

World Journal of *Gastroenterology*

World J Gastroenterol 2022 December 7; 28(45): 6314-6432



EVIDENCE REVIEW

- 6314 COVID-19 drug-induced liver injury: A recent update of the literature
Saha L, Vij S, Rawat K

REVIEW

- 6328 Gastrointestinal microbiota: A predictor of COVID-19 severity?
Neag MA, Vulturar DM, Gherman D, Burlacu CC, Todea DA, Buzoianu AD

MINIREVIEWS

- 6345 The mononuclear phagocyte system in hepatocellular carcinoma
Qiao DR, Shan GY, Wang S, Cheng JY, Yan WQ, Li HJ
- 6356 Rifabutin as salvage therapy for *Helicobacter pylori* eradication: Cornerstones and novelties
Borraccino AV, Celiberto F, Pricci M, Girardi B, Iannone A, Rendina M, Ierardi E, Di Leo A, Losurdo G
- 6363 Deep learning based radiomics for gastrointestinal cancer diagnosis and treatment: A minireview
Wong PK, Chan IN, Yan HM, Gao S, Wong CH, Yan T, Yao L, Hu Y, Wang ZR, Yu HH

ORIGINAL ARTICLE**Observational Study**

- 6380 Best therapy for the easiest to treat hepatitis C virus genotype 1b-infected patients
Zarebska-Michaluk D, Brzdęk M, Jaroszewicz J, Tudrujek-Zdunek M, Lorenc B, Klapaczyński J, Mazur W, Kazek A, Sitko M, Berak H, Janocha-Litwin J, Dybowska D, Supronowicz Ł, Krygier R, Citko J, Piekarska A, Flisiak R

Randomized Clinical Trial

- 6397 Endoscopic mucosal resection-precutting *vs* conventional endoscopic mucosal resection for sessile colorectal polyps sized 10-20 mm
Zhang XQ, Sang JZ, Xu L, Mao XL, Li B, Zhu WL, Yang XY, Yu CH

META-ANALYSIS

- 6410 Meta-analysis on the epidemiology of gastroesophageal reflux disease in China
Lu TL, Li SR, Zhang JM, Chen CW

CASE REPORT

- 6421 Germline BRCA2 variants in advanced pancreatic acinar cell carcinoma: A case report and review of literature
Lee CL, Holter S, Borgida A, Dodd A, Ramotar S, Grant R, Wasson K, Elimova E, Jang RW, Moore M, Kim TK, Khalili K, Moulton CA, Gallinger S, O'Kane GM, Knox JJ

ABOUT COVER

Editorial Board of *World Journal of Gastroenterology*, Giovanni Domenico De Palma, FACS, FASGE, MD, Full Professor, Department of Clinical Medicine and Surgery, "Federico II" University of Naples, Naples 80131, Italy. giovanni.depalma@unina.it

AIMS AND SCOPE

The primary aim of *World Journal of Gastroenterology* (WJG, *World J Gastroenterol*) is to provide scholars and readers from various fields of gastroenterology and hepatology with a platform to publish high-quality basic and clinical research articles and communicate their research findings online. WJG mainly publishes articles reporting research results and findings obtained in the field of gastroenterology and hepatology and covering a wide range of topics including gastroenterology, hepatology, gastrointestinal endoscopy, gastrointestinal surgery, gastrointestinal oncology, and pediatric gastroenterology.

INDEXING/ABSTRACTING

The WJG is now abstracted and indexed in Science Citation Index Expanded (SCIE, also known as SciSearch®), Current Contents/Clinical Medicine, Journal Citation Reports, Index Medicus, MEDLINE, PubMed, PubMed Central, Scopus, Reference Citation Analysis, China National Knowledge Infrastructure, China Science and Technology Journal Database, and Superstar Journals Database. The 2022 edition of Journal Citation Reports® cites the 2021 impact factor (IF) for WJG as 5.374; IF without journal self cites: 5.187; 5-year IF: 5.715; Journal Citation Indicator: 0.84; Ranking: 31 among 93 journals in gastroenterology and hepatology; and Quartile category: Q2. The WJG's CiteScore for 2021 is 8.1 and Scopus CiteScore rank 2021: Gastroenterology is 18/149.

RESPONSIBLE EDITORS FOR THIS ISSUE

Production Editor: *Ying-Yi Yuan*; Production Department Director: *Xiang Li*; Editorial Office Director: *Jia-Ru Fan*.

NAME OF JOURNAL

World Journal of Gastroenterology

ISSN

ISSN 1007-9327 (print) ISSN 2219-2840 (online)

LAUNCH DATE

October 1, 1995

FREQUENCY

Weekly

EDITORS-IN-CHIEF

Andrzej S Tarnawski

EDITORIAL BOARD MEMBERS

<http://www.wjgnet.com/1007-9327/editorialboard.htm>

PUBLICATION DATE

December 7, 2022

COPYRIGHT

© 2022 Baishideng Publishing Group Inc

INSTRUCTIONS TO AUTHORS

<https://www.wjgnet.com/bpg/gerinfo/204>

GUIDELINES FOR ETHICS DOCUMENTS

<https://www.wjgnet.com/bpg/GerInfo/287>

GUIDELINES FOR NON-NATIVE SPEAKERS OF ENGLISH

<https://www.wjgnet.com/bpg/gerinfo/240>

PUBLICATION ETHICS

<https://www.wjgnet.com/bpg/GerInfo/288>

PUBLICATION MISCONDUCT

<https://www.wjgnet.com/bpg/gerinfo/208>

ARTICLE PROCESSING CHARGE

<https://www.wjgnet.com/bpg/gerinfo/242>

STEPS FOR SUBMITTING MANUSCRIPTS

<https://www.wjgnet.com/bpg/GerInfo/239>

ONLINE SUBMISSION

<https://www.f6publishing.com>

Deep learning based radiomics for gastrointestinal cancer diagnosis and treatment: A minireview

Pak Kin Wong, In Neng Chan, Hao-Ming Yan, Shan Gao, Chi Hong Wong, Tao Yan, Liang Yao, Ying Hu, Zhong-Ren Wang, Hon Ho Yu

Specialty type: Gastroenterology and hepatology

Provenance and peer review: Invited article; Externally peer reviewed.

Peer-review model: Single blind

Peer-review report's scientific quality classification

Grade A (Excellent): 0
Grade B (Very good): B, B
Grade C (Good): 0
Grade D (Fair): 0
Grade E (Poor): 0

P-Reviewer: Gnetti L, Italy; Litvin A, Russia

Received: September 20, 2022

Peer-review started: September 20, 2022

First decision: October 18, 2022

Revised: October 25, 2022

Accepted: November 16, 2022

Article in press: November 16, 2022

Published online: December 7, 2022



Pak Kin Wong, In Neng Chan, Liang Yao, Department of Electromechanical Engineering, University of Macau, Taipa 999078, Macau, China

Hao-Ming Yan, School of Clinical Medicine, China Medical University, Shenyang 110013, Liaoning Province, China

Shan Gao, Department of Gastroenterology, Xiangyang Central Hospital, Affiliated Hospital of Hubei University of Arts and Science, Xiangyang 441021, Hubei Province, China

Chi Hong Wong, Faculty of Medicine, Macau University of Science and Technology, Taipa 999078, Macau, China

Tao Yan, Zhong-Ren Wang, School of Mechanical Engineering, Hubei University of Arts and Science, Xiangyang 441053, Hubei Province, China

Liang Yao, Ying Hu, Shenzhen Institute of Advanced Technology, Chinese Academy of Sciences, Shenzhen 518055, Guangdong Province, China

Hon Ho Yu, Department of Gastroenterology, Kiang Wu Hospital, Macau 999078, China

Corresponding author: Tao Yan, PhD, Instructor, School of Mechanical Engineering, Hubei University of Arts and Science, No. 296 Longzhong Road, Xiangyang 441053, Hubei Province, China. yantao@hbuas.edu.cn

Abstract

Gastrointestinal (GI) cancers are the major cause of cancer-related mortality globally. Medical imaging is an important auxiliary means for the diagnosis, assessment and prognostic prediction of GI cancers. Radiomics is an emerging and effective technology to decipher the encoded information within medical images, and traditional machine learning is the most commonly used tool. Recent advances in deep learning technology have further promoted the development of radiomics. In the field of GI cancer, although there are several surveys on radiomics, there is no specific review on the application of deep-learning-based radiomics (DLR). In this review, a search was conducted on Web of Science, PubMed, and Google Scholar with an emphasis on the application of DLR for GI cancers, including esophageal, gastric, liver, pancreatic, and colorectal cancers. Besides, the challenges and recommendations based on the findings of the review are comprehensively analyzed to advance DLR.

Key Words: Radiomics; Deep learning; Gastrointestinal cancer; Medical imaging

©The Author(s) 2022. Published by Baishideng Publishing Group Inc. All rights reserved.

Core Tip: Radiomics, especially deep-learning-based radiomics (DLR), has revolutionized the diagnosis, assessment and prognosis of gastrointestinal (GI) cancer. This review provides an analysis and status of DLR in GI cancer and identifies future challenges and recommendations.

Citation: Wong PK, Chan IN, Yan HM, Gao S, Wong CH, Yan T, Yao L, Hu Y, Wang ZR, Yu HH. Deep learning based radiomics for gastrointestinal cancer diagnosis and treatment: A minireview. *World J Gastroenterol* 2022; 28(45): 6363-6379

