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Review

#### Where Do We Stand with Immunotherapy for Advanced Pancreatic Ductal Adenocarcinoma: A Synopsis of Clinical Outcomes

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Abstract: Pancreatic cancer is the seventh leading cause of cancer-related mortality in both sexes across the globe. It is associated with extremely poor prognosis and remains a critical burden worldwide due to its low survival rates. Histologically, pancreatic ductal adenocarcinoma (PDAC) accounts for 80% of all pancreatic cancers; the majority of which are diagnosed at advanced stages, which makes them ineligible for curative surgery. Conventional chemotherapy provides a five-year overall survival rate of less than 8% forcing scientists and clinicians to search for better treatment strategies. Recent discoveries in cancer immunology have resulted in the incorporation of immunotherapeutic strategies for cancer treatment. Particularly, immune-checkpoint inhibitors, adoptive cell therapies and cancer vaccines have already shifted guidelines for some malignancies, although their efficacy in PDAC has yet to be elucidated. In this review, we summarize the existing clinical data on immunotherapy clinical outcomes in patients with advanced or metastatic PDAC.

Keywords: pancreatic cancer, immunotherapy; immune-checkpoint inhibitors; CAR T-cell therapy; cancer vaccines

#### 1. Introduction

Pancreatic cancer is the seventh leading cause of cancer-related mortality in both sense across the globe [1]. It remains a critical burden worldwide due to its low survival rates and extremely aggressive nature [2]. A total of 95% of pancreatic malignancies arise from exocrine parts (ductal epithelium, acinar cells and connective tissue), and another 5% develop from endocrine parenchyma [3]. Histologically, pancreatic ductal adenocarcinoma (PDAC) accounts for 80% of all pancreatic cancers [4], and the majority of cases are diagnosed at advanced stages (Figure 1) [5]. Localized cases can be treated with surgery, however, the five-year overall survival (OS) rate does not exceed 25% [6]. Unfortunately, there are no curative strategies for advanced stages, and palliative chemoradiotherapy reaches a five-year OS rate of less than 8% [7]. Chemotherapeutic choices of treatment include either modified FOLFIRINOX (5-fluorouracil, leucovorin, irinotecan, oxaliplatin) [8], gemcitabine monotherapy or gemcitabine in combination with nab-paclitaxel [9].

Recent discoveries in cancer immunology resulted in the incorporation of immunotherapeutic strategies for the treatment of various solid and hematologic malignancies [10]. Multiple clinical trials showed the higher efficacy of immune-checkpoint inhibitors (ICIs) for melanoma, lung cancer, renal cell carcinoma, colorectal cancer and hepatocellular carcinoma [11–15]. Adoptive chimeric antigen receptor (CAR) T-cell therapy is another immunotherapeutic tool being established as effective for hematologic malignancies, parfortunately, early-phase clinical trials have shown limited responses to immunotherapy



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# **Synopsis Of Clinical Cancer**

**Stewart Marshall Brooks** 

# **Synopsis Of Clinical Cancer:**

Clinical Cancer: a Synopsis. Synopsis of Clinical Cancer. 2nd Ed ,1970 **Synopsis of Clinical Oncology** Michael W. O'Dell, MD, Michael D. Stubblefield, MD, 2010-04-09 Authored by some of the world's leading cancer experts from wide ranging disciplines including oncology radiation oncology neurosurgery orthopedic surgery and radiology this compact volume provides authoritative state of the art primer level overviews of the various cancer types their evaluation and treatment Every aspect of oncology is covered with clear up to date descriptions of the general principles and concepts fundamental to attaining a clinical appreciation of the foundations of cancer care Each chapter is concise but highlights the basic tenets health care professionals working with cancer patients need to know to understand the diseases and current treatment options To facilitate review Key Points are boxed in each chapter to summarize important clinical information and concepts at a glance This book represents a synthesis of information from some of the top physicians in oncology that is not as easily accessed anywhere else Features of Synopsis of Clinical Oncology include Multi specialty expert authorship Focus **Synopsis of Clinical Cancer** on therapeutics management of cancer Key Points boxed in each chapter for guick review Condict Moore, 1970 Clinical Cancer Condict Moore, 1968 Synopsis of clinical cancer Condict Moore, 1973

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