# Cases from the Community Clinical Investigators Provide Perspectives on Actual Patients with Pancreatic Cancer (Audio Program)

# **CME** Information

## TARGET AUDIENCE

This activity is intended for medical oncologists, hematologyoncology fellows, surgeons and other healthcare providers involved in the treatment of pancreatic cancer.

## **OVERVIEW OF ACTIVITY**

Pancreatic cancer is the fourth most common cause of cancer-related death among US men and women. The overwhelming majority (approximately 90%) of pancreatic cancers are ductal adenocarcinomas, and many patients diagnosed with pancreatic adenocarcinoma (PAD) do not exhibit distinctive symptoms until the disease has reached an advanced stage. For all stages of PAD the combined 1-year survival rate for patients who do not receive surgery is approximately 29%, and the 5-year rate is an appalling 7%. Published results from ongoing trials have led to the emergence of new therapeutic targets and regimens, and the poor clinical course for so many patients with progressive PAD mandates the consideration of these approaches. In order to offer optimal patient care — including the option of clinical trial participation — the practicing medical oncologist must be well informed of these advances.

This CME program was developed from the proceedings of a satellite symposium held during the 2019 Gastrointestinal Cancers Symposium. It explores the most significant therapeutic advances in the field of pancreatic cancer by using the perspectives of leading experts on challenging cases and questions submitted by community oncologists to frame a discussion of how this information has aided in the refinement of current clinical practice and ongoing research. This activity will help medical oncologists and other allied healthcare professionals find answers to the individualized questions and concerns they frequently encounter, and in turn it will help them to provide high-quality cancer care.

#### LEARNING OBJECTIVES

• Develop an evidence-based strategy for the treatment of resectable or borderline-resectable PAD, exploring the role of neoadjuvant and adjuvant chemotherapy and/or radiation therapy.

- Consider patient and disease characteristics and available clinical trial data in the selection and sequencing of systemic therapy for locally advanced or metastatic PAD.
- Design and implement a plan of care to recognize and manage side effects and toxicities associated with the use of approved systemic regimens for the management of locally advanced or metastatic PAD and thus to support quality of life and continuation of therapy.
- Recall available and emerging data with investigational agents currently in clinical testing for PAD, and where applicable refer eligible patients for trial participation.

# ACCREDITATION STATEMENT

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#### AMERICAN BOARD OF INTERNAL MEDICINE (ABIM) — MAINTENANCE OF CERTIFICATION (MOC)

Successful completion of this CME activity, which includes participation in the evaluation component, enables the participant to earn up to 1.25 Medical Knowledge MOC points in the American Board of Internal Medicine's (ABIM) Maintenance of Certification (MOC) program. Participants will earn MOC points equivalent to the amount of CME credits claimed for the activity. It is the CME activity provider's responsibility to submit participant completion information to ACCME for the purpose of granting ABIM MOC credit.

Please note, this program has been specifically designed for the following ABIM specialty: **medical oncology**.

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## HOW TO USE THIS CME ACTIVITY

This CME activity consists of a audio component. To receive credit, the participant should review the CME information, listen to the MP3s, complete the Post-test with a score of 80% or better and fill out the Educational Assessment and Credit Form located at **ResearchToPractice.com/GICancers19/ Pancreatic/Audio/CME**. The corresponding video program is available as an alternative at **ResearchToPractice.com/ GICancers19/Pancreatic**.

## CONTENT VALIDATION AND DISCLOSURES

Research To Practice (RTP) is committed to providing its participants with high-quality, unbiased and state-of-theart education. We assess conflicts of interest with faculty, planners and managers of CME activities. Conflicts of interest are identified and resolved through a conflict of interest resolution process. In addition, all activity content is reviewed by both a member of the RTP scientific staff and an external, independent physician reviewer for fair balance, scientific objectivity of studies referenced and patient care recommendations.

**FACULTY** — The following faculty (and their spouses/partners) reported relevant conflicts of interest, which have been resolved through a conflict of interest resolution process:

#### Andrew E Hendifar, MD

Medical Director, Pancreatic Cancer Gastrointestinal and Neuroendocrine Malignancies Cedars-Sinai Medical Center Los Angeles, California

Advisory Committee: Ipsen Biopharmaceuticals Inc; Consulting Agreements: Ipsen Biopharmaceuticals Inc, Novartis; Contracted Research: Halozyme Inc, Ipsen Biopharmaceuticals Inc.

