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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



Citatione Akholno, L.; Tigai, Z.; Shok, D. Where Do We Stand with Immunotherapy for Advanced Parcovatic Ductal Adenocarcinomus A Synogosis of Clintool Outcomes. Biomedicines 2022, 10, 319s. https://doi.org/10.3090/

Academic Editore: Stefania Croci, Giulio Innamotati and Amodeo Amodei

Received: 6 November 2022 Accepted: 7 December 2022 Published: 9 December 2022

Belowensed by Service 10% 2.5(19%)

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Synopsis Of Clinical Cancer

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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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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 on the rapeutics management of cancer Key Points boxed in each chapter for quick review Clinical Cancer Research Loren K. Mell, Phuoc T. Tran, James B. Yu, Qiang (Ed) Zhang, 2018-11-28 Principles of Clinical Cancer Research provides comprehensive coverage of the fundamentals of clinical cancer research including the full spectrum of methodologies used in the field For those involved in research or considering research careers this book offers a mix of practical advice and analytical tools for effective training in theoretical principles as well as specific usable teaching examples The clinical oncologist or trainee will find a high yield practical guide to the interpretation of the oncology literature and the application of data to real world settings Valuable for both researchers and clinicians who wish to sharpen their skills this book contains all of the cornerstones and explanations needed to produce and recognize quality clinical science in oncology Written from the physician scientist's perspective the book lays a strong foundation in preclinical sciences that is highly relevant to careers in translational oncology research along with coverage of population and outcomes research and clinical trials It brings together fundamental principles in oncology with the statistical concepts one needs to know to design and interpret studies successfully With each chapter including perspectives of both clinicians and scientists or biostatisticians Principles of Clinical Cancer Research provides balanced instructive and high quality topic overviews and applications that are accessible and thorough for anyone in the field KEY FEATURES Gives real world examples and rationales behind which research methods to use when and why Includes numerous tables featuring key statistical methods and programming commands used in everyday clinical research Contains illustrative practical examples and figures in each chapter to help the reader master concepts Provides tips and pointers for structuring a career avoiding pitfalls and achieving success in the field of clinical cancer research Access to fully downloadable eBook The Cancer Story Stewart Marshall Cancer Principles and Practice of Oncology: Handbook of Clinical Cancer Genetics Ellen Matloff, 2013-03-25 Brooks, 1973 This practical user friendly guidebook will allow the clinician to search under disease site for the hereditary cancer syndromes relevant for his her patient's cancer For example a gynecologist oncologist whose patient has ovarian cancer can turn to the Ovary chapter and quickly read a summary of all of the hereditary cancer syndromes that include ovarian cancer She can learn the questions she should be asking when expanding that patient s personal and family history which genes are most relevant whether to refer that patient on for genetic counseling and testing and how to manage that patient long term if the patient is mutation positive or negative The same holds true for the practicing oncologist surgeon urologist endocrinologist gynecologist primary care physician physician s assistant advanced practice nurse and any other clinician seeing a patient who has had cancer This guidebook also contains an overview article on genetic counseling and testing and several in depth articles on issues that are up and coming in the field of hereditary cancer Health Science Books,

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