URL: <https://www.wjgnet.com/1007-9327/full/v28/i45/6363.htm>

DOI: <https://dx.doi.org/10.3748/wjg.v28.i45.6363>

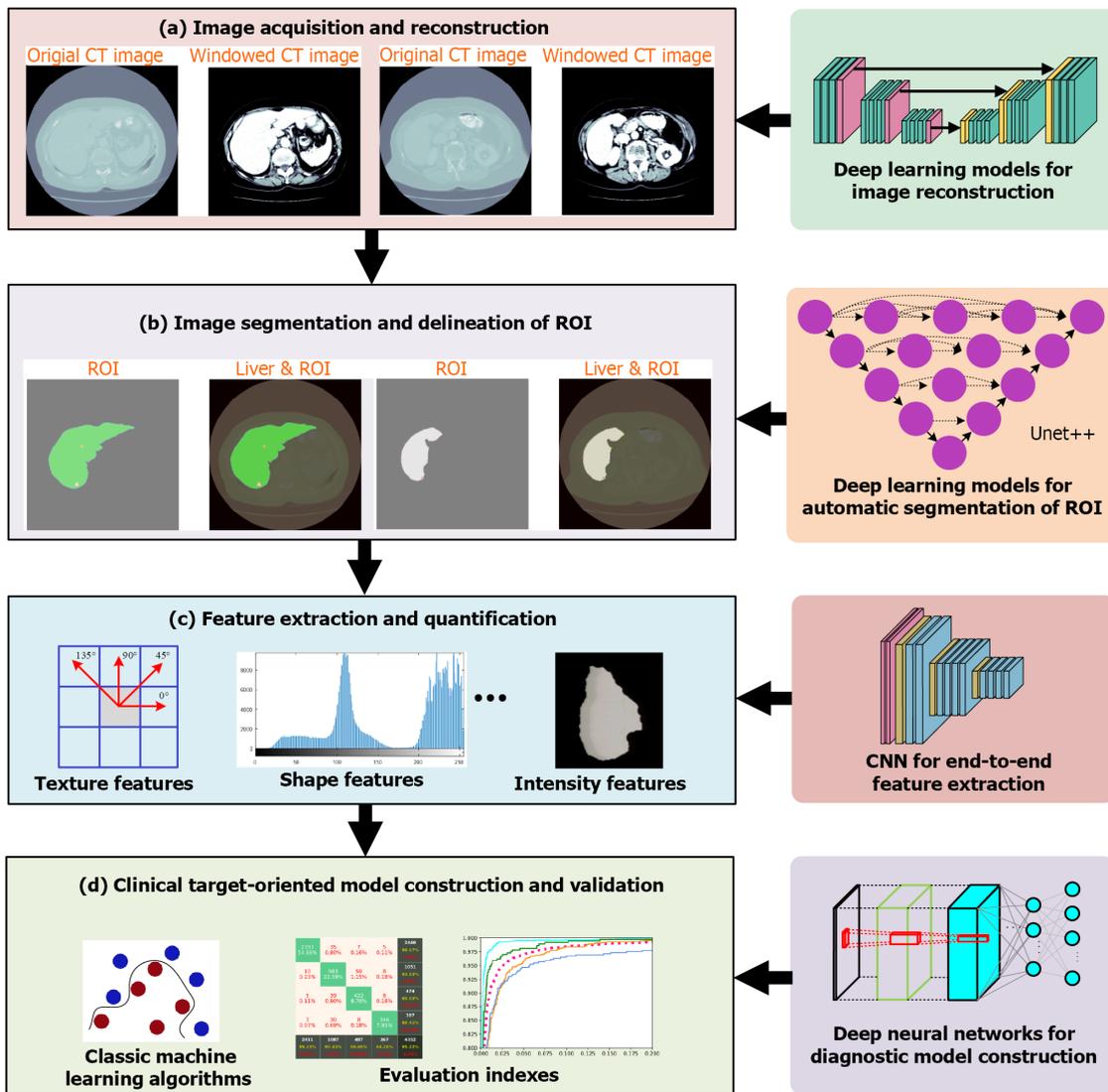
INTRODUCTION

Gastrointestinal (GI) cancers, mainly include colorectal, gastric, liver, esophageal, and pancreatic cancers, and are the leading cause of cancer-related mortality globally[1]. According to CANCER TOMORROW[2], a forecast of the global burden of cancer mortality and incidence, by 2040, new cases of GI cancer and deaths will increase significantly. In recent years, computed tomography (CT), magnetic resonance imaging (MRI), positron emission tomography (PET), ultrasound (US) and other medical imaging techniques have been widely used in GI cancer diagnosis and treatment[3,4]. It is foreseeable that with the increase in GI cancer, the amount of medical imaging data will continue to grow. However, manual reading cannot cope with this growth, and the disparity in expertise among radiologists causes a high rate of missed diagnosis and misdiagnosis. In addition, traditional CT, MRI, PET, US, and other imaging examinations cannot observe changes in tumor heterogeneity, which can provide a better understanding of the causes and progression of cancer[5]. The development of radiomics technology provides new opportunities and methods to solve these dilemmas.

Radiomics is an emerging method for quantitative analysis and prediction of tumor phenotypes using machine learning or statistical models, and was proposed by Lambin *et al*[6] in 2012. In recent years, radiomics has been widely used in GI cancer and showed notable outcomes in tumor characterization, therapy response assessment, and prediction of survival rate after surgery[7-11]. Compared with the conventional method of using only manual inspection, radiomics can extract high-dimensional features that are difficult to be quantitatively described by the doctors from massive radiological images, and to correlate them with clinical and pathological data of patients in order to improve diagnosis and prognostication[12]. The fundamental premise of radiomics is that the developed descriptive models may produce useful prognostic, predictive and diagnostic information. Radiomics can be divided into two main categories: conventional radiomics, also referred to handcrafted radiomics (HCR) and deep-learning-based radiomics (DLR), also referred to as discovery radiomics[13]. Given the benefits of these two approaches, hybrid solutions that mix HCR and DLR also exist.

The HCR workflow is divided into multiple steps: (1) Image acquisition and reconstruction; (2) image segmentation and delineation of region of interest (automatic, semi-automatic, or manual delineation); (3) feature extraction and quantification. This is the core step of HCRs. The extracted features are mainly handcrafted features (also referred as pre-designed features), including shape, texture and intensity features. Some features may be highly correlated or redundant, so feature dimensionality reduction is an important step in feature analysis; and (4) Clinical target-oriented model building and validation. At this step, classic machine learning algorithms are usually used to develop high-precision and high-efficiency prediction models, and the models are trained and validated with sufficient data. The workflow of HCR is depicted in Figure 1.

Although HCR has been widely adopted in GI cancer and has achieved significant results, it has some deficiencies, such as low degree of automation and standardization, cumbersome and time-consuming feature extraction steps, and insufficient robustness and accuracy. Recently, deep learning, a promising technique in characterization of medical images, has gained much attention[14-17]. Many researchers have adopted DLR to overcome the limitations of conventional radiomics[18-22]. DL refers to a broad class of algorithms rather than a specific model. As long as a deep neural network structure is used to represent features at a deeper level, it can be called DL model. One of the most popular DL models used in medical imaging is convolutional neural networks (CNNs), which can automatically learn representative features from medical images. The use of CNNs in radiomics makes it easy to build an end-to-end feature extraction process, thereby avoiding the tedious and handcrafted feature extraction process. CNNs can also be used in image reconstruction and segmentation to improve the automation level of



DOI: 10.3748/wjg.v28.i45.6363 Copyright ©The Author(s) 2022.

Figure 1 Overview of steps in handcrafted radiomics workflow and steps that can be done with deep learning models. ROI: Region of interest.

HCR, and the accuracy and reliability of diagnosis and prediction (Figure 1).

DL techniques are revolutionizing radiomics. In the field of GI cancer diagnosis and treatment, while there are several surveys on HCR[9-11], there is no specific review on the application of DLR. To provide a comprehensive overview of DLR in GI cancer, the performance of DLR in gastroenterology is summarized in this review, with an emphasis on the diagnosis and treatment of GI cancers, including esophageal, gastric, liver, pancreatic, and colorectal cancers. The original contributions to knowledge of this review are: (1) A unique interdisciplinary viewpoint on radiomics by discussing state-of-the-art DLR solutions; and (2) the challenges and recommendations based on the findings of the review are thoroughly analyzed to advance the field.

DLR FOR ESOPHAGEAL CANCER

Esophageal cancer is the seventh most prevalent form of cancer and the sixth most lethal cancer globally [1], and it is classified into esophageal squamous cell carcinoma (ESCC) or esophageal adenocarcinoma according to the type of cells. In consideration of the low overall 5-year survival rate of patients and the variation in responsiveness of patients to the current treatments such as neoadjuvant chemotherapy (NAC) and neoadjuvant chemoradiotherapy (NCRT) due to tumor heterogeneity, it is vital to have accurate diagnosis, pretreatment evaluation and survival rate prediction. The number of DLR studies regarding esophageal cancer has been growing, with most of the studies exploring treatment response, and the others investigating disease classification and survival rate prediction.

An important preoperative topic of esophageal cancer is diagnosis, yet the number of relevant DLR studies for diagnosis is minimal. Takeuchi *et al*[23] fine-tuned VGG16 to develop a DLR model for the diagnosis of esophageal cancer from CT scans, and its performance was comparable to that of the radiologists during testing, with a higher accuracy of 84.2% and specificity of 90.0%.

Response to treatment, especially NAC and NCRT, is one of the most popular research interests in the field of esophageal cancer. Hu *et al*[24] designed a CT-based model to predict the pathological complete response to NCRT of patients with ESCC using DL features, in which the support vector machine (SVM) classifier executed the classification action. The DL features were extracted using pretrained models, and the optimal one used ResNet50 that achieved an area under the receiver operating characteristic curve (AUC) of 0.805 and accuracy of 77.1% for the testing cohort, which achieved better results than using handcrafted features. Ypsilantis *et al*[25] designed a 3S-CNN model that extracted DL features from PET scans and predicted whether the patient with esophageal cancer was non-responsive to NAC. This model was also compared with other competitive machine learning algorithms and results showed that it surpassed the other models with an average specificity, sensitivity and accuracy of 80.7%, 81.6%, and 73.4% respectively. Amyar *et al*[26] presented a novel 3D CNN model named 3D RPET-NET that predicted the response to CRT using esophageal cancer images of FDG-PET scans, and a comparative analysis with other approaches in the literature was also carried out. Three-dimensional RPET-NET obtained the best results with an accuracy of around 72% and even reached 75% when using tumor volume with an isotropic margin of 2 cm. Li *et al*[27] proposed a CT-based 3D DLR model (3D-DLRM), which was modified from ResNet34. Its aim was to predict whether patients with locally advanced thoracic ESCC had an objective or nonobjective response to concurrent CRT, achieving a validation AUC and positive predictive value of 0.833 and 100%, respectively. They also evaluated a model integrating the 3D-DLRM with clinical selected factors that even outperformed the individual 3D-DLRM, reaching a validation AUC of 0.861.

Other research interests of esophageal cancer include patient survival rate prediction. Wang *et al*[28] compared the use of an HCR model, DLR model and DLR nomogram for the prediction of the survival rate of esophageal cancer patients after 3 years of CRT, in which DL features were extracted and selected by DenseNet-169 to build the DLR model. This DLR nomogram attained the highest validation AUC of 0.942 and Harrell's concordance index (C-index) of 0.784, surpassing the results produced by the sole use of HCR and DLR models. Yang *et al*[29] proposed a 3D-CNN model based on ResNet18 to predict esophageal cancer patient survival rate using PET scans. The model was initially pretrained to classify abnormal and healthy esophagus, and then trained to classify whether patients survived or expired within a year after diagnosis in the second stage, and the model obtained an AUC of 0.738. Gong *et al* [30] developed a hybrid radiomics nomogram to predict local recurrence-free survival (LRFS) of locally advanced ESCC patients who received definitive CRT from contrast-enhanced CT (CECT) scans, and it was combined with radiomic features, features extracted by 3D-DenseNet and prognostic clinical risk factors. The final model achieved a C-index of 0.76 for its external validation set, indicating the effectiveness of the addition of DL features for better prediction performances.

Some studies also discuss the application of DLR to prediction of lymph node (LN) metastasis, which is an effective prognosis factor of ESCC. Wu *et al*[31] built a model involving HCR, computer vision and DLR to predict the LN status of ESCC patients, and they also constructed two simpler models for efficacy comparison, and they exploited Convolution Neural Network-Fast (CNN-F) to extract DL features from CT images. The model with all signatures involved performed the best with C-statistic of 0.875, 0.874, and 0.840 for training, internal validation, and external validation cohorts, and those demonstrate its satisfactory discriminative ability.

The studies about the application of DLR for esophageal cancer are summarized in [Table 1](#).

DLR FOR GASTRIC CANCER

Gastric cancer (GC) is the fifth most prevalent form of cancer and the fourth most lethal cancer globally [1]. To ameliorate the low survival rate of patients, early diagnosis of disease and systematic treatment methods are necessary. The application of DLR in GC has been a promising area for research with a rising number of relevant studies published every year, that aim to tackle or refine the existing concerns regarding GC.

