ENDOUROLOGY

Invited Review

Artificial intelligence (AI) in urology-Current use and future directions: An iTRUE study

Milap Shah^{1,2} , Nithesh Naik^{2,3} , Bhaskar K. Somani^{2,4} , BM Zeeshan Hameed^{1,2,5}

Cite this article as: Shah M, Naik N, Somani BK, Hameed BMZ. Artificial intelligence (AI) in urology-Current use and future directions: An iTRUE study. Turk J Urol 2020; 46(Supp. 1): S27-S39.

ABSTRACT

Objective: Artificial intelligence (AI) is used in various urological conditions such as urolithiasis, pediatric urology, urogynecology, benign prostate hyperplasia (BPH), renal transplant, and uro-oncology. The various models of AI and its application in urology subspecialties are reviewed and discussed.

Material and methods: Search strategy was adapted to identify and review the literature pertaining to the application of AI in urology using the keywords "urology," "artificial intelligence," "machine learning," "deep learning," "artificial neural networks," "computer vision," and "natural language processing" were included and categorized. Review articles, editorial comments, and non-urologic studies were excluded.

Results: The article reviewed 47 articles that reported characteristics and implementation of AI in urological cancer. In all cases with benign conditions, artificial intelligence was used to predict outcomes of the surgical procedure. In urolithiasis, it was used to predict stone composition, whereas in pediatric urology and BPH, it was applied to predict the severity of condition. In cases with malignant conditions, it was applied to predict the treatment response, survival, prognosis, and recurrence on the basis of the genomic and biomarker studies. These results were also found to be statistically better than routine approaches. Application of radiomics in classification and nuclear grading of renal masses, cystoscopic diagnosis of bladder cancers, predicting Gleason score, and magnetic resonance imaging with computer-assisted diagnosis for prostate cancers are few applications of AI that have been studied extensively.

Conclusions: In the near future, we will see a shift in the clinical paradigm as AI applications will find their place in the guidelines and revolutionize the decision-making process.

Keywords: Artificial intelligence; deep learning; machine learning; prostate cancer; urolithiasis; urology.

Introduction

Artificial intelligence (AI) refers to the computational capability of the machine to mimic and perform human cognitive tasks. It is causing a paradigm shift in terms of providing health care and decision-making for the clinicians. The advances in the medical technologies used in health care, such as electronic medical records (EMRs), are providing humongous amounts of data.^[1] This large amount of data allows computer-based predictions and decisions to be made to aid in better patient care (Figure 1). By 2025, the growth rate of AI applications in health care is expected to be 29.3%, and the global revenue is estimated to increase by 40%.^[2] With the available patient data, the future health care system is likely to move toward AI outpatient clinics and preventive medicine. AI provides more accuracy and reliable clinical decisions; hence, it is possibly going to be an integral part of the health care system.

The four subfields of AI in health care are as follows:

1. Machine learning (ML): ML is statistical technique-based programming that allows a computer system to learn and recognize patterns to model without explicit instructions. ML uses procedural computer programs wherein machines are trained to learn, detect data patterns, compute, and infer from the

ORCID iDs of the authors: M.S. 0000-0003-1668-7006; N.N. 0000-0003-0356-7697; B.K.S. 0000-0002-6248-6478; B.M.Z.H. 0000-0002-2904-351X

¹Department of Urology, Kasturba Medical College Manipal, Manipal Academy of Higher Education, Manipal, Karnataka, India ²i-TRUE: International Training and Research in Uro-oncology and Endourology, Manipal, Karnataka, India ³Department of Mechanical and Manufacturing Engineering, Manipal Institute of Technology, Manipal Academy of Higher Education, Manipal, Karnataka. India ⁴Department of Urological Surgery, University Hospital Southampton NHS Trust, Southampton, UK ⁵KMC Innovation Centre. Manipal Academy of Higher Education, Manipal, Karnataka, India

Submitted: 09.04.2020

Accepted: 12.04.2020

Available Online Date: 27.05.2020

Corresponding Author:

BM Zeeshan Hameed E-mail: zeeshanhameedbm@gmail.com

©Copyright 2020 by Turkish Association of Urology

Available online at www.turkishjournalofurology.com



Figure 1. A schematic process chart for building an artificial intelligence model

datasets provided. It is observed that the machines are able to generate results similar to those generated by human intelligence.

