

Accuracy of convolutional neural network-based artificial intelligence in diagnosis of gastrointestinal lesions based on endoscopic images: A systematic review and meta-analysis



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ABSTRACT

Background and study aims Recently, a growing body of evidence has been amassed on evaluation of artificial intelligence (AI) known as deep learning in computer-aided diagnosis of gastrointestinal lesions by means of convolutional neural networks (CNN). We conducted this meta-analysis to study pooled rates of performance for CNN-based AI in diagnosis of gastrointestinal neoplasia from endoscopic images.

Methods Multiple databases were searched (from inception to November 2019) and studies that reported on the performance of AI by means of CNN in the diagnosis of gastrointestinal tumors were selected. A random effects model was used and pooled accuracy, sensitivity, specificity, positive predictive value (PPV) and negative predictive value (NPV) were calculated. Pooled rates were categorized based on the gastrointestinal location of lesion (esophagus, stomach and colorectum).

Results Nineteen studies were included in our final analysis. The pooled accuracy of CNN in esophageal neoplasia was 87.2% (76–93.6) and NPV was 92.1% (85.9–95.7); the accuracy in lesions of stomach was 85.8% (79.8–90.3) and NPV was 92.1% (85.9–95.7); and in colorectal neoplasia the accuracy was 89.9% (82–94.7) and NPV was 94.3% (86.4–97.7).

Conclusions Based on our meta-analysis, CNN-based AI achieved high accuracy in diagnosis of lesions in esophagus, stomach, and colorectum.

Introduction

Early detection of gastrointestinal neoplasia by endoscopy is a widely adopted strategy to prevent cancer-related morbidity

and/ or mortality. The disease prognosis greatly depends on the stage of cancer at diagnosis. Gastrointestinal neoplastic conditions are frequently detected by direct endoscopic visualization by a trained endoscopist and endoscopists use their

knowledge, gathered from experience of endoscopic appearance, to detect these lesions.

To maximize detection and/or differentiation of a lesion, a clean mucosal surface and a meticulous mechanical exploration are paramount. Apart from detecting a lesion, predicting its potential to be carcinogenic is difficult. In addition, both lesion detection and its assessment are subject to substantial operator dependence. To improve detection of lesion by human eye, various optical enhancements of the endoscope have been made. High-definition white light endoscopy with or without chromo-endoscopy, narrow-band imaging (NBI) with or without magnification, confocal laser endomicroscopy, and endocytoscopic imaging system are some of the examples.

Recently, a growing body of evidence has been amassed on use of artificial intelligence (AI) known as deep learning in computer-aided diagnosis (CAD) of health-related conditions based on medical imaging [1]. A convolutional neural network (CNN) is a type of deep learning method that enables machines to analyze various training images and extract specific clinical features using a back-propagation algorithm. CNN data-driven systems are trained on datasets containing large numbers of images with their corresponding labels. CNN can be seen as a system that first extracts relevant features from the input images and it subsequently uses those learned features to classify a given image. The network uses convolutions of the input image to extract the most relevant information that helps to classify the image into different entities. Based on the accumulated data features, machine algorithms can diagnose newly acquired clinical images prospectively [2–4].

CNN-based CAD has been reported as being highly beneficial in the field of endoscopy, including EGD, colonoscopy and capsule endoscopy. [2, 5, 6] CNN has transformed the field of computer vision and has been shown to work in real-time with raw, unprocessed frames from the video sequence. [2] In this systematic review and meta-analysis, we aim to quantitatively appraise the current reported data on the diagnostic performance of CNN based computer aided diagnosis of gastrointestinal neoplasia.

Methods

Search strategy

The literature was searched by a medical librarian for the concepts of AI with endoscopy for gastrointestinal lesions. The search strategies were created using a combination of keywords and standardized index terms. Searches were run in November 2019 in ClinicalTrials.gov, Ovid EBM Reviews, Ovid Embase (1974+), Ovid Medline (1946+ including epub ahead of print, in-process & other non-indexed citations), Scopus (1970+) and Web of Science (1975+). Results were limited to English language. All results were exported to Endnote X9 (Clarivate Analytics) where obvious duplicates were removed leaving 4245 citations. Search strategy is listed in **Appendix 1**. The MOOSE checklist was followed and is listed in **Appendix 2**. Reference lists of evaluated studies were examined to identify other potential studies of interest.

Study selection

In this meta-analysis, we included studies that developed or validated a deep CNN learning model for diagnosis of neoplasia of the gastrointestinal tract (esophagus, stomach, and colorectum) using either one or a combination of white-light endoscopy (WLE), narrow-band imaging (NBI) endoscopy (magnifying and/ or non-magnifying), and chromoendoscopy. Study selection was restricted to only those that used CNN-based deep machine learning models. Studies were included irrespective of inpatient/outpatient setting, study sample-size, follow-up time, abstract/ manuscript status, and geography as long as they provided the appropriate data needed for the analysis.

Our exclusion criteria were as follows: (1) studies that used non-CNN-based machine learning algorithms (like support vector machine etc); (2) studies that used endoscopic optics other than standard WLE and/or NBI-based images as their training and testing platform; and (3) studies not published in English language. In cases of multiple publications from a single research group reporting on the same patient cohort and/or overlapping cohorts, each reported contingency table was treated as being mutually exclusive. When needed, authors were contacted via email for clarification of data and/or study-cohort overlap.