Many studies focused on prediction of treatment response of patients. Cui *et al*[32] constructed a pretreatment venous-phase CT-based DLR nomogram that combined handcrafted features, DL features and remarkable clinicopathological factors to identify locally advanced GC patients with good response to NAC. The nomogram achieved better than the clinical model and the separate use of two features that were built for comparison, attaining C-index values of 0.829, 0.804, and 0.827 in its internal validation cohort and two external validation cohorts, respectively. Li *et al*[33] developed a combined artificial intelligence (AI) model that incorporated feature outputs from HCR and DLR models, which aimed to determine whether the patients had signet ring cell carcinoma (SRCC) of GC and predict survival and treatment response to postoperative chemotherapy from CECT images. They also compared its efficacy with the clinical, HCR and DLR models, and the AI model obtained the best

Table 1 Summary of studies using deep-learning-based radiomics for esophageal cancer

Ref.	Imaging	Study design	Study aim	DL model	Dataset	Outcomes
Takeuchi <i>et al</i> [23], 2021	CT	Retrospective	Detection of esophageal cancer	VGG16	1646 CT images (1500 images for training and validation, 146 for testing)	Accuracy: 84.2%; F value: 74.2%; Sensitivity: 71.1%; Specificity: 90%) in test set
Hu <i>et al</i> [24], 2021	CT	Retrospective	Evaluation of response to NCRT to ESCC	ResNet50	231 patients (161 in training cohort, 70 in testing cohort)	AUC: 0.805; C-index: 0.805; Accuracy: 77.1%; Sensitivity: 83.9%; Specificity: 71.8%) for the testing cohort
Ypsilantis <i>et al</i> [25], 2015	PET	Retrospective	Prediction of response to NAC in patients with esophageal cancer	3S-CNN	107 patients	Sensitivity: 80.7%; Specificity: 81.6%; Accuracy: 73.4%
Amyar <i>et al</i> [26], 2019	PET	Retrospective	Prediction of response to radio-chemotherapy in patients with esophageal cancer	3D RPET-NET	97 patients	Accuracy: 75.0%; Sensitivity: 76.0%; Specificity: 74.0%; AUC: 0.74
Li <i>et al</i> [27], 2021	CT	Retrospective	Prediction of treatment response to CCRT among patients with locally advanced TESCC	ResNet34	306 patients (203 in training cohort, 103 in validation cohort)	AUC: 0.833; PPV: 100%
Wang <i>et al</i> [28], 2022	CT	Retrospective	Prediction of survival rates for patients with esophageal cancer after 3 yr with chemoradio-therapy	DenseNet- 169	154 patients (116 in training cohort, 38 in validation cohort)	AUC: 0.942; C-index: 0.784
Yang <i>et al</i> [29], 2019	PET	Retrospective	Identification of esophageal cancer patients with poor prognosis	3D-CNN based on ResNet18	1107 scans	AUC: 0.738
Gong <i>et al</i> [30], 2022	CECT	Retrospective	Prediction of LRFS in esophageal cancer patients after 1 yr of definitive chemoradio-therapy	3D-Densenet	397 patients	C-index: 0.76
Wu <i>et al</i> [31], 2019	CT	Retrospective	Prediction of LN status of patients with ESCC	CNN-F	411 patients	C-index: 0.840

CT: Computed tomography; PET: Positron emission tomography; CECT: Contrast-enhanced computed tomography; NCRT: Neoadjuvant chemoradiotherapy; NAC: Neoadjuvant chemotherapy; ESCC: Esophageal squamous cell carcinoma; CCRT: Concurrent chemo-radiation therapy; TESCC: Thoracic esophageal squamous cell carcinoma; LRFS: Local recurrence-free survival; LN: Lymph node; CNN-F: Convolutional neural network-fast; AUC: Area under the receiver operating characteristic curve; PPV: Positive predictive value; DLR: Deep-learning-based radiomics.

results with an AUC of 0.786 and accuracy of 71.6% for diagnosing SRCC for the test cohort. The AI model also evaluated that SRCC patients with higher risks had shorter median overall survival (OS) and insignificant improvements in median OS after receiving adjuvant chemotherapy than those of lower risk, indicating its good capability to predict survival and response to treatment. Tan *et al*[34] built a dual-energy CT delta radiomics model to predict the treatment response to chemotherapy of patients with far-advanced GC. They developed a V-Net segmentation model, and the application of this semi-automatic segmentation model to the delta radiomics model shortened the diagnostic time and achieved better results in terms of mean AUC (0.728 *vs* 0.687 in the testing cohort, 0.828 *vs* 0.749 in the independent validation cohort) than using manual segmentation.

Survival rate prediction is also a popular topic for DLR of GC. Hao *et al*[35] combined clinical variables, radiomic features and DL features to build a CT-based prediction Cox proportional-hazard model, which served to predict the OS and progression-free survival (PFS) of patients with GC. The model acquired the highest C-index of 0.783 and 0.770 for OS and PFS when using postoperative clinical variables, and the most dominant variables for survival prediction were identified as important prognostic factors in the subsequent survival analysis. Some studies only made use of DL techniques to build predictive models for similar purposes. Zhang *et al*[36] proposed a multi-focus and multi-level fusion feature pyramid network (MMF-FPN) to predict OS risks of GC patients from CT images, and other models using existing methods in the literature were used for comparison. The experimental results showed that MMF-FPN was the finest model that attained the highest C-indexes (validation: 0.74, testing: 0.76) and hazard ratios (validation: 3.50, testing: 9.46).

To do a preoperative prediction of early recurrence of patients with advanced GC from CT images, Zhang *et al*[37] designed a radiomics nomogram that utilized clinical characteristics and radiomics signature containing handcrafted and DL features as input. The radiomics nomogram reached an AUC and accuracy of 0.806 and 0.723, respectively, while having considerable *k* values of 0.932 for both intra-

and inter-reader agreement, exceeding the results obtained by the radiomics signature and clinical modal built for comparison.

Accurate prediction of LN status of GC, which is a remarkable prognostic factor, is of importance to determine the appropriate treatment. Guan *et al*[38] explored the efficacy of using different DL models to extract features and machine learning classifiers (*i.e.*, SVM and random forest) to build a CT-based predictive model for the evaluation of LN status. Other models using radiomic features and integrated features were built for comparison, and the best model was ResNet50-RF with an AUC and accuracy of 0.9803 and 98.10%, respectively. A nomogram based on DL feature scores and clinical risk factors was also developed and a higher AUC of 0.9914 was achieved in the testing cohort. Dong *et al*[39] proposed a similar DLR nomogram to evaluate the number of LN metastases of locally advanced GC patients before surgery, in which radiomics signatures that contained handcrafted and DL features and clinical characteristics were used. The performance of the model was evaluated with four validation sets; three of which were collected from China and one from Italy. The model showed its good discriminative capability to identify N-staging of GC with higher C-indexes of 0.797 in the validation sets from China and 0.822 in the set from Italy, and it outperformed other predictors such as clinical models and single signatures. To predict the LN status and prognosis of patients, the dual-energy CT-based DLR nomogram created by Li *et al*[40] incorporated CT-reported LN and two radiomics signatures for arterial-phase and venous-phase CT images, in which DL features were extracted *via* CNN. The nomogram performed better and gained a higher AUC of 0.82 than the clinical model built alongside for comparative analysis, and the associated prognosis prediction was satisfactory in terms of PFS (C-index: 0.64) and OS (C-index: 0.67). Jin *et al*[41] developed a DLR model that adopted ResNet-18 to evaluate the LN status in nodal stations using CECT, and the high value of the median AUC of the 11 stations (0.876) proved the excellent prediction ability of the model. The authors attempted to build a nomogram combining the DL features with clinical features, but no significant improvements in the results were observed.

Other prognostic factors of GC have also been investigated in previous studies. Sun *et al*[42] exploited DL techniques to build a CT-based radiomics nomogram for evaluating the status of serosal invasion of advanced GC patients. Three radiomics signatures were generated based on the three phases of CT images with their DL features extracted using CNNs, and they were integrated with clinical characteristics to form the nomogram. The final model outperformed other models, such as clinical and phenotypic models, and its AUC for test sets I and II was 0.87 and 0.90, respectively. Li *et al*[43] compared the use of DL features and radiomic features to create a CECT-based GC risk (GRISK) model using similar procedures for the prediction of the status of lymphovascular invasion in patients with localized GC. The team explored the use of deep transfer learning models to build a gastric imaging marker, in which five pretrained models and an auto-encoder were utilized for feature extraction and reduction, respectively. Then, it was integrated with patient clinical and radiological characteristics to construct its own GRISK model. The GRISK model with deep transfer learning gastric imaging marker obtained comparable AUC (0.722 *vs* 0.725) and accuracy (0.671 *vs* 0.710) with the other model with the radiomics gastric imaging marker but did not surpass the latter model.

The studies investigating the usage of DLR for GC are summarized in [Table 2](#).

DLRs FOR LIVER CANCER

Primary liver cancer is the sixth most prevalent form of cancer and the third most lethal cancer globally, and some of its common phenotypes are hepatocellular carcinoma (HCC) and intrahepatic cholangiocarcinoma[1]. Taking the high mortality caused by this disease into account, the clinical application of early diagnosis, individualized evaluation and prognosis prediction are valued. The exploitation of DLR technology in liver cancer has been rapidly developing, and various solutions for the issues in different phases of diagnosis and treatment are emerging.

Computer-aided diagnosis does not only aid radiologists such as shortening the diagnosis time, but also allow them to evaluate appropriate treatments at earlier stages of liver cancer. Ding *et al*[44] constructed a CT-based DLR model that fused a radiomics signature and a DL model, to differentiate HCC into low or high grade. The DL model was an alteration of VGG19 and it performed better than the radiomics signature, with better AUC (0.7513 *vs* 0.7475) and accuracy (66.31% *vs* 65.78%). The fused DLR model was the optimal model with observable improvements in the results, achieving an AUC of 0.8042 and accuracy of 72.73%.

Accurate prediction of patient response to different therapies is critical to realize personalized treatment at different stages of HCC. Peng *et al*[45] developed a multi-class DL model from ResNet50 to predict four treatment responses to transarterial chemoembolization (TACE) therapy of HCC patients using CECT scans. Its performance was assessed using confusion matrices and receiver operating characteristic curves, and the model attained an AUC over 0.90 for all four classes in both validation sets, and accuracies of 85.1% and 82.8% for validation sets 1 and 2, respectively. In the next year, they combined conventional radiomics and DL to build a new CECT-based DLR model that served to predict the initial treatment response to TACE of HCC patients preoperatively[46]. Different from their prior

Table 2 Summary of studies using deep-learning-based radiomics for gastric cancer

Ref.	Imaging	Study design	Study aim	DL model	Dataset	Outcomes
Cui <i>et al</i> [32], 2022	CT	Retrospective	Prediction of response to NAC in patients with LAGC	DenseNet-121	719 patients	C-index: 0.829
Li <i>et al</i> [33], 2022	CT	Retrospective	Diagnosis and prediction of chemotherapy response to SRCC patients	Modified U-Net	855 patients (598 in training cohort; 257 in testing cohort)	For diagnosis, AUC: 0.786; accuracy: 71.6%; sensitivity: 77.3%; specificity: 69.2% for testing cohort
Tan <i>et al</i> [34], 2020	CT	Retrospective	Prediction of response to chemotherapy in patients with gastric cancer	V-Net	116 patients	Mean AUC: 0.728 (testing cohort); 0.828 (validation cohort) when using semi-segmentation
Hao <i>et al</i> [35], 2021	CT	Retrospective	Prediction of OS and PFS after gastrectomy; evaluation of effects of variables on survival prediction	Attention-guided VAE	1061 patients (743 for training; 318 for testing)	C-index of OS: 0.783; C-index of PFS: 0.770 when only using postoperative variables
Zhang <i>et al</i> [36], 2021	CT	Retrospective	Prediction of OS risks of patients with gastric cancer	MMF-FPN	640 patients (337 in training set; 181 in validation set; 122 in test set)	C-index: 0.76; hazard ratio: 9.46 in test set
Zhang <i>et al</i> [37], 2020	CT	Retrospective	Prediction of early recurrence of patients with AGC	DCNNs	669 patients	AUC: 0.806; accuracy: 0.723; sensitivity: 0.827; specificity: 0.667
Guan <i>et al</i> [38], 2022	CT	Retrospective	Prediction of preoperative status of LNM of gastric cancer patients	ResNet50-RF	347 patients (242 for training; 105 for testing)	AUC: 0.9803; accuracy: 98.10%; sensitivity: 98.39%; specificity: 0.9767% for testing of ResNet50-RF
Dong <i>et al</i> [39], 2020	CT	Retrospective	Prediction of the number of LNM in LAGC	DenseNet-201	730 patients	C-index: 0.822 in validation set
Li <i>et al</i> [40], 2020	CT	Retrospective	Prediction of LNM and prognosis in gastric cancer patients	DCNNs	204 patients (136 in training set, 68 in test set)	AUC: 0.82 in test set; C-index of OS: 0.67; C-index of PFS: 0.64
Jin <i>et al</i> [41], 2021	CT	Retrospective	Prediction of LNM status in LN stations of gastric cancer patients	ResNet-18	1699 patients	Median AUC: 0.876; median Sensitivity: 0.743; median Specificity: 0.936 in validation cohort
Sun <i>et al</i> [42], 2020	CT	Stage I: Retrospective; stage II: Validation	Prediction of serosa invasion of AGC patients	DCNNs	572 patients (252 in training set, 176 in test set I, 144 in test set II)	AUC: 0.87; accuracy: 80%; sensitivity: 0.73; specificity: 0.85 in test set I. AUC: 0.90; accuracy: 85%; sensitivity: 0.75; specificity: 0.93 in test set II
Li <i>et al</i> [43], 2022	CT	Retrospective	Evaluation of lymphovascular invasion of localized gastric cancer patients	SqueezeNet, ResNet50, Inception V3, VGG19, DeepLoc	1062 patients (728 for training, 334 for testing)	AUC: 0.725; sensitivity: 73.2%; specificity: 60.3%; accuracy: 71.0% for radiomics GRISK model (final model) in testing cohort

CT: Computed tomography; NAC: Neoadjuvant chemotherapy; LAGC: Locally advanced gastric cancer; SRCC: Signet ring cell carcinoma; OS: Overall survival; PFS: Progression-free survival; AGC: Advanced gastric cancer; LNM: Lymph node metastasis; LN: Lymph node; DCNNs: Deep convolutional neural networks; MMF-FPN: Multi-focus and multi-level fusion feature pyramid network; UC: Area under the receiver operating characteristic curve; C-index: Harrell's concordance index; GRISK: Gastric Risk; RF: Random forest; VAE: Variational auto-encoder; DLR: Deep-learning-based radiomics.

work, they designed their own CNN for feature extraction and prediction, and the DL model was integrated with five radiomics models built with different classic machine learning algorithms or tumor size feature to build integrated models for efficacy comparison. The DL model outperformed all individual radiomics models with an AUC of 0.972, while all integrated models yielded higher values of AUC than merely using DL model. The combination of DL with random forest classifier obtained the highest AUC of 0.994.