2. Natural language processing (NLP): It illustrates the ability of a computer to comprehend the written and spoken language. Some applications that are possible through NLP include language translation, text processing, and speech recognition. In order to extract useful information and reliable details from patient services and provide "virtual assistance" for physicians, a comprehensive data research such as electronic medical record (EMR), doctor's notes, pharmaceutical products, and medical imaging can also be analyzed.

3. Deep learning (DL) and artificial neural networks (ANNs): In the network architectural layers, the ANN comprises of individual units that function like artificial neurons programmed to accomplish computer tasks and recognize complex patterns. DL requires training massive datasets of multilayered neural networks. Deep neural convolution network (DCNN) is a commonly used ANN, which is effective when used in digitized image pattern identification or recognition.

Main Points:

- AI is widely used in the diagnosis, treatment, and outcome prediction in various urological conditions.
- In urolithiasis, AI is used to detect stone composition and to predict spontaneous passage of stone.
- AI applications in prostate carcinoma are used for the diagnosis-Gleason scoring, treatment decisions-making, and even predicting the disease-free survival.

4. Computer vision: Computer vision technology is used for visual search, trend forecasting, augmented reality, and virtual reality. The radiological and pathological images and simple and complex endoscopic videos can be used by machines to understand the details and patterns in the images in order to identify the tumors or malignancy present in the diagnostic imaging has already shown that AI has extensive "knowledge" to identify tumors. Computer vision can also be used for analysis and grading of pathological tissue slides.

AI is increasingly applied not only to the diagnosis of urological conditions but also its management and predictive analysis. ^[3] This article focuses on addressing the application of AI and AI algorithms in urological subspecialties. This article reviews the use of AI in various benign and malignant conditions such as urolithiasis, pediatric urology, urogynecology, benign enlargement of the prostate, renal transplant, and uro-oncology pertaining to the kidneys, bladder, prostate, and testes.

Clinical and Research Consequences

Application of AI in Benign Urological Conditions

Urolithiasis

In the past few decades, there has been a rapid transition in the analysis, treatment and monitoring of cases with urolithiasis, the recent entry being applications of AI to identify the stone from computed tomography (CT) and ultrasound (US) images^[4,5], detecting stone composition^[6,7], predicting spontaneous stone passage^[8,9] and even the outcomes of endourological procedures (Table 1).^[10-13]

Table 1. Studies looking at applications of AI in benign urological conditions						
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity
A. Urolithiasis						
Parakh et al. ^[4]	Urinary stone detection on unen- hanced CT images	 535 patients (279 stones present; 256 stones absent) 100 scans (test data) 	Convolutional neural network (CNN)	>90%	NA	NA
Chiang et al. ^[5]	To identify asso- ciation between stone disease and genetic polymorp- hisms, patient habits	 151 (calcium oxalate stone) patients 105 healthy controls 	 Discriminant analysis Artificial Neural network (ANN) 	 Genetic factors DA: 64% ANN: 65% Genetic and Env. Factors DA: 75% ANN: 89% 	NA	NA
Kazemi et al. ^[6]	For early detecti- on of the type of kidney stone and the most influenti- al parameters	936 patients dataset42 features	 -Bayesian model-Decision Trees ANN Rule-based classifiers 	97.1% (ensemble model)	NA	NA
Kriegshauser et al. ^[7]	To investigate use of single-source dual-energy com- puted tomography (ssDECT) for the characterization of renal stones.	• 32 stone dataset	-Multiparametric algorithms	 97% to distinguish uric acid and non-uric acid stones 72% to distinguish non-uric acid stone subtypes 	NA	NA
Eken et al. ^[8]	To compare AI models with LR by applying on medical dataset	 227 patients 176 urinary stones 51 no stones 	 ANN Genetic algorithm (GA) Logistic regression analysis (LR) 	NA	ANN: 94.9% GA: 67.6% LR: 95.5%	ANN: 78% GA: 76% LR: 48%
Dal Moro et al. ^[9]	To predict the spontaneous passage of ureteral stones in patients with renal colic	 1163 patients (402 found valuable) nine clinical factors 	Linear program- ming support vector machine (LPSVM)	NA	84.5%	86.9%
Shabaniyan et al. ⁽¹⁰⁾	To predict posto- perative outcome of PCNL	 254 patients 26 variables	• Machine learning (ML) techniques such as sequential forward selecti- on and Fisher's discriminant analysis	94.8%	Requirement of Stent placement: 85.2% Require- ment of blood transfusion: 95%	NA