#### Eileen M O'Reilly, MD

Winthrop Rockefeller Chair in Medical Oncology Section Head Hepatopancreaticobiliary/Neuroendocrine Cancers Gastrointestinal Oncology Service Associate Director David M Rubenstein Center for Pancreatic Cancer Attending Physician, Member Memorial Sloan Kettering Cancer Center Professor of Medicine Weill Cornell Medical College New York, New York

**Consulting Agreements:** 3DMedcare, Agios Pharmaceuticals Inc, Alignmed, Amgen Inc, Antengene, Aptus Clinical, Arbutus Biopharma, ASLAN Pharmaceuticals, Astellas Pharma Global Development Inc, AstraZeneca Pharmaceuticals LP, Bayer HealthCare Pharmaceuticals, BeiGene, BioLineRx, Boston Scientific Corporation, BridgeBio, Bristol-Myers Squibb Company, CARsgen Therapeutics, CASI Pharmaceuticals, Celgene Corporation, Cipla Limited, CytomX Therapeutics, Daiichi Sankyo Inc, Debiopharm Group, Delcath Systems Inc, Eisai Inc, Exelixis Inc, Genoscience Pharma, Gilead Sciences Inc, Halozyme Inc, Hengrui Therapeutics Inc, Incyte Corporation, Inovio Pharmaceuticals Inc, Ipsen Biopharmaceuticals Inc, Janssen Biotech Inc, Jazz Pharmaceuticals Inc, Kyowa Hakko Kirin Co Ltd, LAM, Lilly, Loxo Oncology, Merck, MINAPHARM Pharmaceuticals, NewLink Genetics Corporation, Novella Clinical, Onxeo, PCI Biotech, Pfizer Inc. Pharmacyclics LLC, an AbbVie Company, PharmaCyte Biotech, Pieris Pharmaceuticals, QED Therapeutics, RedHill, Sanofi Genzyme, Servier, Silenseed Ltd, SillaJen, Sobi, Targovax, twoXAR, Vicus Therapeutics, Yakult Pharmaceutical Industry CO LTD, Yiviva; Contracted Research: Acta Biologica, Agios Pharmaceuticals Inc, Array BioPharma Inc, AstraZeneca Pharmaceuticals LP, Bayer HealthCare Pharmaceuticals, BeiGene, Bristol-Myers Squibb Company, CASI Pharmaceuticals, Celgene Corporation, Exelixis Inc, Genentech, Halozyme Inc, Incyte Corporation, Lilly, MabVax, Novartis, OncoQuest Inc, Polaris Group, Puma Biotechnology Inc, QED Therapeutics, Roche Laboratories Inc.

#### Philip A Philip, MD, PhD

Kathryn Cramer Endowed Chair in Cancer Research Professor of Oncology and Pharmacology Leader, GI and Neuroendocrine Oncology Vice President of Medical Affairs Karmanos Cancer Institute Wayne State University Detroit, Michigan

Advisory Committee: AbbVie Inc., Biolinx, Caris Life Sciences, Celgene Corporation, Eisai Inc, Forty Seven Inc, Halozyme Inc, Ipsen Biopharmaceuticals Inc, Lexicon Pharmaceuticals Inc, Lilly, Merck, Novartis, Rafael Pharmaceuticals Inc, Taiho Oncology Inc; Consulting Agreements: AbbVie Inc, Celgene Corporation, Lilly, Merck, Rafael Pharmaceuticals Inc; Contracted Research: AAA Pharmaceutical, Astellas Pharma Global Development Inc, Bayer HealthCare Pharmaceuticals, Biolinx, Boston Biomedical Pharma Inc, Bristol-Myers Squibb Company, Celgene Corporation, Daiichi Sankyo Inc, Eisai Inc, Forty Seven Inc, Gilead Sciences Inc, Halozyme Inc, Incyte Corporation, Lilly, Merck, Novocure, QED Therapeutics, Taiho Oncology Inc, Tyme Technologies Inc; Data and Safety Monitoring Board/Committee: ASLAN Pharmaceuticals, Blueprint Medicines, Erytech Pharma, Lexicon Pharmaceuticals Inc; Speakers Bureau: Bristol-Myers Squibb Company, Celgene Corporation, Ipsen Biopharmaceuticals Inc, Merck.

#### Margaret A Tempero, MD

Director, UCSF Pancreas Center The Rombauer Family Distinguished Professorship in Pancreas Cancer Clinical and Translational Science Leader, Pancreas Cancer Program Professor of Medicine, Division of Hematology and Oncology University of California, San Francisco San Francisco, California

Advisory Committee: AstraZeneca Pharmaceuticals LP, Cancer Prevention Research Institute of Texas, Immunovia; Consulting **Agreements:** Advance Medical, Astellas Pharma Global Development Inc, BioPharm Communications, Bristol-Myers Squibb Company, Celgene Corporation, EcoR1 Capital, Merck; **Contracted Research:** FibroGen, Gossamer Bio, Halozyme Inc.