Survival prediction is also an important research area to facilitate individualized HCC treatment. To predict the OS of HCC patients who were treated with stereotactic body radiation therapy, Wei *et al* [47] established a CECT-based DL network model that comprised two variational-autoencoder-based survival models and one CNN-based model for extracting radiomic features, clinical features and CT features. The performance of the separate models and the integrated radiomics model using either DL network or Cox hazard model was compared by C-index, in which the integrated model produced the

highest C-index of 0.650 in repeated cross-validation among all models. Liu *et al*[48] developed two separate DLR models to differentiate HCC patients who received radiofrequency ablation (RFA) or surgical resection into high or low risks using CEUS images, and the corresponding radiomics signatures were built. Afterwards, two radiomics nomograms were constructed by combining the signatures with clinical variables to predict the 2-year PFS of patients and both models. Both DLR models achieved satisfactory values of C-index (0.726 for RFA, 0.741 for surgical resection). The good agreement of the survival predictions of the nomograms was demonstrated from the calibration curves.

Postoperative recurrence of cancer is one of the primary causes of death, which extends to the increase in recurrence risk assessment using DLR. To predict the early recurrence of HCC patients using multi-phase CECT scans, Wang *et al*[49] explored the predictive ability of various kinds of models, and they included a DLR model based on ResNet, a clinical model extracting features from clinical data and three combined CNN-based models of different structures. Experimental results demonstrated that the integration of DL features and clinical features improved the prediction accuracy, and one combined model obtained the highest AUC of 0.825. The team improved their study by comparing the DL model with a conventional radiomics model, and one more combined model of another structure was added to the comparative analysis of their previous work[50]. The DL model performed better than the radiomics model with an average AUC of 0.7233 and accuracy of 69.52%, while one of the combined models surpassed the rest in the comparative analysis and reached 0.8248 and 78.66% in its average AUC and accuracy, respectively. They also investigated the effect of attaching a joint loss function to the best model on the average AUC and accuracy, and the two metrics were improved to 0.8331 and 80.49%. He *et al*[51] presented an intelligent-augmented DL model for Risk Assessment of Post Liver Transplantation (i-RAPIT) model in their study, which was a multi-network model that estimated the recurrence risk of HCC patients after liver transplantation. The i-RAPIT model was composed of two deep CapsNet networks for feature extraction from MR and pathological images, and a natural-language-processing-based radial basis function (NLP-based RBF) for extracting clinical features. Before the MR images were entered into the model, U-Net was also exploited for tumor and liver detection in the images. The model achieved a total accuracy of 82%, and AUC of 0.87 and F-1 score of 84% when comparing with other network combinations.

Early detection of microvascular and macrovascular invasion is another practical approach to select the proper therapy for HCC patients and reduce mortality. Jiang *et al*[52] adopted 3D-CNN to build a CT-based DL model for predicting the status of microvascular invasion of HCC patients, and three models based on radiomics features, radiologic features, and integration of the two kinds of features and clinical characteristics was also used for comparison. The results produced by the four models were excellent, with the DL model achieving better results for a few metrics such as AUC (0.906), and sensitivity (93.2%) in the validation set. Wang *et al*[53] devised a new DL model named MVI-Mind that consisted of a light-weight transformer for segmentation and a CNN for prediction of microvascular invasion, and several DL techniques were used to compare the proposed methods. The MVI-Mind attained highest mean intersection over union of 0.9006 and accuracy of 99.47% as compared with other DL segmentation algorithms, and it maintained its superiority in prediction and obtained AUC values of 0.9223, 0.8962, and 0.9100 for arterial phase, portal venous phase and delayed period CT images, respectively. For estimating the status of macrovascular invasion using CT scans, Fu *et al*[54] utilized the concept of multi-task DL neural network (MTnet) to build predictive models. Radiomic features from CT images, clinical and radiological factors were fused to construct the proposed model, and it was modified from U-Net that contained modules engaged in tumor segmentation, feature extraction and prediction. It exhibited the most outstanding performance with an AUC of 0.836 among all models built for comparison.

The studies investigating the implementation of DLR for liver cancer are summarized in [Table 3](#).

DLR FOR PANCREATIC CANCER

Pancreatic cancer is the seventh most deadly cancer worldwide, in which pancreatic adenocarcinoma or pancreatic ductal adenocarcinoma (PDAC) are the most prevalent, accounting for the high mortality rate [1]. The number of deaths caused by this disease is almost equivalent to the number of cases due to the overall poor prognosis, so the introduction of advanced AI technologies is essential and urgent to rectify the situation. In these few years, the field of DLR in pancreatic cancer has flourished and more critical issues such as disease differentiation and survival prediction have been discussed.

Achieving an accurate diagnosis of PDAC gives a great contribution to avoiding false predictions and improving the survival outcomes of patients. For distinguishing between PDAC and autoimmune pancreatitis using CT scans, Ziegelmeier *et al*[55] developed a DLR model that utilized VGG19 to extract DL features, and its efficacy was compared with a model trained on handcrafted radiomic features. The former model performed better with higher mean values in AUC, sensitivity, and specificity (0.90, 89% and 83%) over the cross-validation procedure. Liao *et al*[56] used a DL model based on the coarse-to-fine network architecture search (C2FNAS) to perform segmentation of CECT images for radiomic feature extraction, and they were used for training the machine learning model for

Table 3 Summary of studies using deep-learning-based radiomics for liver cancer

Ref.	Imaging	Study design	Study aim	DL model	Dataset	Outcomes
Ding <i>et al</i> [44], 2021	CT	Retrospective	Evaluation of HCC differentiation	VGG19	1234 patients (799 in training cohort, 248 in validation cohort; 187 in independent testing cohort)	AUC: 0.8042; accuracy: 72.73%; sensitivity: 70.75%; specificity: 75.31% in testing cohort for the fused DLRs model
Peng <i>et al</i> [45], 2020	CT	Retrospective	Prediction of different treatment responses to TACE in HCC patients	ResNet50	789 patients (562 in training cohort; 89 and 138 in validation cohorts 1 and 2)	Accuracy: 85.1% in validation cohort 1; accuracy: 82.8% in validation cohort 2
Peng <i>et al</i> [46], 2021	CT	Retrospective	Prediction of initial response to TACE in HCC patients	CNN	310 patients (139 in training cohort; 171 in validation cohort)	AUC: 0.994
Wei <i>et al</i> [47], 2021	CT	Retrospective	Prediction of OS of HCC patients treated with SBRT	CNN	167 patients	C-index: 0.650 in cross validation
Liu <i>et al</i> [48], 2020	US	Retrospective	Prediction of PFS of HCC patients treated with RFA or surgical resection	CNN	214 RFA patients (149 for training; 65 for validation), 205 SR patients (144 for training; 61 for validation)	C-index of RFA: 0.726; C-index of surgical resection: 0.726
Wang <i>et al</i> [49], 2019	CT	Retrospective	Prediction of early recurrence of HCC patients	ResNet	167 patients	AUC of best model: 0.825
Wang <i>et al</i> [50], 2020	CT	Retrospective	Prediction of early recurrence of HCC patients	ResNet	167 patients	For the best model with joint loss function, AUC: 0.8331; accuracy: 80.49%
He <i>et al</i> [51], 2021	MRI and pathological data	Retrospective	Evaluation of HCC recurrence risk of liver transplantation recipients	U-net, CapsNet	109 patients (87 for training; 22 for testing)	Total accuracy: 82%; recall: 80%; precision: 89%; AUC: 0.87; F-1 score: 84%
Jiang <i>et al</i> [52], 2021	CT	Retrospective	Prediction of microvascular invasion status of HCC patients	3D-CNN	405 patients (324 in training set, 81 in validation set)	AUC: 0.906; sensitivity: 75.7%; specificity: 93.2%; accuracy: 85.2%; F-1 score: 87.2% in validation set
Wang <i>et al</i> [53], 2022	CT	Retrospective	Prediction of microvascular invasion status of HCC patients	Transformer, CNN	138 patients	For arterial phase images in validation set, AUC: 0.9223; Average accuracy: 86.78%
Fu <i>et al</i> [54], 2021	CT	Retrospective	Prediction of macrovascular invasion status in HCC patients	Modified U-Net	366 patients (281 in training cohort, 85 in validation cohort)	AUC: 0.836 in validation cohort

CT: Computed tomography; MRI: Magnetic resonance imaging; US: Ultrasound; HCC: Hepatocellular carcinoma; TACE: Transarterial chemoembolization; PFS: Progression-free survival; RFA: Radiofrequency ablation; SBRT: Stereotactic body radiation therapy; CNN: Convolutional neural network; AUC: Area under the receiver operating characteristic curve; C-index: Harrell's concordance index; DLR: Deep-learning-based radiomics.

prediction. The DL segmentation model obtained a mean Dice score of 0.773 for segmentation while the prediction model yielded an AUC of 0.960 when distinguishing between PDAC and the control group (non-cancerous diseases and normal pancreas). Tong *et al* [57] constructed a ResNet-50-based DLRs model to classify PDAC and chronic pancreatitis patients from CEUS images, and the outputs were the probability of being PDAC or chronic pancreatitis, and heatmaps with highlighted regions that displayed the detected lesions. A two-round reader study was conducted to test the effectiveness of the model. The model achieved an AUC of 0.967 and 0.953 in two validation sets and outperformed the radiologists in the first round, while radiologists could obtain higher accuracies in determining the disease with the aid of the model in the second round.

Prediction of treatment response is also a critical aspect in the field of DLR in pancreatic cancer. Watson *et al* [58] built a CNN model based on LeNet to classify, using CT scans, whether PDAC patients had a pathological response or no response to NAC. It was compared with two models: a hybrid DL model that had the same architecture as the pure DL model but captured both CT image features and one clinical feature [\geq to 10% decrease in carbohydrate antigen (CA)-19], and a CA-19 model only taking in the feature regarding CA-19 decrease. Both DL models could produce superior results than the CA-19 model, and the hybrid DL model obtained a slightly higher AUC than the pure DL model (0.784 *vs* 0.738).