Table 1. Studies	looking at applica	tions of AI in ben	ign urological con	ditions (Continue)	
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity
Aminsharifi et al. ^[11]	To predict mul- tiple outcomes after percutaneous nephrolithotomy (PCNL) and compare with GSS and CROES nomogram	• 146 adult pati- ents	• ML-based SVM model	80%-95.1%	Stone free status: 92% Need for repeat PCNL: 97% Need for ESWL: 82% Need for URS: 91%	NA
Kadlec et al. ^[12]	For outcome prediction after various forms of endourological intervention	• 382 renal units	• nonlinear LR model	Classification ac- curacy of 69.6%	Stone free status: 75.3% Need for sec. procedure: 30%	Stone free status: 60.4% Need for sec. procedure: 98.3%
Seckiner et al. ^[13]	To predict stone free status after ESWL	 139 patients (training set) 32 patients (validation set) 32 patients (test set) 	ANN	88.7% in the test group	NA	NA
B. Benign Prostation	c Hyperplasia					
Torshizi et al. ^[14]	To diagnose the severity of BPH and suggest app- ropriate treatment	• 44 patients	Fuzzy system expert	~90%	NA	NA
Sonke et al. ^[15]	To diagnose BPH and compare with regression analysis	• 1903 patients	ANN	NA	71%	69%
C. Pediatric Urolog	gy					
Bagli et al. ^[16]	To predict sonog- raphic outcome after pyeloplasty in children with ureteropelvic junc- tion obstruction	 84 children training set 16 children test set 	ANN	100%	100%	100%
Logvinenko et al. ^[17]	To predict patients at high risk of VCUG abnorma- lities, based on RBUS findings	• 2259 patients	ANN Multivariate LR analysis	NA	For any grade VUR • ANN: 64% • MLR: 84%	For any grade VUR • ANN: 60% • MLR: 25%
Blum et al. ^[18]	To predict the need for surgery in UPJO cases based on dynamics of renogram	55 patients45 features	• Linear support vector machine (SVM)	93%	91%	96%
D. Urogynecology						
Sabadell et al. ^[20]	To predict occur- rence of SUI after prolapse surgery and as a diagnostic tool	• 169 patients	ML algorithm	NA	NA	NA

Table 1. Studies looking at applications of AI in benign urological conditions (Continue)							
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity	
Jelovsek et al. ^[21]	To predict recur- rence, complica- tions, and health status improve- ment after prolapse surgery	• 1301 patients	LR models	NA	NA	NA	
E. Renal Transplan	nt						
Atallah et al. ^[22]	To predict 5-year graft survival	• 2728 patients (70% training and 30% testing)	 Naïve Bayes Based Feature Selector (NBBFS) Algorithm K-nearest neighbor Algorithm (KNN) 	80.77%	81.2%	NA	
Greco et al. ^[23]	To predict graft fa- ilure and associa- tion with BMI and other risk factors	• 194 patients	ML algorithmsDecisional Trees	NA	88.2%	73.8%	
Goldfarb et al. ^[24]	To predict cadave- ric graft survival over three years based on pretrans- plant variables	37,407 patients dataset	Logistic regression based model Tree- based model	65%	NA	NA	

CT: Computed Tomography; CNN: Convolutional Neural Network; ANN: Artificial Neural Network; DA: Discriminant Analysis; ssDECT: Single-Source Dual-Energy Computed Tomography; LR: Logistic regression; GA: Genetic Algorithm; LP SVM: Linear Programming Support Vector Machine; PCNL: Percutaneous Nephrolithotomy; ML: Machine Learning; GSS: Guys Stone Score; CROES: Clinical Research Office of the Endourological Society; SVM: Support Vector Machine; ESWL: Extracorporeal Shock Wave Lithotripsy; URS: Ureterorenoscopy; BPH: Benign Prostatic Hyperplasia; VCUG: Voiding Cystourethrogram; RBUS: Renal Bladder Ultrasound; NBBFS: Naïve Bayes Based Feature Selector; KNN: K-Nearest Neighbor; BMI: Body Mass Index; UPJO: Uretero-Pelvic Junction Obstruction

