jeddah, Saudi Arabia

<sup>3</sup> King Abdullah International Medical Research Center, Jeddah, Saudi Arabia

<sup>4</sup> Division of Dermatology, Department of Medicine, Ministry of the National Guard-Health Affairs, Jeddah, Saudi Arabia

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