Survival prediction is another vital feature of PDAC that occupies a substantial portion of the existing DLR studies. Muhammad *et al* [59] designed a CNN architecture modified from AlexNet to evaluate the survival risk of PDAC patients that received radiomic features extracted from CECT images, and the

model reached a C-index of 0.85, indicating itself as a good survival model. Zhang *et al*[60] also made use of a CNN that was pretrained with non-small cell lung cancer images to construct their CT-based survival model for patients with resectable PDAC, in which a modified loss function was used. The proposed model accomplished finer prognostic predictions than the conventional radiomic model with an index of prediction accuracy of 11.81% and C-index of 0.651. They released another paper in the same year and compared the efficacy of DL and radiomic features from CECT images by feeding them separately to a random forest classifier to build a DLR model for predicting OS[61]. Similar to their prior work, the DL features were extracted by a pretrained CNN model but with a different structure. The model that used DL features attained an AUC of 0.81, which was higher than the other model based on radiomic features and gained a hazard ratio of 1.38 when the respective risk scores (predicted probabilities of deaths) were tested in survival analyses. Later, they further modified their previous DLR model to a risk score-based feature fusion model to predict 2-year OS[62]. Two small models based on DL and radiomic features separately were embodied in the framework to generate their corresponding risk scores, and these risk scores were used to train the main prediction model. The performance of the proposed model was later assessed with other models using different feature reduction techniques, and the risk score model achieved the highest AUC of 0.84. Yao *et al*[63] devised a new multi-task network model to perform both survival and tumor surgical margin prediction of resectable PDAC patients simultaneously using multi-phase CECT scans. Inside the model, a 3D-CNN model incorporated with a nnUNet for pancreas segmentation was exploited for the margin prediction part, while the combination of 3D-ResNet18 and Contrast-Enhanced 3D Convolutional Long Short-Term Memory (CE-ConvLSTM) network was responsible for survival prediction. The model achieved the results exceeding all other deep models in the comparative analysis, which yielded a C-index of 0.705 in predicting survival outcome and a balanced accuracy of 73.6% in determining the resection margin. They revised their preliminary work by incorporating pancreatic anatomical features into the model and switching to implement an automatically self-learning segmentation method that used 3D UNet as the network architecture and nnUNet as the backbone model for training[63]. The new model attained the highest survival C-index of 0.667 and balanced accuracy of 67.1% for resection margin prediction among all the models including their previous model and other DL and radiomics models.

LN metastasis also possesses a high prognostic value in pancreatic cancer and it is noteworthy to have an early and accurate prediction of its status. An *et al*[64] developed a DLR model with different radiomics signatures extracted from dual-energy CT scans for the prediction of LN metastasis by a pretrained ResNet-18 model. Experiments of adding key clinical features were conducted to compare the effectiveness of using different approaches. The combined model integrated DL features and key clinical features yielded the highest AUC of 0.92 and accuracy of 86%.

The expression of various genes is an influential factor for patient prognosis and preoperative prediction of these prognostic factors can assist the diagnosis and treatment evaluation process. To predict the status of *HMGA2* and *C-MYC* gene expression of PDAC patients, Li *et al*[65] compared the use of radiomic features, DL features (extracted by pretrained CNN) and integration of both features in a CT-based model using an SVM classifier. Region of interest segmentation was conducted by two experienced radiologists individually, and the model was tested with different segmented images for improving the validity of the study. A model using DL features and all features achieved similar values in all evaluation metrics for both *C-MYC* and *HMGA2* tests, while DL features selected by Doctor B obtained outstanding average AUC scores (*C-MYC*: 0.90, *HMGA2*: 0.91) and accuracies (*C-MYC*: 95%, *HMGA2*: 88%) in the two gene tests.

The studies investigating the application of DLR for pancreatic cancer are summarized in [Table 4](#).

DLR FOR COLORECTAL CANCER

Colorectal cancer (CRC) is the third most common kind of cancer and the second leading cause of cancer-related fatalities worldwide[1]. It is crucial to carry out research on the diagnosis, treatment response prediction, and survival prediction of CRC, which can improve the prognosis of patients and significantly reduce the social and medical burden. In recent years, promising research results have emerged in the preoperative, intraoperative and postoperative stages of CRC using DLRs technology, covering the entire process of CRC diagnosis and treatment.

DLR is revolutionizing the treatment options for CRC. When making treatment decisions for CRC patients, identifying *KRAS* mutations, which may contribute to the continued proliferation of tumors, can help personalize treatment and care for CRC patients[66]. For preoperative prediction of *KRAS* mutations in patients with CRC, HCR and DLR were merged into a noninvasive model created by Wu *et al*[67]. The model, which mixed the handcrafted and DLR radiomics features, produced a C-index for the original cohort of 0.815 and the validation cohort of 0.832, which was higher than using HCR or DLR alone. For the individualized treatment decision-making in colorectal liver metastases (CRLM) management, the prediction of chemotherapeutic response is crucial. To predict the response to chemotherapy in CRLM, Wei *et al*[68] developed a ResNet10-based DLR model that used contrast-enhanced multidetector CT images as inputs. They also developed an HCR model for comparison. The

Table 4 Summary of studies using deep-learning-based radiomics for pancreatic cancer

Ref.	Imaging	Study design	Study aim	DL model	Dataset	Outcomes
Ziegelmayr <i>et al</i> [55], 2020	CT	Retrospective	Identification of PDAC and AIP	VGG19	86 patients (44 AIP patients and 42 PDAC patients)	Sensitivity: 89%; specificity: 83%; AUC: 0.90
Liao <i>et al</i> [56], 2022	CT	Retrospective	Identification of PDAC, non-cancerous pancreatic diseases and normal pancreas	CNN	3120 images (1872 for training, 624 for validation, 624 for testing)	Sensitivity: 89.9%; specificity: 91.3%; AUC: 0.960 when distinguishing between PDAC and control group
Tong <i>et al</i> [57], 2022	US	Retrospective	Identification of PDAC and CP	ResNet-50	558 patients	AUC: 0.967; sensitivity: 87.2%; specificity: 100%
Watson <i>et al</i> [58]	CT	Retrospective	Prediction of pathologic response of PDAC patients to NAC	LeNet	81 patients (65 for training and validation; 16 for testing)	AUC: 0.785; brier score: 0.174; sensitivity: 81.4%; specificity: 60.4% in test set of hybrid deep learning model
Muhammad <i>et al</i> [59], 2018	CT	Retrospective	Evaluation of survival hazard of PDAC patients	AlexNet	159 patients	C-index: 0.76; hazard ratio: 9.46 in test set
Zhang <i>et al</i> [60], 2020	CT	Retrospective	Evaluation of survival probability of PDAC patients	CNN	520 patients	IPA: 11.81%, C-index: 0.651 in testing cohort
Zhang <i>et al</i> [61], 2020	CT	Retrospective	Prediction of OS of PDAC patients; Evaluation of risk scores to distinguish patients with high or low risk	CNN	98 patients (68 in training cohort; 30 in testing cohort)	AUC: 0.81; hazard ratio: 1.86
Zhang <i>et al</i> [62], 2021	CT	Retrospective	Prediction of 2-yr OS of resectable PDAC patients	CNN	98 patients (68 in training cohort; 30 in testing cohort)	AUC: 0.84; specificity: 68%; sensitivity: 91%
Yao <i>et al</i> [63], 2021	CT	Retrospective	Prediction of survival risk and tumor resection margin of resectable PDAC patients	CNN	205 patients	C-index: 0.705 for survival prediction; balanced accuracy: 73.6%, sensitivity: 81.3%, specificity: 65.9% for resection margin prediction
Yao <i>et al</i> [63], 2021	CT	Retrospective	Prediction of survival risk and tumor resection margin of resectable PDAC patients	CNN	1209 patients	C-index: 0.667 for survival prediction; balanced accuracy: 67.1%; sensitivity: 59.8%; specificity: 74.3% for resection margin prediction
An <i>et al</i> [64], 2022	CT	Retrospective	Prediction of LNM status and OS in PDAC patients	ResNet-18	148 patients (88 in training cohort, 25 in validation cohort, 35 in testing cohort)	For combined model, AUC: 0.92; accuracy: 86%; sensitivity: 94%; specificity: 78% in testing cohort
Li <i>et al</i> [65], 2019	CT	Retrospective	Prediction of <i>HMG2A</i> and <i>C-MYC</i> gene expression status of PDAC patients; Prediction of survival time of patients	CNN	111 patients	Average AUC score: 0.90; accuracy: 95%; sensitivity: 92%; specificity: 98% in <i>C-MYC</i> test with deep features selected by Doctor B; average AUC score: 0.91; accuracy: 88%; sensitivity: 89%; specificity: 88% in <i>HMG2A</i> test with deep features selected by Doctor B

CT: Computed tomography; US: Ultrasound; PDAC: Pancreatic ductal adenocarcinoma; AIP: Autoimmune pancreatitis; CP: Chronic pancreatitis; NAC: Neoadjuvant chemotherapy; LNM: Lymph node metastasis; OS: Overall survival; AUC: Area under the receiver operating characteristic curve; C-index: Harrell's concordance index; CNN: Convolutional neural network; IPA: Index of prediction accuracy; DLR: Deep-learning-based radiomics.

DLR model achieved a higher AUC than the HCR model when predicting the response to chemotherapy in CRLM (training: 0.903 *vs* 0.745; validation: 0.820 *vs* 0.598). Microsatellite instability (MSI) function is a predictive biomarker for clinical outcomes and predicts responses to adjuvant 5-fluorouracil and immunotherapy in CRC. A DL model that was created using the MobileNetV2 architecture by Zhang *et al*[69] was adopted to predict the MSI status of CRC based on MR images. With AUC values of 0.868, the best model successfully identified 85.4% of the MSI status, indicating that the suggested model may aid in locating individuals who might benefit from chemotherapy or immunotherapy.

DLR in CRC also emphasizes the need to predict treatment response. For improving NCRT response prediction in locally advanced rectal cancer, Fu *et al*[70] compared the handcrafted and DL features extracted from pre-treatment diffusion-weighted MR images. The DLR approach produced a mean AUC of 0.73, while the HCR method yielded a mean AUC of 0.64, which demonstrated that DLR may achieve higher classification performance compared with HCRs. To predict the distant metastasis in

locally advanced rectal cancer patients receiving NCRT, Liu *et al*[71] exploited the use of a DLR model based on MR images. DLR achieved a C-index of 0.747 and AUC of 0.894 at 3 years. In order to define tumor morphological change for response evaluation in patients with metastatic CRC, Lu *et al*[72] offered a DLR study using CNN and recurrent neural network. They discovered that the DL network performed better than the size-based equivalent with C-index (0.649 *vs* 0.627), and was capable of predicting the early on-treatment response in metastatic CRC. The predictive performance could be improved by the integration of DL network with size-based methodology.

LN metastasis, which is a key prognostic factor for CRC, is among the other study topics of CRC. Ding *et al*[73] adopted a DLR nomogram based on faster region-based CNN (Faster R-CNN) to predict LN metastasis in patients with CRC. Patient age, Faster R-CNN-detected LN metastasis, and tumor differentiation were predictors in the Faster R-CNN nomogram for predicting LN metastasis, with AUCs in the training and validation sets of 0.862 and 0.920, respectively. Zhao *et al*[74] applied a DLR model related with genomics phenotypes for predicting LN metastasis in CRC and showed good performance with AUCs of 0.81, 0.77, and 0.73 in the training, testing and validation sets, respectively. Li *et al*[75] examined the performance of the three most popular classification techniques-DL, conventional machine learning, and deep transfer learning-to determine the most efficient way for automatic classification of CRC LN metastases. Deep transfer learning was the most successful, with an accuracy of 0.7583 and AUC of 0.7941. All of these studies have shown that DLR technology has good performance in the prediction and classification of LN metastasis.

The studies exploring the creation of DLR for CRC are summarized in [Table 5](#).

CHALLENGES AND RECOMMENDATIONS

In the past several years, with the development of DL technology, the research and application of DLR in tumor diagnosis, treatment and prognosis have been increasing. To perform a systematic evaluation of the status of DLRs for GI cancer, we conducted an extensive review on all original publications between January 1, 2015, and August 30, 2022. Even though several published articles have confirmed the exceptional performance of DLR, there are still many issues that algorithm designers and doctors must address. Below is a list of the challenges and recommendations for DLR in future research summarized by our team.