Parakh et al.^[4] studied the diagnostic performance of the convolution neural network (CNN) on CT images for detection of urinary stones in 535 adult patients assumed to have renal calculi using two scanners. The first scanner identified the urinary tract, and the next one detected the stone. Using nine different variation models, it achieved an accuracy of more than 90%. The study concluded that the efficiency of CNNs can be improved by the use of transfer learning with datasets augmented with labeled images. Shabaniyan et al.^[10] developed a decision support system using ML techniques to predict the outcomes of surgical treatment for renal calculus. The algorithm was trained with a dataset of 254 patients and 26 parameters, which comprised variables from patients' history, renal calculus composition, and laboratory investigations. This model achieved an accuracy of 94.8%, 85.2%, and 95% in predicting outcomes of a procedure, predicting whether patient will require a stent after the procedure, and predicting the need for blood transfusion, respectively. Aminsharifi et al.^[11] studied data of 146 adult patients in whom percutaneous nephrolithotomy (PCNL) was done to validate efficiency of a machine-based learning algorithm for predicting the outcomes after PCNL and to compare the results with Clinical Research Office of Endourological

Society nomogram and Guy's Stone Score (GSS). This program predicted the PCNL results with an accuracy of up to 95%.

Benign Enlargement of the Prostate

Many questionnaires are available for the clinical prediction of benign prostatic hyperplasia (BPH), yet the results are unreliable and inaccurate. Various AI techniques and ANN models such as multilayered back propagation method to predict the severity of obstruction on the basis of noninvasive tests have been used (Table 1).^[14,15] Torshizi et al.^[14] applied fuzzy intelligent systems in predicting the severity of BPH and also recommended the treatment required for it. The study consisted of two models. The first model predicted the severity, whereas the second model helped to make a treatment decision. The results were then compared for accuracy and validation with an expert panel. The accuracy achieved was nearly 90%.

Pediatric Urology

AI has been used in the field of pediatric urology for predicting the outcome of surgical procedures^[16], severity of the condition on the basis of imaging as well as detecting abnormalities in imaging (Table 1).^[17,18] Bagri et al.^[16] applied computerized ANN to predict the outcomes after pyeloplasty on the basis of US findings in children with uretero-pelvic junction obstruction. The prediction was based on whether postoperatively the results were "significantly improved," "improved," "same," or "worse." The results showed 100% sensitivity and specificity for all four-outcome measures. Both multivariate analysis and ML algorithms were used by Logvinenko et al.^[17] to evaluate whether the renal and bladder ultrasound (RBUS) could predict the abnormalities on voiding cystourethrogram (VCUG) for conditions such as vesico-ureteric reflux and congenital ure-thral abnormalities. The results showed that RBUS was a poor predictor of the abnormalities of VCUG and both could only complement each other but cannot replace them.

Urogynecology

On the basis of urinary incontinence data obtained from the wearable devices, AI techniques were applied to predict the time and number of incontinence episodes and the outcome of conservative or medical management for stress urinary incontinence (SUI).^[19] Models to predict the occurrence of complications such as SUI after prolapse surgery, recurrence, and overall outcomes of surgery were also studied on the basis of the data available from various randomized controlled trials (Table 1).^[20,21] In the near future, AI applications can be used to provide personalized care based on the patient demographics and clinical characteristics of every individual.

Renal Transplant

The outcome of kidney transplant prediction is very important. Various studies have been conducted to predict the outcomes of kidney transplantation using ANN and ML algorithms, as described (Table 1).^[22-24] Atallah et al.^[22] proposed a prediction method by combining two methods-Bayes and k-nearest neighbor-which achieved more accuracy by choosing minimum number of features. It was based on data mining techniques to predict five-year graft survival after transplantation. This new proposed prediction method comprises three stages: data preparation stage, feature selection stage, and prediction stage. This prediction method can be used in other transplant datasets to measure the graft survival.

Application of AI in Uro-Oncology

Testicular Malignancy

Not much has been studied about the applications of AI in testicular malignancy. Baessler et al.^[25] applied ML-based CT radiomics to determine whether the lymph nodes dissected in patients with metastatic or advanced nonseminomatous testicular germ cell tumor were malignant or benign. The model correctly classified with an accuracy of 0.81 (area under the curve [AUC]), 88% sensitivity, and 72% specificity.