Data abstraction and analysis

Data on study-related outcomes from the individual studies were abstracted independently onto a predefined standardized form by at least two authors (BPM, SRK). Disagreements were resolved by consultation with a senior author (GK). Diagnostic performance data were extracted and contingency tables were created at the reported thresholds. Contingency tables consisted of reported accuracy, sensitivity, specificity, positive predictive value (PPV) and negative predictive value (NPV). The results from testing of the algorithm were collected for the pooled analysis.

Definitions are as follows: (1) Accuracy: number of lesions detected by CNN/total number of lesions; (2) Sensitivity: detected number of correct neoplastic lesions by CNN (true positives)/histologically confirmed number of neoplastic lesions (total positives); (3) Specificity: detected number of correct non-neoplastic lesions by CNN (true negatives)/number of histologically proven non-neoplastic lesions (total negatives); (4) PPV: detected number of correct neoplastic lesions by CNN (true positives)/number of neoplastic lesions diagnosed by CNN (true positives + false positives); and (5) NPV: number of lesions correctly diagnosed as non-neoplastic lesions by CNN (true negatives)/number of lesions diagnosed as non-neoplastic by CNN (true negatives + false negatives).

If a study provided multiple contingency tables for the same or for different algorithms, we assumed these to be independent from each other. This assumption was accepted, as the goal of the study was to provide an overview of the pooled rates of various studies rather than providing precise point estimates. This methodology has been used and reported in literature [1]. A formal assessment of study quality was not done, due to the non-clinical nature of the studies.

► Table 1 Study characteristics.

Study	Aim	Endoscopy technique	Machine learning model	Training strategy	Testing strategy	Accuracy	Sensitivity	Specificity	PPV	NPV	Remarks
Ahmad, 2019 [11]	Colorectal adenoma detection	Standard colonoscopy	CNN	Multicenter colonoscopy images and videos of 4664 polyp test frames	17 video datasets of complete colonoscopy withdrawal with 83 polyps consisting of 83716 frames (14634 polyp & 69082 non-polyp)	92.5	84.5	92.5	NR	NR	Conference abstract
Byrne, 2019 [2]	Colorectal polyp detection in real-time endoscopic video images	NBI endoscopy	CNN	Unaltered video frames	125 videos of consecutively encountered diminutive polyps	94	98	83	90	97	■
Cai, 2019 [12]	Detect early ESCC under conventional endoscopic white light imaging	White light endoscopy	CNN	1332 abnormal and 1096 normal images	187 images from 57 patients	91.4	97.8	85.4	86.4	97.6	■
Chen, 2018 [13]	Colorectal polyp detection	NBI endoscopy	CNN (TensorFlow algorithm)	1476 images of neoplastic and 681 of hyperplastic polyps	96 hyperplastic and 188 neoplastic polyps smaller than 5 mm	90.1	96.3	78.1	89.6	91.5	■
Cho, 2019 [14]	Classify gastric neoplasms based on endoscopic white-light images	White light endoscopy	CNN (Inception-v4, Resnet-152, Inception-Resnet-v2)	5017 images from 1269 individuals	200 images from 200 patients	93	60.7	98.3	85	93.9	Advanced gastric cancer
						74.5	28.3	88.3	41.9	80.5	Early gastric cancer
						86.4	0	99.4	0	86.9	High-grade dysplasia
						78.5	6.7	91.2	11.3	84.7	Low-grade dysplasia
						66.5	95.7	50.8	51.1	95.7	Non-neoplastic
Guo, 2019 [15]	Real time automated diagnosis of precancerous and early ESCC in both non-magnifying and magnifying settings	NBI endoscopy NBI endoscopy videos	CNN (SegNet architecture)	2770 images of precancerous lesions and early ESCC in 191 cases and 3703 images of non-cancerous lesions in 358 cases	1480 malignant images in 59 cases, 5191 non-cancerous images in 2004 cases, 27 precancerous and early ESCC videos, and 33 normal videos	nr	98.04	95.03	NR	NR	■
						nr	60.8	99.9	NR	NR	■

► **Table 1** (Continuation)