Prospective and multi-center studies

According to the most recent research, most studies on DLR were retrospective and single center. Retrospective studies may have sample selection bias and cannot truly reflect the distribution of clinical cases, which could jeopardize the precision of DLR models. As different centers have different machine parameters, scanning settings, and diagnostic rules, single-center studies limit the generalization of the DLR models. Prospective and multi-center studies can evaluate the reliability and accuracy of the DLR models, enhance their generalization, and bridge the gap between academic studies and clinical applications. Thus, carrying out prospective and multi-center studies is the key to accelerating the clinical application of DLRs models.

Development of user-friendly DL models

We found that many physicians do not really want to use DLR methods for related research because the models usually have complex structures, large parameters, poor interpretability, non-existence of gradients, overfitting, and other problems, which limit the promotion and use of DLR technology. Therefore, it is necessary to develop simple and user-friendly models and training schemes for non-professional users. Publication of more source codes and pre-training weights are ways to reduce the development and training difficulty of DL models. For overfitting problems, development of automatic data augmentation schemes and image synthesis schemes can increase the amount of training data. For the black box nature of DL models, attention maps and network dissection schemes can be integrated into the model to improve interpretability.

Establishment of accessible datasets

For DLR, the dataset is the new oil. DLR analysis requires a large amount of data to train and validate models; however, most studies are based on private datasets and do not use uniform construction standards, which will hinder the reproducibility of the studies and deployment of DLR models. Thus, a professional data development organization that combines multi-center data should be established. The organization should also standardize the development process of multiple kinds of datasets and make the datasets publicly accessible. Additionally, to reward data contributors, researchers who use these datasets could charge appropriate fees.

Efficient fusion of multiple features

DLR is a new technology in the field of AI for medical image analysis. Although its performance is satisfactory, it is not a panacea, especially in the case of extreme shortage of data. Numerous studies

Table 5 Summary of studies using deep-learning-based radiomics for colorectal cancer

Ref.	Imaging	Study design	Study aim	DL model	Dataset	Outcomes
Wu <i>et al</i> [67], 2020	CT	Retrospective	Predicting KRAS status in patients with CRC	CNN	Primary cohort: 279 patients; validation cohort: 119 patients	C-index of 0.815 for the primary cohort and 0.832 for the validation cohort
Wei <i>et al</i> [68], 2021	CT	Retrospective	Predicting the response to chemotherapy in CRLM	ResNet10	192 patients	AUC of DLR: 0.820; AUC of HCR: 0.598
Zhang <i>et al</i> [69], 2021	MRI	Retrospective	Predicting the MSI status of CRC	MobileNetV2	491 patients	Accuracy: 85.4%; AUC: 0.868
Fu <i>et al</i> [70], 2020	MRI	Retrospective	Predicting NCRT response in patients with LARC	VGG19	43 patients	AUC of DLR: 0.73; AUC of HCR: 0.64
Liu <i>et al</i> [71], 2021	MRI	Retrospective	Predicting the distant metastasis of LARC patients receiving NCRT	ResNet18	235 patients	C-index of 0.747 and AUC of 0.894 in the validation cohort
Lu <i>et al</i> [72], 2021	CT	Retrospective	Prediction of early on-treatment response in mCRC	CNN + RNN	1028 patients	C-index: 0.649
Ding <i>et al</i> [73], 2020	MRI	Retrospective	Prediction of metastatic LN in CRC	Faster RCNN	545 patients	AUC for training: 0.862; AUC for validation: 0.920
Zhao <i>et al</i> [74], 2022	CT	Retrospective	Prediction of metastatic LN in CRC	Autoencoder	423 patients	AUC for training: 0.81; AUC for validation: 0.73; AUC for testing: 0.77
Li <i>et al</i> [75], 2020	MRI	Retrospective	Classification of CRC LN Metastasis images	AlexNet	3364 samples (1646 positive; 1718 negative)	Accuracy: 75.83%; AUC: 0.7941

CRC: Colorectal cancer; CT: Computed tomography; MRI: Magnetic resonance imaging; CRLM: Colorectal liver metastases; MSI: Microsatellite instability; NCRT: Neoadjuvant chemoradiotherapy; LARC: Locally advanced rectal cancer; mCRC: Metastatic colorectal cancer; LN: Lymph node; CNN: Convolutional neural network; RNN: Recurrent neural network; AUC: Area under the receiver operating characteristic curve; C-index: Harrell's concordance index; DLR: Deep-learning-based radiomics; HCR: Handcrafted radiomics; Faster R-CNN: Faster region-based convolutional neural network.

have demonstrated that combining HCR and DLR, can result in better performance. Thus, we suggest integrating other clinical features, genomics, handcrafted features, and DL features to build an optimal solution. Moreover, a suitable feature dimensionality reduction scheme should also be adopted to reduce the redundancy of the integrated features. In addition to imaging features, features extracted from clinical data sources, such as gene expression, clinical characteristics, and blood biomarkers, can also be combined to enhance radiomic features.

CONCLUSION

Globally, GI cancers account for a large portion of cancer-related fatalities. For the diagnosis and treatment of GI cancer, DLR can offer a simpler, quicker and more reliable approach. This article is the first comprehensive review on DLR in the GI tract. The status, difficulties, and suggestions discussed in this review can help engineers create optimal radiomics products to support clinical decision-making and offer guidance for diagnosis and treatment of other tumors. Despite the success of DLR in GI cancer, prospective and multi-center studies are still needed. Development of user-friendly DL models, the creation of large public databases, and the fusion of multiple features are also necessary to encourage the clinical application of radiomics.

FOOTNOTES

Author contributions: Wong PK, Chan IN, Yan HM, Gao S, Wong CH, and Yan T collected the literature and wrote the initial manuscript, conceptualized the table and figures, and contributed equally to this work; Yao L, Hu Y, Wang ZR, and Yu HH conceptualized the structure of the text and critically revised the manuscript for important intellectual content; all authors read and approved the final version of the manuscript.

Supported by the Guangdong Basic and Applied Basic Research Fund, Shenzhen Joint Fund (Guangdong-Shenzhen Joint Fund) Guangdong-Hong Kong-Macau Research Team Project, No. 2021B1515130003; Science and Technology

Development Fund of Macau, No. 0026/2022/A; and Project of Xiangyang Science and Technology on Medical and Health Field, No. 2022YL05A.

Conflict-of-interest statement: All authors declare no conflict of interest.

Open-Access: This article is an open-access article that was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution NonCommercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: <https://creativecommons.org/licenses/by-nc/4.0/>

Country/Territory of origin: China

ORCID number: Pak Kin Wong 0000-0002-7623-6904; In Neng Chan 0000-0002-7015-4510; Hao-Ming Yan 0000-0002-2223-4671; Shan Gao 0000-0002-8232-4134; Chi Hong Wong 0000-0003-2183-4525; Tao Yan 0000-0002-8929-015X; Liang Yao 0000-0003-3377-1509; Ying Hu 0000-0002-3807-3649; Zhong-Ren Wang 0000-0002-3488-7119; Hon Ho Yu 0000-0002-9580-345X.

S-Editor: Chen YL

L-Editor: A

P-Editor: Chen YL

REFERENCES

- 1 **Sung H**, Ferlay J, Siegel RL, Laversanne M, Soerjomataram I, Jemal A, Bray F. Global Cancer Statistics 2020: GLOBOCAN Estimates of Incidence and Mortality Worldwide for 36 Cancers in 185 Countries. *CA Cancer J Clin* 2021; **71**: 209-249 [PMID: 33538338 DOI: 10.3322/caac.21660]
- 2 **World Health Organization.** Cancer Tomorrow. [cited 4 August 2022]. Available from: <https://gco.iarc.fr/tomorrow/en>. [DOI: 10.32755/sjcriminal.2022.01]
- 3 **Kinkel K**, Lu Y, Both M, Warren RS, Thoeni RF. Detection of hepatic metastases from cancers of the gastrointestinal tract by using noninvasive imaging methods (US, CT, MR imaging, PET): a meta-analysis. *Radiology* 2002; **224**: 748-756 [PMID: 12202709 DOI: 10.1148/radiol.2243011362]
- 4 **Dimitrakopoulou-Strauss A**, Ronellenfitsch U, Cheng C, Pan L, Sachpekidis C, Hohenberger P, Henzler T. Imaging therapy response of gastrointestinal stromal tumors (GIST) with FDG PET, CT and MRI: a systematic review. *Clin Transl Imaging* 2017; **5**: 183-197 [PMID: 29104864 DOI: 10.1007/s40336-017-0229-8]
- 5 **Davnall F**, Yip CS, Ljungqvist G, Selmi M, Ng F, Sanghera B, Ganeshan B, Miles KA, Cook GJ, Goh V. Assessment of tumor heterogeneity: an emerging imaging tool for clinical practice? *Insights Imaging* 2012; **3**: 573-589 [PMID: 23093486 DOI: 10.1007/s13244-012-0196-6]
- 6 **Lambin P**, Rios-Velazquez E, Leijenaar R, Carvalho S, van Stiphout RG, Granton P, Zegers CM, Gillies R, Boellard R, Dekker A, Aerts HJ. Radiomics: extracting more information from medical images using advanced feature analysis. *Eur J Cancer* 2012; **48**: 441-446 [PMID: 22257792 DOI: 10.1016/j.ejca.2011.11.036]
- 7 **Du KP**, Huang WP, Liu SY, Chen YJ, Li LM, Liu XN, Han YJ, Zhou Y, Liu CC, Gao JB. Application of computed tomography-based radiomics in differential diagnosis of adenocarcinoma and squamous cell carcinoma at the esophagogastric junction. *World J Gastroenterol* 2022; **28**: 4363-4375 [PMID: 36159013 DOI: 10.3748/wjg.v28.i31.4363]
- 8 **Zhang YC**, Li M, Jin YM, Xu JX, Huang CC, Song B. Radiomics for differentiating tumor deposits from lymph node metastasis in rectal cancer. *World J Gastroenterol* 2022; **28**: 3960-3970 [PMID: 36157536 DOI: 10.3748/wjg.v28.i29.3960]
- 9 **Hou M**, Sun JH. Emerging applications of radiomics in rectal cancer: State of the art and future perspectives. *World J Gastroenterol* 2021; **27**: 3802-3814 [PMID: 34321845 DOI: 10.3748/wjg.v27.i25.3802]
- 10 **Cannella R**, La Grutta L, Midiri M, Bartolotta TV. New advances in radiomics of gastrointestinal stromal tumors. *World J Gastroenterol* 2020; **26**: 4729-4738 [PMID: 32921953 DOI: 10.3748/wjg.v26.i32.4729]
- 11 **Stanzione A**, Verde F, Romeo V, Boccadifuoco F, Mainenti PP, Maurea S. Radiomics and machine learning applications in rectal cancer: Current update and future perspectives. *World J Gastroenterol* 2021; **27**: 5306-5321 [PMID: 34539134 DOI: 10.3748/wjg.v27.i32.5306]
- 12 **Aerts HJ**, Velazquez ER, Leijenaar RT, Parmar C, Grossmann P, Carvalho S, Bussink J, Monshouwer R, Haibe-Kains B, Rietveld D, Hoebers F, Rietbergen MM, Leemans CR, Dekker A, Quackenbush J, Gillies RJ, Lambin P. Decoding tumour phenotype by noninvasive imaging using a quantitative radiomics approach. *Nat Commun* 2014; **5**: 4006 [PMID: 24892406 DOI: 10.1038/ncomms5006]
- 13 **Afshar P**, Mohammadi A, Plataniotis KN, Oikonomou A, Benali H. From handcrafted to deep-learning-based cancer radiomics: Challenges and opportunities. *IEEE Signal Process Mag* 2019; **36**: 132-160 [DOI: 10.1109/msp.2019.2900993]
- 14 **LeCun Y**, Bengio Y, Hinton G. Deep learning. *Nature* 2015; **521**: 436-444 [PMID: 26017442 DOI: 10.1038/nature14539]
- 15 **Shin HC**, Roth HR, Gao M, Lu L, Xu Z, Nogues I, Yao J, Mollura D, Summers RM. Deep Convolutional Neural Networks for Computer-Aided Detection: CNN Architectures, Dataset Characteristics and Transfer Learning. *IEEE Trans Med Imaging* 2016; **35**: 1285-1298 [PMID: 26886976 DOI: 10.1109/TMI.2016.2528162]
- 16 **Chan HP**, Samala RK, Hadjiiski LM, Zhou C. Deep Learning in Medical Image Analysis. *Adv Exp Med Biol* 2020; **1213**: 3-21 [PMID: 32030660 DOI: 10.1007/978-3-030-33128-3_1]