Renal Cell Carcinoma

ML and DL algorithms based on CT-texture analysis were applied for differentiating renal masses such as angiomyolipoma, clear cell renal cell carcinoma (ccRCC), papillary renal cell carcinoma, and oncocytoma^[26-28] to predict the nuclear grade and to identify certain genetic mutations to predict the prognosis, recurrence, and survival outcomes (Table 2). Kocak et al.^[26] used CT-texture analysis, applied ML techniques to predict and identify the nuclear grade (Furhman) of ccRCC, and compared the results with those obtained with percutaneous biopsy. The results were comparable, and the maximum predictive value was achieved with the use of the support vector machine (SVM). The algorithm could differentiate nuclear grades in 85.1% of ccRCC cases. Ding et al.^[29] also conducted a similar study showing increased precision in classifying the grade of ccRCC.

Biomarkers and signatures based on more than one gene expression have been developed in recent years for predicting the ccRCC overall survival (OS) and prognosis of the disease. Li et al.^[30] developed a model based on 15 genes, which could help predict the prognosis and survival. They found that the group with a higher risk had substantially poorer prognosis and survival than the group with patients having lower risk. The risk groups were not associated with patient characteristics such as sex or age but were related to hemoglobin levels. They were also associated with tumor features such as size and grade.

PBRM1 mutations are the second most common mutations found in ccRCC. Kocak et al.^[31] applied ANN- and ML-based algorithms to identify PBRM1 mutations based on CT scan texture analysis. Overall, 88% of ccRCC with PBRM1 mutation status was correctly identified by ANN. On the basis of these results, future studies can be conducted to develop noninvasive biomarkers for identifying histopathological subtypes to predict the prognosis and response to treatment.

Bladder Cancer

ML algorithms, DCNN models, genetic algorithms, and SVMs have been applied in bladder cancer for improving cystoscopic diagnosis and prediction of prognosis and survival (Table 2).^[32-35] Ikeda et al.^[32] made a competent CNN by training it with 2102 cystoscopic pictures with an aim to increase the efficiency in diagnosis of bladder cancer using AI. It achieved sensitivity and specificity of 89.7% and 94.0%, respectively. Lorencin et al.^[33] used the data of 1997 and 986 images with and without bladder cancer, respectively, to train multilayer perceptron along with DCNN for the diagnosis of bladder malignancy. It showed promising results, with AUC value reaching up to 0.99. Wang et al.^[36] achieved more than 75% accuracy by using least squares SVM in predicting the five-year overall and cancerspecific mortality of patients post radical cystectomy.

Table 2. Studies looking at application of AI in urological malignancies						
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity
A. Renal Cell Car	cinoma(RCC)					
Kocak et al. ^[26]	To distinguish the major subt- ypes of RCC	 68 RCC patients for internal validation 26 RCC patients for external validation 275 CT images for texture features 	Artificial Neural Network (ANN) Support vector machine (SVM)	ANN: 84.60% SVM: 69%	ANN: 69% SVM: 71%	ANN: 100% SVM: 100%
Feng et al. ^[27]	To differentiate angiomyolipoma (AML) and RCC based on texture analysis of CT images	58 patients42 features	 Machine Learning (ML) based quanti- tative texture analysis SVM with re- cursive feature elimination Synthetic minority oversampling technique (SMOTE) 	93.9%	87.8%	100%
Coy et al. ^[28]	To distinguish ccRCC and on- cocytoma from MDCT images	 4000 iterations (90% training and 10% vali- dation) 179 patients 	Deep Learning (DL) based Go- ogle TensorFlow software	74.4%	85.8%	NA
Ding et al. ^[29]	To preoperatively distinguish high nuclear grade from low nuclear grade in ccRCC	• 92 cases (for validation)	 Logistic Regression (LR) model Least absolute shrinkage and selection operator (LASSO) for texture score 	NA	NA	NA
Li et al. ^[30]	To predict sur- vival in patients with ccRCC based on gene expression	N=533 (training dataset) Risk score model based on 15 ge- nes N=101 (test dataset)	• ML-based ran- dom forest va- riable hunting Cox regression analysis	NA	NA	NA
Kocak et al. ^[31]	To identify the mutation status of PBRM1 gene in ccRCC pati- ents	 45 patients (29 without mutation; 16 with mutation) 161 labeled segmentations (87 without mutation; 74 with mutation) 	 ML-based quantitative CT-texture analysis such as Random Forest (RF) algorithm ANN 	ANN: 88.22% RF: 95%	NA	NA