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# Hardware/Software Requirements:

A high-speed Internet connection A monitor set to 1280 x 1024 pixels or more Internet Explorer 11 or later, Firefox 56 or later, Chrome 61 or later, Safari 11 or later, Opera 48 or later Adobe Flash Player 27 plug-in or later Adobe Acrobat Reader (Optional) Sound card and speakers for audio

Last review date: April 2019

Expiration date: April 2020

# **Select Publications**

#### Margaret A Tempero, MD

Conroy T et al. Unicancer GI PRODIGE 24/CCTG PA.6 trial: A multicenter international randomized phase III trial of adjuvant mFOLFIRINOX versus gemcitabine (gem) in patients with resected pancreatic ductal adenocarcinomas. *Proc ASCO* 2018; Abstract LBA4001.

Cunningham D et al. Phase III randomized comparison of gemcitabine versus gemcitabine plus capecitabine in patients with advanced pancreatic cancer. *J Clin Oncol* 2009;27(33):5513-8.

Herrmann R et al. Gemcitabine plus capecitabine compared with gemcitabine alone in advanced pancreatic cancer: A randomized, multicenter, phase III trial of the Swiss Group for Clinical Cancer Research and the Central European Cooperative Oncology Group. *J Clin Oncol* 2007;25(16):2212-7.

Kalser MH, Ellenberg SS. Pancreatic cancer: Adjuvant combined radiation and chemotherapy following curative resection. *Arch Surg* 1985;120(8):899-903.

Klinkenbijl JH et al. Adjuvant radiotherapy and 5-fluorouracil after curative resection of cancer of the pancreas and periampullary region: Phase III trial of the EORTC Gastrointestinal Tract Cancer Cooperative Group. *Ann Surg* 1999;230(6):776-82.

Neoptolemos JP et al. Comparison of adjuvant gemcitabine and capecitabine with gemcitabine monotherapy in patients with resected pancreatic cancer (ESPAC-4): A multicentre, open-label, randomised, phase 3 trial. *Lancet* 2017;389(10073): 1011-24.

Neoptolemos JP et al. Adjuvant chemotherapy with fluorouracil plus folinic acid vs gemcitabine following pancreatic cancer resection: A randomized controlled trial. *JAMA* 2010;304(10):1073-81.

Neoptolemos JP et al. A randomized trial of chemoradiotherapy and chemotherapy after resection of pancreatic cancer. *N Engl J Med* 2004;350(12):1200-10.

Oettle H et al. Adjuvant chemotherapy with gemcitabine and long-term outcomes among patients with resected pancreatic cancer: The CONKO-001 randomized trial. *JAMA* 2013;310(14):1473-81.

Regine WF et al. Fluorouracil vs gemcitabine chemotherapy before and after fluorouracil-based chemoradiation following resection of pancreatic adenocarcinoma: A randomized controlled trial. *JAMA* 2008;299(9):1019-26.

Smeenk HG et al. Long-term survival and metastatic pattern of pancreatic and periampullary cancer after adjuvant chemoradiation or observation: Long-term results of EORTC trial 40891. *Ann Surg* 2007;246(5):734-40.

Van Tienhoven G et al. Preoperative chemoradiotherapy versus immediate surgery for resectable and borderline resectable pancreatic cancer (PREOPANC-1): A randomized, controlled, multicenter phase III trial. *Proc ASCO* 2018; Abstract LBA4002.

#### Philip A Philip, MD, PhD

Burris HA III et al. Improvements in survival and clinical benefit with gemcitabine as first-line therapy for patients with advanced pancreas cancer: A randomized trial. *J Clin Oncol* 1997;15(6):2403-13.

Conroy T et al. FOLFIRINOX versus gemcitabine for metastatic pancreatic cancer. N Engl J Med 2011;364(19):1817-25.

Goldstein D et al. *Nab*-paclitaxel plus gemcitabine for metastatic pancreatic cancer: Long-term survival from a phase III trial. *J Natl Cancer Inst* 2015;107(2):pii.

Scheithauer W et al. Dose modification and efficacy of *nab*-paclitaxel plus gemcitabine vs gemcitabine for patients with metastatic pancreatic cancer: Phase III MPACT trial. *J Gastrointest Oncol* 2016;7(3):469-78.

Sohal DPS et al. Metastatic pancreatic cancer: ASCO Clinical Practice Guideline update. J Clin Oncol 2018;36(24):2545-56.