- 17 **Yan T**, Wong PK, Qin YY. Deep learning for diagnosis of precancerous lesions in upper gastrointestinal endoscopy: A review. *World J Gastroenterol* 2021; **27**: 2531-2544 [PMID: 34092974 DOI: 10.3748/wjg.v27.i20.2531]
- 18 **Tamada D**, Kromrey ML, Ichikawa S, Onishi H, Motosugi U. Motion Artifact Reduction Using a Convolutional Neural Network for Dynamic Contrast Enhanced MR Imaging of the Liver. *Magn Reson Med Sci* 2020; **19**: 64-76 [PMID: 31061259 DOI: 10.2463/mrms.mp.2018-0156]
- 19 **Liu Z**, Jiang Z, Meng L, Yang J, Liu Y, Zhang Y, Peng H, Li J, Xiao G, Zhang Z, Zhou R. Handcrafted and Deep Learning-Based Radiomic Models Can Distinguish GBM from Brain Metastasis. *J Oncol* 2021; **2021**: 5518717 [PMID: 34188680 DOI: 10.1155/2021/5518717]
- 20 **Hatt M**, Parmar C, Qi J, El Naqa I (2019). Machine (deep) learning methods for image processing and radiomics. *IEEE Trans Radiat Plasma Med Sci* **3**: 104-108 [DOI: 10.1109/trpms.2019.2899538]
- 21 **Zhang X**, Zhang Y, Zhang G, Qiu X, Tan W, Yin X, Liao L. Deep Learning With Radiomics for Disease Diagnosis and Treatment: Challenges and Potential. *Front Oncol* 2022; **12**: 773840 [PMID: 35251962 DOI: 10.3389/fonc.2022.773840]
- 22 **Avanzo M**, Wei L, Stancanello J, Vallières M, Rao A, Morin O, Mattonen SA, El Naqa I. Machine and deep learning methods for radiomics. *Med Phys* 2020; **47**: e185-e202 [PMID: 32418336 DOI: 10.1002/mp.13678]
- 23 **Takeuchi M**, Seto T, Hashimoto M, Ichihara N, Morimoto Y, Kawakubo H, Suzuki T, Jinzaki M, Kitagawa Y, Miyata H, Sakakibara Y. Performance of a deep learning-based identification system for esophageal cancer from CT images. *Esophagus* 2021; **18**: 612-620 [PMID: 33635412 DOI: 10.1007/s10388-021-00826-0]
- 24 **Hu Y**, Xie C, Yang H, Ho JWK, Wen J, Han L, Lam KO, Wong IYH, Law SYK, Chiu KWH, Vardhanabhuti V, Fu J. Computed tomography-based deep-learning prediction of neoadjuvant chemoradiotherapy treatment response in esophageal squamous cell carcinoma. *Radiother Oncol* 2021; **154**: 6-13 [PMID: 32941954 DOI: 10.1016/j.radonc.2020.09.014]
- 25 **Ypsilantis PP**, Siddique M, Sohn HM, Davies A, Cook G, Goh V, Montana G. Predicting Response to Neoadjuvant Chemotherapy with PET Imaging Using Convolutional Neural Networks. *PLoS One* 2015; **10**: e0137036 [PMID: 26355298 DOI: 10.1371/journal.pone.0137036]
- 26 **Amyar A**, Ruan S, Gardin I, Chatelain C, Decazes P, Modzelewski R. 3-D RPET-NET: Development of a 3-D PET Imaging Convolutional Neural Network for Radiomics Analysis and Outcome Prediction. *IEEE Trans Radiat Plasma Med Sci* 2019; **3**: 225-231 [DOI: 10.1109/trpms.2019.2896399]
- 27 **Li X**, Gao H, Zhu J, Huang Y, Zhu Y, Huang W, Li Z, Sun K, Liu Z, Tian J, Li B. 3D Deep Learning Model for the Pretreatment Evaluation of Treatment Response in Esophageal Carcinoma: A Prospective Study (ChiCTR2000039279). *Int J Radiat Oncol Biol Phys* 2021; **111**: 926-935 [PMID: 34229050 DOI: 10.1016/j.ijrobp.2021.06.033]
- 28 **Wang J**, Zeng J, Li H, Yu X. A Deep Learning Radiomics Analysis for Survival Prediction in Esophageal Cancer. *J Healthc Eng* 2022; **2022**: 4034404 [PMID: 35368956 DOI: 10.1155/2022/4034404]
- 29 **Yang CK**, Yeh JC, Yu WH, Chien LI, Lin KH, Huang WS, Hsu PK. Deep Convolutional Neural Network-Based Positron Emission Tomography Analysis Predicts Esophageal Cancer Outcome. *J Clin Med* 2019; **8** [PMID: 31200519 DOI: 10.3390/jcm8060844]
- 30 **Gong J**, Zhang W, Huang W, Liao Y, Yin Y, Shi M, Qin W, Zhao L. CT-based radiomics nomogram may predict local recurrence-free survival in esophageal cancer patients receiving definitive chemoradiation or radiotherapy: A multicenter study. *Radiother Oncol* 2022; **174**: 8-15 [PMID: 35750106 DOI: 10.1016/j.radonc.2022.06.010]
- 31 **Wu L**, Yang X, Cao W, Zhao K, Li W, Ye W, Chen X, Zhou Z, Liu Z, Liang C. Multiple Level CT Radiomics Features Preoperatively Predict Lymph Node Metastasis in Esophageal Cancer: A Multicentre Retrospective Study. *Front Oncol* 2019; **9**: 1548 [PMID: 32039021 DOI: 10.3389/fonc.2019.01548]
- 32 **Cui Y**, Zhang J, Li Z, Wei K, Lei Y, Ren J, Wu L, Shi Z, Meng X, Yang X, Gao X. A CT-based deep learning radiomics nomogram for predicting the response to neoadjuvant chemotherapy in patients with locally advanced gastric cancer: A multicenter cohort study. *EClinicalMedicine* 2022; **46**: 101348 [PMID: 35340629 DOI: 10.1016/j.eclinm.2022.101348]
- 33 **Li C**, Qin Y, Zhang WH, Jiang H, Song B, Bashir MR, Xu H, Duan T, Fang M, Zhong L, Meng L, Dong D, Hu Z, Tian J, Hu JK. Deep learning-based AI model for signet-ring cell carcinoma diagnosis and chemotherapy response prediction in gastric cancer. *Med Phys* 2022; **49**: 1535-1546 [PMID: 35032039 DOI: 10.1002/mp.15437]
- 34 **Tan JW**, Wang L, Chen Y, Xi W, Ji J, Xu X, Zou LK, Feng JX, Zhang J, Zhang H. Predicting Chemotherapeutic Response for Far-advanced Gastric Cancer by Radiomics with Deep Learning Semi-automatic Segmentation. *J Cancer* 2020; **11**: 7224-7236 [PMID: 33193886 DOI: 10.7150/jca.46704]
- 35 **Hao D**, Li Q, Feng QX, Qi L, Liu XS, Arefan D, Zhang YD, Wu S. Identifying Prognostic Markers From Clinical, Radiomics, and Deep Learning Imaging Features for Gastric Cancer Survival Prediction. *Front Oncol* 2021; **11**: 725889 [PMID: 35186707 DOI: 10.3389/fonc.2021.725889]
- 36 **Zhang L**, Dong D, Zhong L, Li C, Hu C, Yang X, Liu Z, Wang R, Zhou J, Tian J. Multi-Focus Network to Decode Imaging Phenotype for Overall Survival Prediction of Gastric Cancer Patients. *IEEE J Biomed Health Inform* 2021; **25**: 3933-3942 [PMID: 34101609 DOI: 10.1109/JBHI.2021.3087634]
- 37 **Zhang W**, Fang M, Dong D, Wang X, Ke X, Zhang L, Hu C, Guo L, Guan X, Zhou J, Shan X, Tian J. Development and validation of a CT-based radiomic nomogram for preoperative prediction of early recurrence in advanced gastric cancer. *Radiother Oncol* 2020; **145**: 13-20 [PMID: 31869677 DOI: 10.1016/j.radonc.2019.11.023]
- 38 **Guan X**, Lu N, Zhang J. Computed Tomography-Based Deep Learning Nomogram Can Accurately Predict Lymph Node Metastasis in Gastric Cancer. *Dig Dis Sci* 2022 [PMID: 35909203 DOI: 10.1007/s10620-022-07640-3]
- 39 **Dong D**, Fang MJ, Tang L, Shan XH, Gao JB, Giganti F, Wang RP, Chen X, Wang XX, Palumbo D, Fu J, Li WC, Li J, Zhong LZ, De Cobelli F, Ji JF, Liu ZY, Tian J. Deep learning radiomic nomogram can predict the number of lymph node metastasis in locally advanced gastric cancer: an international multicenter study. *Ann Oncol* 2020; **31**: 912-920 [PMID: 32304748 DOI: 10.1016/j.annonc.2020.04.003]
- 40 **Li J**, Dong D, Fang M, Wang R, Tian J, Li H, Gao J. Dual-energy CT-based deep learning radiomics can improve lymph node metastasis risk prediction for gastric cancer. *Eur Radiol* 2020; **30**: 2324-2333 [PMID: 31953668 DOI: 10.1007/s00330-019-06621-x]
- 41 **Jin C**, Jiang Y, Yu H, Wang W, Li B, Chen C, Yuan Q, Hu Y, Xu Y, Zhou Z, Li G, Li R. Deep learning analysis of the