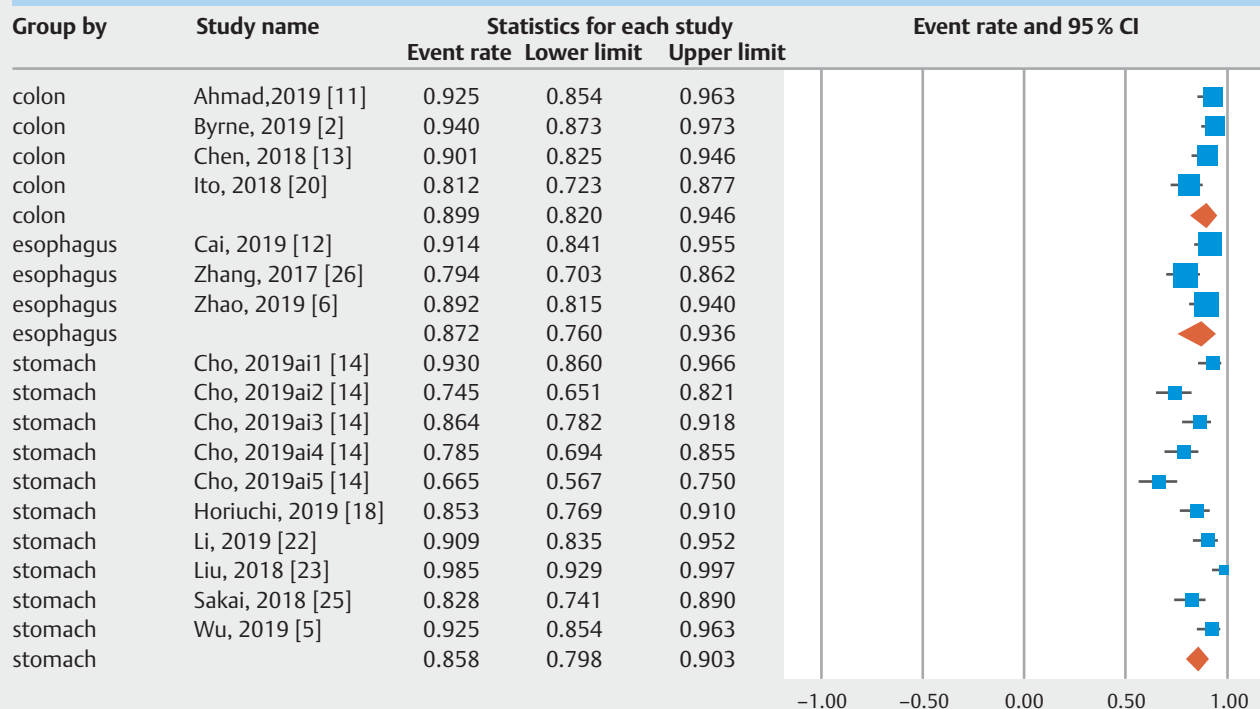
Study	Aim	Endoscopy technique	Machine learning model	Training strategy	Testing strategy	Accuracy	Sensitivity	Specificity	PPV	NPV	Remarks
Hirasawa, 2018 [16]	Detect early and advanced gastric cancer	Standard white-light, chromoendoscopy, NBI	CNN (Single Shot MultiBox Detector)	13584 EGD images for 2639 histologically proven gastric cancer	2296 images from 77 gastric cancer lesions of 69 patients	nr	92.2	nr	30.6	NR	■
Horie, 2018 [17]	Detect esophageal cancer	White-light, NBI	CNN (Single Shot MultiBox Detector)	8428 histologically proven EGD images of cancer in 384 patients	162 images of cancer and 376 images without cancer from 47 patients with 49 cancer lesions, 573 images of non-cancerous areas from 50 patients with no cancer	nr	77	79	39	95	■
Horiuchi, 2019 [18]	Differentiate gastric cancer from gastritis	Magnifying NBI endoscopy	CNN (GoogLeNet)	1492 cancer and 1078 gastritis images	151 cancer and 107 gastritis images	85.3	95.4	71	82.3	91.7	■
Ikenoyama, 2019 [19]	Detect gastric cancer	Standard EGD	CNN	13584 images from 2639 lesions	2940 images from 140 cases (209 early cancer images, 273 non-neoplastic images)	nr	65.6	NR	14.6	NR	Conference abstract
Ito, 2018 [20]	Assist in cT1b colorectal cancer diagnosis	White-light colonoscopy	CNN (AlexNet & Caffe)	Group 1: 2520 cT1s+ cT1a, 2418 cT1b images; Group 2: 2604 cT1s+ cT1a, 2400 cT1b images; Group 3: 2604 cT1s+ cT1a, 2418 cT1b images	190 conventional white-light images	81.2	67.5	89	NR	NR	■
Komeda, 2019 [21]	Colorectal polyp classification	White-light, NBI, chromoendoscopy	CNN	29572 adenoma images, 62999 non-adenoma images	60 cases of colon polyps	nr	97.5	97.9	NR	NR	Conference abstract, white light
						nr	94.8	96.5	NR	NR	Conference abstract, NBI
						nr	90.1	99.5	NR	NR	Conference abstract, chromoendoscopy

► Table 1 (Continuation)