Table 2. Studies looking at application of AI in urological malignancies (Continue)						
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity
B. Bladder Carcin	ioma					
Ikeda et al. ^[32]	To improve cystoscopic diag- nosis of bladder cancer using AI	 2102 images (1671 normal tissue; 431 tumor lesions) 8:2 (training: test set) 	Convolutional Neural Network (CNN)	NA	89.7%	94%
Lorencin et al. ^[33]	To use multilayer perceptron met- hod for diagnosis of bladder cancer from cystoscopic images	 1997 bladder cancer images 986 non- cancer tissues images 	Multilayer Per- ceptron (MLP)Laplacian edge detector	NA	NA	NA
Hashemi et al. ^[34]	To classify cystoscopic bladder images using AI	• 540 cystos- copic bladder images	Multilayer ne- ural networksGenetic algo- rithm (GA)	7% decrease in error on classification as compared with other methods	NA	NA
Eminaga et al. ^[35]	To perform diagnostic clas- sification based on cystoscopic images using DL-CNN	 479 patients 18,681 images (generated with 10 degree grades) 60% training set-10% vali- dation set 30% test set 	 Deep Learning CNN (DL- CNN) Xception model ResNet50 model InceptionV3 VGG-19 VGG-16 	F1 scores Xcep- tion: 99.52% ResNet: 99.48%	NA	NA
Wang et al. ^[36]	To predict blad- der cancer prog- nosis in terms of five-year overall and cancer-speci- fic mortality	• 117 bladder cancer patients	• Output-based transfer lear- ning approach with least square support vector machine (LS-SVM)	 5 years overall mortality Proposed classifier(v1): 76.97% Proposed classifier(v2): 76.18% 5-year cancer- specific morta- lity Proposed classifier(v1): 74.85% Proposed classifier(v2): 75.15% 	 5 years overall mortality Proposed classifier(v1): 78.48% Proposed classifier(v2): 78.29% 5-year cancer- specific morta- lity Proposed classifier(v1): 90.26% Proposed classifier(v2): 92.38% 	 5 years overall mortality Proposed classifier(v1): 75.79% Proposed classifier(v2): 74.33% 5-year cancer- specific morta- lity Proposed classifier(v1): 38% Proposed classifier(v2): 31%
Gavriel et al. ^[37]	To predict five- year prognosis of bladder cancer	• 78 patients diagnosed with MIBC	• ML-based en- semble model	94.8%	89.5%	97.4%

Table 2. Studies looking at application of AI in urological malignancies (Continue)							
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity	
Hasnain et al. ^[38]	To predict postcystectomy recurrence and survival	• Dataset of 3503 patients	 Ensemble ML-based models Support vector machine (SVM) K-nearest neighbor Algorithm (KNN) Random Forest Gradient-boosted trees (GBT) 	NA	>70%	>70%	
Bartsch et al. ^[39]	To predict recur- rence of NMIBC based on genome profile	 112 frozen NMIBC speci- mens 21 gene classi- fier set 	• ML cased genetic programming algorithm	NA	Test Set Five gene combined rule: 69% Three gene com- bined rule: 71%	Test Set Five gene combined rule: 62% Three gene com- bined rule: 67%	
Wu et al. ^[40]	To compare different DL- CNN models to predict response to treatment in bladder cancer (T0 prediction)	• 123 CT scans (pre and post- treatment)	Multiple DL- CNN models with structure modification and layer freezing	Base DL-CNN 70%	Base DL-CNN 60%	Base DL-CNN 80%	
C. Prostate Carci	noma						
Ström et al. ^[41]	To diagnose and grade prostate cancer in biop- sies	Training set-976 patients (6682 slides) Test set-246 patients (1631 slides)	ANN	NA	99%	94.9%	
Bulten et al. ^[42]	To assign Gle- ason grade to prostate biopsies using AI	1243 patients (5759 biopsies)	DL system	Benign versus malignant: 96%- 97% Grade group 2 or more: 79%-83% Grade group 3 or more: 76%-82%	Benign versus malignant: 97.4% Grade group 2 or more: 86%-95% Grade group 3 or more: 76%-92%	Benign versus malignant: 83%- 100% Grade group 2 or more: 52%-70% Grade group 3 or more: 72%-782%	
Viswanath et al. ^[44]	To compare va- rious classifier in detecting CaP on t2W MRI images using radiomic texture features	• 85 T2W MRI datasets	 Quadratic Discriminant Analysis (QDA) -Sup- port Vector Machines (SVMs) Naïve Bayes Decision Trees (NBDT) 	NA	NA	NA	