- primary tumour and the prediction of lymph node metastases in gastric cancer. *Br J Surg* 2021; **108**: 542-549 [PMID: 34043780 DOI: 10.1002/bjs.11928]
- 42 Sun RJ, Fang MJ, Tang L, Li XT, Lu QY, Dong D, Tian J, Sun YS. CT-based deep learning radiomics analysis for evaluation of serosa invasion in advanced gastric cancer. *Eur J Radiol* 2020; **132**: 109277 [PMID: 32980726 DOI: 10.1016/j.ejrad.2020.109277]
- 43 Li Q, Feng QX, Qi L, Liu C, Zhang J, Yang G, Zhang YD, Liu XS. Prognostic aspects of lymphovascular invasion in localized gastric cancer: new insights into the radiomics and deep transfer learning from contrast-enhanced CT imaging. *Abdom Radiol (NY)* 2022; **47**: 496-507 [PMID: 34766197 DOI: 10.1007/s00261-021-03309-z]
- 44 Ding Y, Ruan S, Wang Y, Shao J, Sun R, Tian W, Xiang N, Ge W, Zhang X, Su K, Xia J, Huang Q, Liu W, Sun Q, Dong H, Farias MCQ, Guo T, Krylov AS, Liang W, Xiao W, Bai X, Liang T. Novel deep learning radiomics model for preoperative evaluation of hepatocellular carcinoma differentiation based on computed tomography data. *Clin Transl Med* 2021; **11**: e570 [PMID: 34841694 DOI: 10.1002/ctm2.570]
- 45 Peng J, Kang S, Ning Z, Deng H, Shen J, Xu Y, Zhang J, Zhao W, Li X, Gong W, Huang J, Liu L. Residual convolutional neural network for predicting response of transarterial chemoembolization in hepatocellular carcinoma from CT imaging. *Eur Radiol* 2020; **30**: 413-424 [PMID: 31332558 DOI: 10.1007/s00330-019-06318-1]
- 46 Peng J, Huang J, Huang G, Zhang J. Predicting the Initial Treatment Response to Transarterial Chemoembolization in Intermediate-Stage Hepatocellular Carcinoma by the Integration of Radiomics and Deep Learning. *Front Oncol* 2021; **11**: 730282 [PMID: 34745952 DOI: 10.3389/fonc.2021.730282]
- 47 Wei L, Owen D, Rosen B, Guo X, Cuneo K, Lawrence TS, Ten Haken R, El Naqa I. A deep survival interpretable radiomics model of hepatocellular carcinoma patients. *Phys Med* 2021; **82**: 295-305 [PMID: 33714190 DOI: 10.1016/j.ejmp.2021.02.013]
- 48 Liu F, Liu D, Wang K, Xie X, Su L, Kuang M, Huang G, Peng B, Wang Y, Lin M, Tian J. Deep Learning Radiomics Based on Contrast-Enhanced Ultrasound Might Optimize Curative Treatments for Very-Early or Early-Stage Hepatocellular Carcinoma Patients. *Liver Cancer* 2020; **9**: 397-413 [PMID: 32999867 DOI: 10.1159/000505694]
- 49 Wang W, Chen Q, Iwamoto Y, Han X, Zhang Q, Hu H, Lin L, Chen YW. Deep Learning-Based Radiomics Models for Early Recurrence Prediction of Hepatocellular Carcinoma with Multi-phase CT Images and Clinical Data. *Annu Int Conf IEEE Eng Med Biol Soc* 2019; **2019**: 4881-4884 [PMID: 31946954 DOI: 10.1109/EMBC.2019.8856356]
- 50 Wang W, Chen Q, Iwamoto Y, Aonpong P, Lin L, Hu H, Zhang Q, Chen YW. Deep fusion models of multi-phase CT and selected clinical data for preoperative prediction of early recurrence in hepatocellular carcinoma. *IEEE Access* 2020; **8**: 139212-139220 [DOI: 10.1109/access.2020.3011145]
- 51 He T, Fong JN, Moore LW, Ezeana CF, Victor D, Divatia M, Vasquez M, Ghobrial RM, Wong STC. An imageomics and multi-network based deep learning model for risk assessment of liver transplantation for hepatocellular cancer. *Comput Med Imaging Graph* 2021; **89**: 101894 [PMID: 33725579 DOI: 10.1016/j.compmedimag.2021.101894]
- 52 Jiang YQ, Cao SE, Cao S, Chen JN, Wang GY, Shi WQ, Deng YN, Cheng N, Ma K, Zeng KN, Yan XJ, Yang HZ, Huan WJ, Tang WM, Zheng Y, Shao CK, Wang J, Yang Y, Chen GH. Preoperative identification of microvascular invasion in hepatocellular carcinoma by XGBoost and deep learning. *J Cancer Res Clin Oncol* 2021; **147**: 821-833 [PMID: 32852634 DOI: 10.1007/s00432-020-03366-9]
- 53 Wang L, Wu M, Li R, Xu X, Zhu C, Feng X. MVI-Mind: A Novel Deep-Learning Strategy Using Computed Tomography (CT)-Based Radiomics for End-to-End High Efficiency Prediction of Microvascular Invasion in Hepatocellular Carcinoma. *Cancers (Basel)* 2022; **14** [PMID: 35740620 DOI: 10.3390/cancers14122956]
- 54 Fu S, Lai H, Huang M, Li Q, Liu Y, Zhang J, Huang J, Chen X, Duan C, Li X, Wang T, He X, Yan J, Lu L. Multi-task deep learning network to predict future macrovascular invasion in hepatocellular carcinoma. *EClinicalMedicine* 2021; **42**: 101201 [PMID: 34917908 DOI: 10.1016/j.eclinm.2021.101201]
- 55 Ziegelmayer S, Kaissis G, Harder F, Jungmann F, Müller T, Makowski M, Braren R. Deep Convolutional Neural Network-Assisted Feature Extraction for Diagnostic Discrimination and Feature Visualization in Pancreatic Ductal Adenocarcinoma (PDAC) versus Autoimmune Pancreatitis (AIP). *J Clin Med* 2020; **9** [PMID: 33322559 DOI: 10.3390/jcm9124013]
- 56 Liao WC, Chang D, Chen PT, Wang P, Liu KL, Wu MS, Wang W. Ep1153: distinguishing pancreatic cancer from non-cancerous pancreatic diseases and normal pancreas with deep learning-based segmentation and radiomics-based classification. *Gastroenterology* 2022; **162**: S-1071 [DOI: 10.1016/s0016-5085(22)62563-0]
- 57 Tong T, Gu J, Xu D, Song L, Zhao Q, Cheng F, Yuan Z, Tian S, Yang X, Tian J, Wang K, Jiang T. Deep learning radiomics based on contrast-enhanced ultrasound images for assisted diagnosis of pancreatic ductal adenocarcinoma and chronic pancreatitis. *BMC Med* 2022; **20**: 74 [PMID: 35232446 DOI: 10.1186/s12916-022-02258-8]
- 58 Watson MD, Baimas-George MR, Murphy KJ, Pickens RC, Iannitti DA, Martinie JB, Baker EH, Vrochides D, Ocuin LM. Pure and Hybrid Deep Learning Models can Predict Pathologic Tumor Response to Neoadjuvant Therapy in Pancreatic Adenocarcinoma: A Pilot Study. *Am Surg* 2021; **87**: 1901-1909 [PMID: 33381979 DOI: 10.1177/0003134820982557]
- 59 Muhammad H, Häggström I, Klimstra DS, Fuchs TJ. Survival modeling of pancreatic cancer with radiology using convolutional neural networks. In: Simulation, image processing, and ultrasound systems for assisted diagnosis and navigation. Cham: Springer, 2018: 187-192 [DOI: 10.1007/978-3-030-01045-4_23]
- 60 Zhang Y, Lobo-Mueller EM, Karanicolas P, Gallinger S, Haider MA, Khalvati F. CNN-based survival model for pancreatic ductal adenocarcinoma in medical imaging. *BMC Med Imaging* 2020; **20**: 11 [PMID: 32013871 DOI: 10.1186/s12880-020-0418-1]
- 61 Zhang Y, Lobo-Mueller EM, Karanicolas P, Gallinger S, Haider MA, Khalvati F. Prognostic Value of Transfer Learning Based Features in Resectable Pancreatic Ductal Adenocarcinoma. *Front Artif Intell* 2020; **3**: 550890 [PMID: 33733206 DOI: 10.3389/frai.2020.550890]
- 62 Zhang Y, Lobo-Mueller EM, Karanicolas P, Gallinger S, Haider MA, Khalvati F. Improving prognostic performance in resectable pancreatic ductal adenocarcinoma using radiomics and deep learning features fusion in CT images. *Sci Rep* 2021; **11**: 1378 [PMID: 33446870 DOI: 10.1038/s41598-021-80998-y]
- 63 Yao J, Shi Y, Cao K, Lu L, Lu J, Song Q, Jin G, Xiao J, Hou Y, Zhang L. DeepPrognosis: Preoperative prediction of pancreatic cancer survival and surgical margin *via* comprehensive understanding of dynamic contrast-enhanced CT imaging

- and tumor-vascular contact parsing. *Med Image Anal* 2021; **73**: 102150 [PMID: 34303891 DOI: 10.1016/j.media.2021.102150]
- 64 **An C**, Li D, Li S, Li W, Tong T, Liu L, Jiang D, Jiang L, Ruan G, Hai N, Fu Y, Wang K, Zhuo S, Tian J. Deep learning radiomics of dual-energy computed tomography for predicting lymph node metastases of pancreatic ductal adenocarcinoma. *Eur J Nucl Med Mol Imaging* 2022; **49**: 1187-1199 [PMID: 34651229 DOI: 10.1007/s00259-021-05573-z]
- 65 **Li K**, Xiao J, Yang J, Li M, Xiong X, Nian Y, Qiao L, Wang H, Eresen A, Zhang Z, Hu X, Wang J, Chen W. Association of radiomic imaging features and gene expression profile as prognostic factors in pancreatic ductal adenocarcinoma. *Am J Transl Res* 2019; **11**: 4491-4499 [PMID: 31396352]
- 66 **Meng M**, Zhong K, Jiang T, Liu Z, Kwan HY, Su T. The current understanding on the impact of KRAS on colorectal cancer. *Biomed Pharmacother* 2021; **140**: 111717 [PMID: 34044280 DOI: 10.1016/j.biopha.2021.111717]
- 67 **Wu X**, Li Y, Chen X, Huang Y, He L, Zhao K, Huang X, Zhang W, Dong M, Huang J, Xia T, Liang C, Liu Z. Deep Learning Features Improve the Performance of a Radiomics Signature for Predicting KRAS Status in Patients with Colorectal Cancer. *Acad Radiol* 2020; **27**: e254-e262 [PMID: 31982342 DOI: 10.1016/j.acra.2019.12.007]
- 68 **Wei J**, Cheng J, Gu D, Chai F, Hong N, Wang Y, Tian J. Deep learning-based radiomics predicts response to chemotherapy in colorectal liver metastases. *Med Phys* 2021; **48**: 513-522 [PMID: 33119899 DOI: 10.1002/mp.14563]
- 69 **Zhang W**, Yin H, Huang Z, Zhao J, Zheng H, He D, Li M, Tan W, Tian S, Song B. Development and validation of MRI-based deep learning models for prediction of microsatellite instability in rectal cancer. *Cancer Med* 2021; **10**: 4164-4173 [PMID: 33963688 DOI: 10.1002/cam4.3957]
- 70 **Fu J**, Zhong X, Li N, Van Dams R, Lewis J, Sung K, Raldow AC, Jin J, Qi XS. Deep learning-based radiomic features for improving neoadjuvant chemoradiation response prediction in locally advanced rectal cancer. *Phys Med Biol* 2020; **65**: 075001 [PMID: 32092710 DOI: 10.1088/1361-6560/ab7970]
- 71 **Liu X**, Zhang D, Liu Z, Li Z, Xie P, Sun K, Wei W, Dai W, Tang Z, Ding Y, Cai G, Tong T, Meng X, Tian J. Deep learning radiomics-based prediction of distant metastasis in patients with locally advanced rectal cancer after neoadjuvant chemoradiotherapy: A multicentre study. *EBioMedicine* 2021; **69**: 103442 [PMID: 34157487 DOI: 10.1016/j.ebiom.2021.103442]
- 72 **Lu L**, Dercle L, Zhao B, Schwartz LH. Deep learning for the prediction of early on-treatment response in metastatic colorectal cancer from serial medical imaging. *Nat Commun* 2021; **12**: 6654 [PMID: 34789774 DOI: 10.1038/s41467-021-26990-6]
- 73 **Ding L**, Liu G, Zhang X, Liu S, Li S, Zhang Z, Guo Y, Lu Y. A deep learning nomogram kit for predicting metastatic lymph nodes in rectal cancer. *Cancer Med* 2020; **9**: 8809-8820 [PMID: 32997900 DOI: 10.1002/cam4.3490]
- 74 **Zhao J**, Wang H, Zhang Y, Wang R, Liu Q, Li J, Li X, Huang H, Zhang J, Zeng Z, Yi Z, Zeng F. Deep learning radiomics model related with genomics phenotypes for lymph node metastasis prediction in colorectal cancer. *Radiother Oncol* 2022; **167**: 195-202 [PMID: 34968471 DOI: 10.1016/j.radonc.2021.12.031]
- 75 **Li J**, Wang P, Zhou Y, Liang H, Luan K. Different Machine Learning and Deep Learning Methods for the Classification of Colorectal Cancer Lymph Node Metastasis Images. *Front Bioeng Biotechnol* 2020; **8**: 620257 [PMID: 33520971 DOI: 10.3389/fbioe.2020.620257]



Published by **Baishideng Publishing Group Inc**
7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA
Telephone: +1-925-3991568
E-mail: bpgoffice@wjgnet.com
Help Desk: <https://www.f6publishing.com/helpdesk>
<https://www.wjgnet.com>