Study	Aim	Endoscopy technique	Machine learning model	Training strategy	Testing strategy	Accuracy	Sensitivity	Specificity	PPV	NPV	Remarks
Li, 2019 [22]	Early gastric cancer detection	Magnifying NBI endoscopy	CNN (InceptionV3-Keras framework)	386 images of non-cancer lesions, 1702 images of early cancer	341 images (170 early cancer & 171 non-cancer lesions)	90.91	91.18	90.64	90.64	91.18	■
Liu, 2018 [23]	Early gastric cancer detection	Magnifying NBI endoscopy	CNN (VGG16, InceptionV3, InceptionResNetV2)	Magnifying NBI of normal gastric images and early gastric cancer images	Images (number not mentioned)	98.5	98.1	98.9	NR	NR	Conference abstract
Ozawa, 2018 [24]	Automatic endoscopic detection and classification of colorectal polyps	White-light & NBI colonoscopy	CNN (Single Shot MultiBox Detector)	16418 images of 4752 histologically proven colorectal polyps and 4013 images of normal colorectum	3533 images	91	92	91	93	NR	Conference abstract
Sakai, 2018 [25]	Detect early gastric cancer	White light endoscopy	CNN (Single Shot MultiBox Detector)	1000 images of 0-I, 0-IIa, 0-IIc lesions	228 cancer images	82.8	73.6	98.8	NR	NR	■
Wu, 2019 [5]	Detect early gastric cancer	EGD	CNN (VGG-16, ResNet-50, Google's TensorFlow)	2204 early cancer, 326 advanced cancer, 4791 control	170 images	92.5	94	91	91.26	93.81	■
Zhang, 2017 [26]	Early esophageal neoplasia	NBI and magnifying endoscopy	CNN	218 endoscopic images generated 218000 patches, 90% used for training and rest for testing	218000 patches, 90% used for training and rest for testing	79.38	73.41	83.54	72.09	84.44	Conference abstract
Zhao, 2019 [6]	Early ESCC using magnifying NBI	Magnifying NBI endoscopy	VGG16	three test groups with 463, 438 & 449 images	1350 images in 219 patients	89.2	87	84.1	81.9	90.4	■

CNN, convolutional neural networks; PPV, positive predictive value; NPV, negative predictive value; ESCC, esophageal squamous cell carcinoma; NBI, narrow band imaging; EGD, esophagogastroduodenoscopy; PET, positron emission tomography; ESD, endoscopic submucosal dissection; VGG, visual geometry group; NR, not reported

Accuracy



► Fig. 1 Forest plot, accuracy.

We used meta-analysis techniques to calculate the pooled estimates in each case following the random-effects model [8]. We assessed heterogeneity between study-specific domain-effects model [8]. We assessed heterogeneity between study-specific estimates by using Cochran Q statistical test for heterogeneity, 95% prediction interval (PI), which deals with dispersion of the effects, and the I^2 statistics [9,10]. In this, values < 30%, 30%–60%, 61%–75%, and > 75% were suggestive of low, moderate, substantial, and considerable heterogeneity, respectively. A formal publication bias assessment was not done due to the nature of the pooled results being derived from the studies.

All analyses were performed using Comprehensive Meta-Analysis (CMA) software, version 3 (BioStat, Englewood, New Jersey, United States).

Results

Search results and study characteristics

The literature search resulted in 4245 study hits (study search and selection flowchart: **Supplementary Fig. 1**). All 4245 studies were screened and 106 full-length articles and/or abstracts were assessed. Nineteen studies [2, 5, 6, 11–26] reported on the detection and/or classification of gastrointestinal neoplastic lesions by CNN. Among the 19 studies, five [6, 12, 15, 17, 26] reported on efficacy of CNN in diagnosing esophageal neoplasia, eight [5, 14, 16, 18, 19, 22, 23, 25] reported on use of CNN in neoplasia of the stomach and six [2, 11, 13, 20, 21, 24] evaluat-

ed use of CNN in diagnosing colorectal neoplasia. Seven studies [5, 11, 12, 14, 19, 20, 25] used standard WLE, eight used NBI (magnifying and/or non-magnifying) [2, 6, 13, 15, 18, 22, 23, 26] and four [16, 17, 21, 24] used a combination of standard WLE and/or NBI and/or chromo-endoscopy images (► **Table 1**).

From all the included studies, we were able to extract a total of 26 contingency table datasets for CNN performance in diagnosing gastrointestinal lesions (► **Table 1**).

Meta-analysis outcomes

CNN performance by gastrointestinal location:

Esophageal neoplasia:

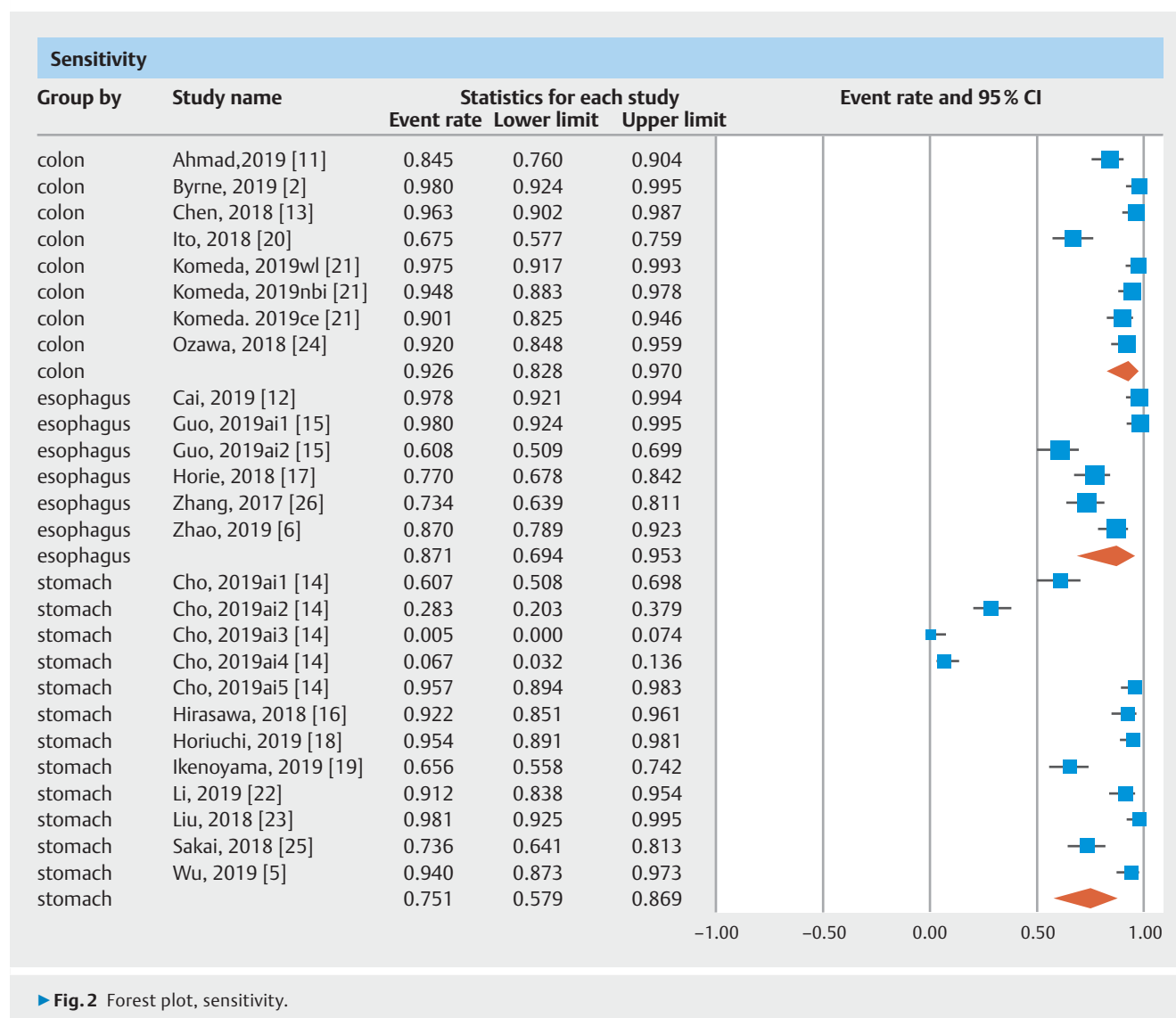
The pooled accuracy of CNN in the computer-aided diagnosis of esophageal neoplasia was 87.2% (95% CI 76–93.6). The sensitivity was 87.1% (95% CI 69.4–95.3), specificity was 87.3% (95% CI 74.3–94.2), PPV was 72.3% (95% CI 41.7–90.5) and NPV was 92.1% (95% CI 85.9–95.7).

Neoplastic lesions in stomach:

The pooled accuracy of CNN in the computer-aided diagnosis of neoplastic lesions of the stomach was 85.8% (95% CI 79.8–90.3). The sensitivity was 75.1% (95% CI 57.9–86.9), specificity was 91.4% (95% CI 84.3–95.4), PPV was 51% (95% CI 30.9–70.8) and NPV was 92.1% (95% CI 85.9–95.7).

Colorectal neoplasia:

The pooled accuracy of CNN in the computer-aided diagnosis of colorectal neoplasia was 89.9% (95% CI 82–94.6). The sensitivity was 92.6% (95% CI 82.8–97), specificity was 92.4%



(95% CI 84.5–96.4), PPV was 91% (95% CI 68.8–97.9) and NPV was 94.3% (95% CI 86.4–97.7).

Results are summarized in ► **Table 1**. Forest plots are shown in ► **Fig. 1**, ► **Fig. 2**, ► **Fig. 3**, ► **Fig. 4**, and ► **Fig. 5**.

Validation of meta-analysis results

Sensitivity analysis

To assess whether any one study had a dominant effect on the meta-analysis, we excluded one study at a time and analyzed its effect on the main summary estimate. On this analysis, no single study significantly affected the outcome or the heterogeneity.

Heterogeneity

A large degree of between-study heterogeneity was expected due to the broad nature of machine learning algorithms and endoscopic optics included in this study. This is reflected in our I^2 values (► **Table 2**). Our subgroup analysis based on tu-