Suker M et al. **FOLFIRINOX for locally advanced pancreatic cancer: A systematic review and patient-level meta-analysis.** *Lancet Oncol* 2016;17(6):801-10.

Von Hoff D et al. Increased survival in pancreatic cancer with *nab*-paclitaxel plus gemcitabine. *N Engl J Med* 2013;369(18):1691-703.

Von Hoff D et al. Randomized phase III study of weekly *nab*-paclitaxel plus gemcitabine versus gemcitabine alone in patients with metastatic adenocarcinoma of the pancreas (MPACT). Gastrointestinal Cancers Symposium 2013;Abstract LBA148.

# Select Publications

#### Eileen M O'Reilly, MD

Conroy T et al. FOLFIRINOX versus gemcitabine for metastatic pancreatic cancer. N Engl J Med 2011;364(19):1817-25.

Gill S et al. PANCREOX: A randomized phase III study of fluorouracil/leucovorin with or without oxaliplatin for second-line advanced pancreatic cancer in patients who have received gemcitabine-based chemotherapy. *J Clin Oncol* 2016;34(32):3914-20.

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Sonbol MB et al. Second-line treatment in patients with pancreatic ductal adenocarcinoma: A meta-analysis. *Cancer* 2017;123(23):4680-6.

Von Hoff DD et al. Increased survival in pancreatic cancer with *nab*-paclitaxel plus gemcitabine. *N Engl J Med* 2013;369(18):1691-703.

Wang-Gillam A et al. Nanoliposomal irinotecan with fluorouracil and folinic acid in metastatic pancreatic cancer after previous gemcitabine-based therapy (NAPOLI-1): A global, randomised, open-label, phase 3 trial. *Lancet* 2016;387(10018):545-57.

Yoo C et al. A randomised phase II study of modified FOLFIRI.3 vs modified FOLFOX as second-line therapy in patients with gemcitabine-refractory advanced pancreatic cancer. *Br J Cancer* 2009;101(10):1658-63.

#### Andrew E Hendifar, MD

Alvarez R et al. Stromal disrupting effects of nab-paclitaxel in pancreatic cancer. Br J Cancer 2013;109(4):926-33.

Beatty GL et al. **CD40** agonists alter tumor stroma and show efficacy against pancreatic carcinoma in mice and humans. *Science* 2011;331(6024):1612-6.

Bekaii-Saab T et al. A phase 1b/II study of cancer stemness inhibitor napabucasin in combination with gemcitabine (gem) & *nab*-paclitaxel (nabptx) in metastatic pancreatic adenocarcinoma (mpdac) patients (pts). Proc ESMO World Congress on Gastrointestinal Cancer 2017; Abstract LBA-002.

Bekaii-Saab T et al. CanStem111P trial: A phase III study of napabucasin (BBI-608) plus *nab*-paclitaxel (*nab*-PTX) with gemcitabine (gem) in adult patients with metastatic pancreatic adenocarcinoma (mPDAC). *Proc ASCO* 2017;Abstract TPS4148.

Bullock AJ et al. Final analysis of stage 1 data from a randomized phase II study of PEGPH20 plus *nab*-paclitaxel/gemcitabine in stage IV previously untreated pancreatic cancer patients (pts), utilizing Ventana companion diagnostic assay. *Proc ASCO* 2016; Abstract 4104.

Hingorani S et al. Randomized phase II study of PEGPH20 plus *nab*-paclitaxel/gemcitabine (PAG) vs AG in patients (Pts) with untreated, metastatic pancreatic ductal adenocarcinoma (mPDA). *Proc ASCO* 2017; Abstract 4008.

Kaufman B et al. **Olaparib monotherapy in patients with advanced cancer and a germline BRCA1/2 mutation.** *J Clin Oncol* 2015;33(3):244-50.

Khelifa S et al. Development of a companion diagnostic assay for tissue hyaluronan detection and treatment with PEGPH20 in metastatic pancreatic ductal adenocarcinoma patients. *Proc ASCO* 2016; Abstract e15749.

Olive KP et al. Inhibition of hedgehog signaling enhances delivery of chemotherapy in a mouse model of pancreatic cancer. *Science* 2009;324(5922):1457-61.

Provenzano PP et al. Enzymatic targeting of the stroma ablates physical barriers to treatment of pancreatic ductal adenocarcinoma. *Cancer Cell* 2012;21(3):418-29.

Sherman MH et al. Vitamin D receptor-mediated stromal reprogramming suppresses pancreatitis and enhances pancreatic cancer therapy. *Cell* 2014;159(1):80-93.