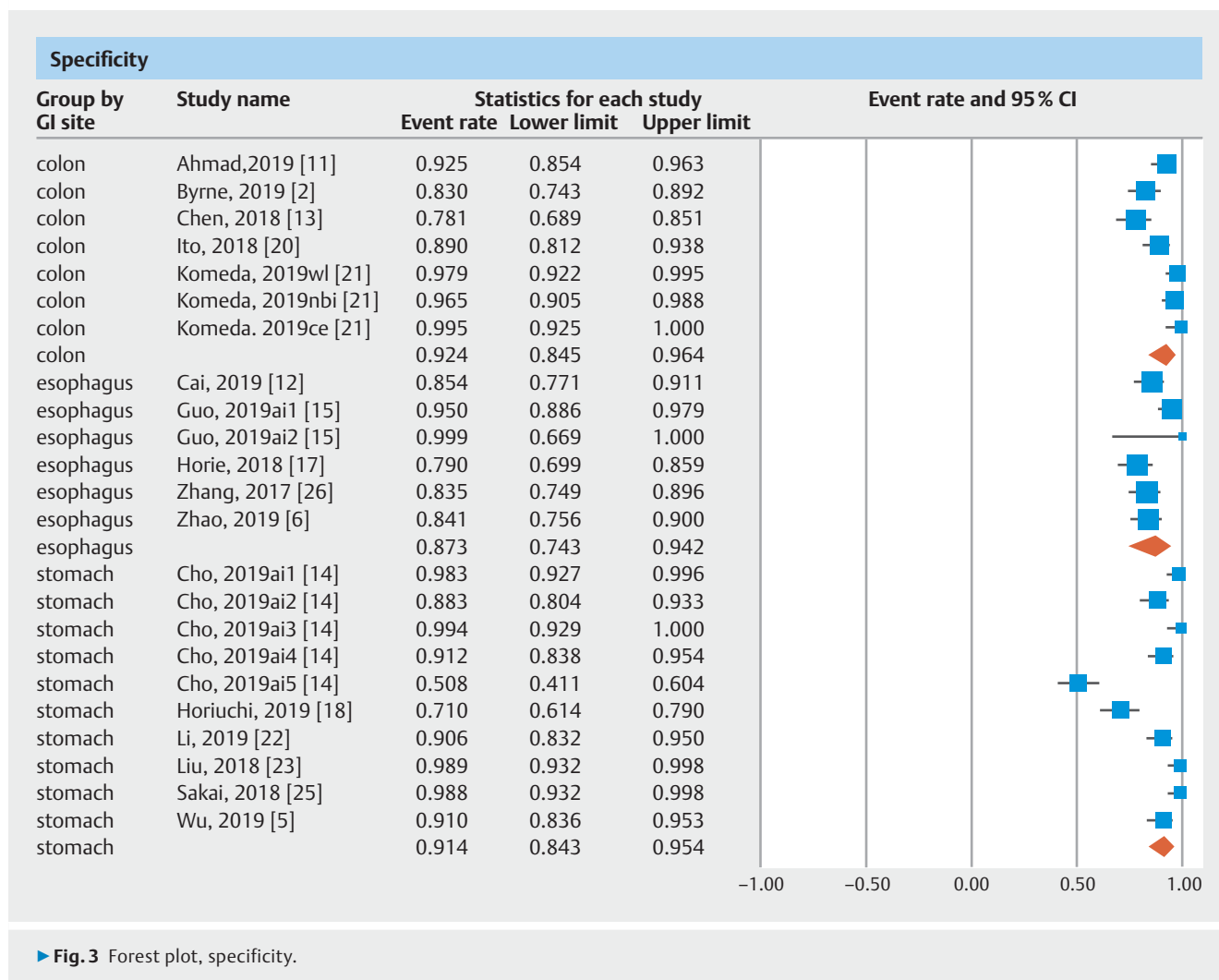
mor location did not affect the observed I^2 values and therefore it can be said that tumor location was not a contributory factor. Prediction interval statistics was not calculated due to the expected large degree of heterogeneity and the fact that the goal was not to provide precise point estimates.

Publication bias

Publication bias assessment largely depends on the sample size and the effect size. A publication bias assessment was deferred in this study due to the fact that the reported effects were independent of the sample size. We, however, do not rule out the possibility of potential publication bias in terms of negative studies being less frequently published.

Quality of evidence

The quality of evidence was rated for results from the meta-analysis according to the GRADE working group approach [27]. Observational studies begin with a low-quality rating, and



based on the risk of bias and heterogeneity, the quality of this meta-analysis would be considered as low-quality evidence.

Discussion

To the best of our knowledge, this is the first systematic review and meta-analysis assessing the accuracy parameters of convolutional neural network (CNN) based computer aided diagnosis of gastrointestinal lesions that includes esophageal, gastric and colorectal data. Based on our analysis, CNN-based deep machine learning demonstrates high accuracy in image-based diagnosis of lesions in esophagus, stomach and colorectum.

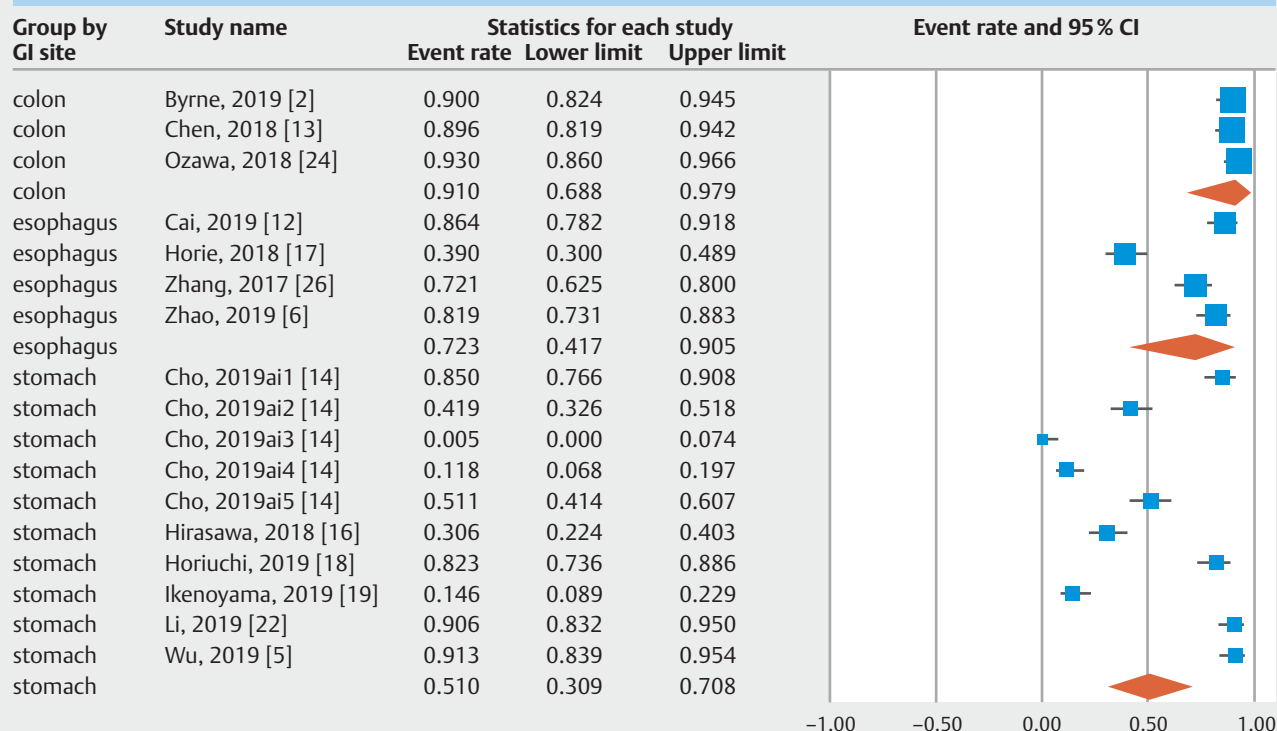
A key finding of our study is that CNN achieved >90% NPV in diagnosis of esophageal, gastric and colorectal lesions. The majority of the included studies evaluated performance of CNN in experimental conditions and not in a real-life clinical scenario. Prospective studies and real-time video analysis of endoscopic images are lacking. Only high-quality images were used to train the CNN. In a real clinical setting, less insufflation of air, post-biopsy bleeding, halation, blur, defocus or mucus can all affect an accurate CAD. There was variability in the choice of threshold used to report sensitivity and specificity. There was lack in

uniformity of validating the training process of the algorithm before using it for testing.

A recent meta-analysis published by Liu et al [28] reported similar diagnostic accuracy results for use of AI in prediction and detection of colorectal polyps. They reported better performance for AI under NBI and performance superior to that of non-expert endoscopists. This study primarily differs in the reported AI parameters for esophageal, gastric, and colorectal lesions. In addition, we did not include studies that primarily assessed the nuances of mathematical formulae behind the CNN algorithm and we did not include studies that used support vector machine-based algorithm.

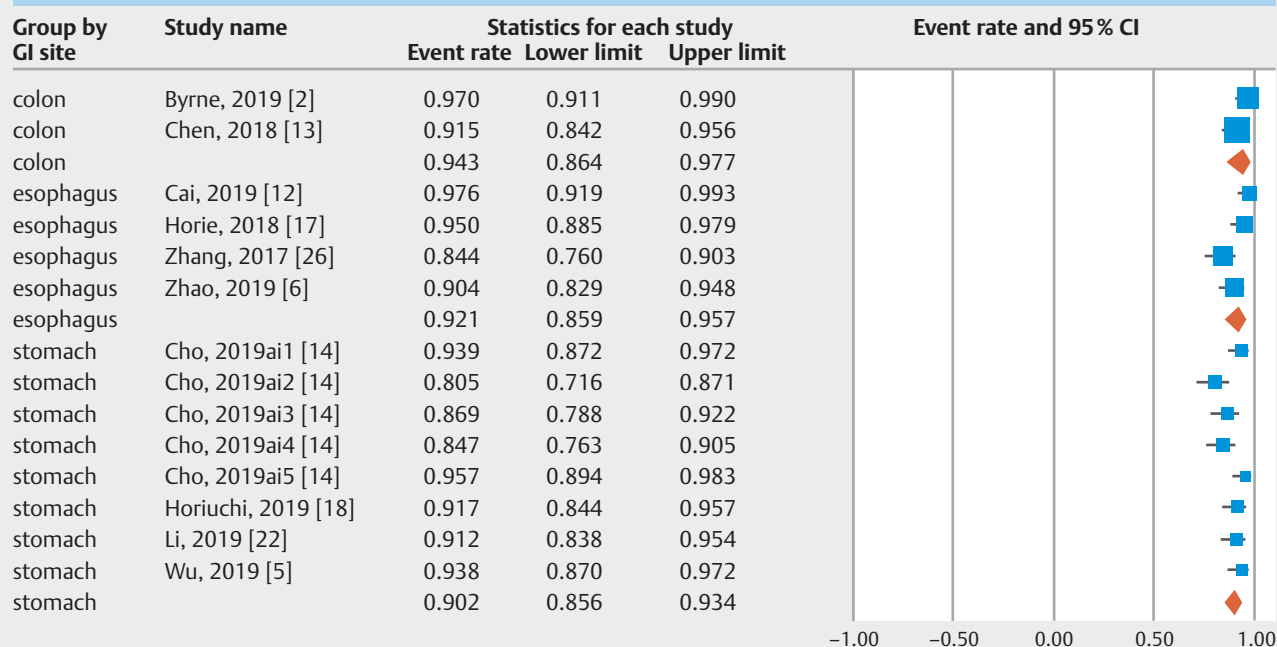
The strengths of this review lie in careful selection of studies reporting on machine-based learning that is solely based on CNN-based algorithms and avoiding other redundant studies. The American Society of Gastrointestinal Endoscopy (ASGE) in its second Preservation Incorporation of Valuable Endoscopic Innovations (PIVI-2) declaration proposed a NPV threshold of 90% or greater for real-time optical diagnosis of diminutive colorectal polyps using advanced endoscopic imaging [29]. We have demonstrated that CNN achieves this threshold in CAD of gastrointestinal lesions regardless of their location.

PPV



► Fig. 4 Forest plot, PPV.

NPV



► Fig. 5 Forest plot, NPV.

Table 2 Summary of results.

Pooled rates	Accuracy	Sensitivity	Specificity	PPV	NPV
Esophagus	87.2 (76–93.6) $I^2=70$ 3 datasets	87.1 (69.4–95.3) $I^2=90$ 6 datasets	87.3 (74.3–94.2) $I^2=60$ 6 datasets	72.3 (41.7–90.5) $I^2=95$ 4 datasets	92.1 (85.9–95.7) $I^2=74$ 4 datasets
Stomach	85.8 (79.8–90.3) $I^2=83$ 10 datasets	75.1 (57.9–86.9) $I^2=96$ 12 datasets	91.4 (84.3–95.4) $I^2=92$ 10 datasets	51 (30.9–70.8) $I^2=97$ 10 datasets	90.2 (85.6–93.4) $I^2=64$ 8 datasets
Colorectal	89.9 (82–94.6) $I^2=69$ 4 datasets	92.6 (82.8–97) $I^2=88$ 8 datasets	92.4 (84.5–96.4) $I^2=81$ 7 datasets	91 (68.8–97.9) $I^2=0$ 3 datasets	94.3 (86.4–97.7) $I^2=61$ 2 datasets

CNN, convolutional neural network; PPV, positive predictive value; NPV, negative predictive value

There are limitations to this study. The included studies were not representative of the general population and community practice, with studies being performed in an experimental environment. Our analysis had studies that were retrospective in nature contributing to selection bias. To capture maximum available data, we included six conference abstracts that have not been published as full manuscripts yet. We were unable to formally conduct a quality assessment, as there is no guidance on how to appropriately score and report quality on items pertaining to machine-based learning. Moreover, we considered individual accuracy tables as independent of each other, which does not reflect real-life case scenario.

Our analysis has the limitation of heterogeneity. We were unable to statistically ascertain a cause for the observed heterogeneity. We hypothesize, however, that the observed heterogeneity is primarily due to the following variables: threshold cut-off used, different training algorithm as well as the training methodology employed, and the variability in endoscopic optics (standard white-light, NBI, chromo-endoscopy). In addition, endoscopic optics differ in their diagnostic accuracy based on the underlying gastrointestinal lesion being assessed. In terms of algorithm training and testing, not all studies employed a validation step to fine-tune the algorithm. Therefore, the possibility of over-fitting in the reported accuracy data is possible.

We only included studies that evaluated the performance of CNN-based algorithms and not others, such as support vector machine algorithms (SVM). This is due to the inherent mathematical differences in the algorithms that make CNNs highly unique and superior performers when compared to SVMs, and due to the fact that SVMs are less likely to be used for image classification in the near future. Although the technology is rapidly advancing in AI, we do not anticipate that CNN-based deep learning will become obsolete before further real-life prospective studies are reported. We do, however, anticipate rapid technical improvements in CNN algorithms in terms of faster processing times despite an increase in number of deep hidden learning layers, and the implementation of positive reinforcement in CNN learning that allows the algorithm to learn from

its errors and encourages it to execute a correct neuron while inhibiting a wrong one.

Conclusions

In conclusion, based on our meta-analysis, deep machine learning by means of CNN-based algorithms demonstrates high accuracy in diagnosis of gastrointestinal lesions. Deep learning in gastroenterology is in its infancy and is witnessing a rapid, steep growth in terms of learning as well as technological development. Future studies are needed to streamline the machine-learning process and define its role in the CAD of gastrointestinal neoplastic conditions in real-life clinical scenarios.

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Competing interests

Dr. Dulai received an American Gastroenterology Association Research Scholar Award. He is a consultant for Takeda, Janssen, Pfizer, and Abbvie. He has also received grant support from Takeda, Janssen, Pfizer, and Abbvie.

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