Table 2. Studies looking at application of AI in urological malignancies (Continue)							
Study	Objective	Study Design	Algorithm/Model	Accuracy	Sensitivity	Specificity	
Wildeboer et al. ^[45]	For automated localization of CaP based on radiomics of TRUS	 50 men with biopsy confir- med CaP 	ML techniques using B-mode, shear-wave elas- tography (SWE), and dynamic contrast-enhan- ced ultrasound (DCE-US) radiomics	NA	NA	NA	
Deng et al. ^[46]	For treatment stratification of patients with me- tastatic castrate resistant CaP	• 78 features associated with the patient cli- nical and me- dical history, lab reports and metastases	ML-based model	NA	NA	NA	
de la Calle et al. ^[47]	To predict recurrence and progression of CaP based on bi- omarker analysis	 648 samples (424 tumors, 224 normal tissue) Tissue micro assays anti Ki-67, ERG antibodies 	AI algorithm	100% in identifi- cation of ERG+ tumor	NA	NA	
Bibault et al. ^[48]	To predict sur- vival in patients with CaP	 Dataset from PLCO trial 8776 patients (diagnosed with CaP on follow-up) n=7021 (training set) N=1755 (test set) 	AI algorithm	10-year OS: 87% 10-year cancer- specific survival: 98%	10-year OS: 60% 10-year cancer- specific survival: 55%	NA	

RCC: Renal Cell Carcinoma; CT: Computed Tomography; ANN: Artificial Neural Network; SVM: Support Vector Machine; AML: Angiomyolipoma; ML: Machine Learning; SMOTE: Synthetic Minority Oversampling Technique; ccRCC: Clear Cell Renal Cell Carcinoma; MDCT: Multiple Detector Computed Tomography; DL: Deep Learning; LR: Logistic Regression; LASSO: Least Absolute Shrinkage and Selection Operator; PBRM1: Polybromo1; RF: Random Forest; CNN: Convolutional Neural Network; MLP: Multi Layer Perceptron; AI: Artificial Intelligence; GA: Genetic Algorithm; DL-CNN: Deep Learning Convolutional Neural Network; LS-SVM: Least Square Support Vector Machine; MIBC: Muscle Invasive Bladder Cancer; KNN: K-Nearest Neighbor; GBT: Gradient-Boosted Trees; NMIBC: Non-Muscle Invasive Bladder Cancer; CaP: Carcinoma Prostate; T2W MRI: T2 Weighted Magnetic Resonance Imaging; QDA: Quadratic Discriminant Analysis; NBDT: Naïve Bayes Decision Trees; TRUS: Trans Rectal Ultrasonogram; SWE: Shear-Wave Elastography; DCE-US: Dynamic Contrast-Enhanced Ultrasound; PLCO: Prostate Lung Colorectal Ovarian; OS: Overall Survival

Gavriel et al.^[37] proposed an ensemble system comprising ML-based algorithms to predict five-year prognosis with different combinations of image, clinical, and spatial features and quantify potential prognostic markers related to lymphocytes, macrophages, tumor buds, and PD-L1. The method successfully classified 71.4% of the patients who succumbed to muscle invasive bladder cancer (MIBC) within five years, significantly higher than the 28.6% of the current clinical gold standard, the tumour, node, metastasis (TNM) staging system.

Several studies have applied ML-based algorithms and models to identify genes that could predict the recurrence of disease or the future progression. Slides of patients diagnosed with MIBC were labeled with immunofluorescence (IF) and used for measuring the tumor buds, to determine the effectiveness of neoadjuvant chemotherapy, and to identify patients who were not responding to the treatment. This was done to stop the treatment prematurely to avoid the adverse effects of chemotherapy.^[38-40]

Prostate Carcinoma

AI applications are on the verge of revolutionizing the current practice in carcinoma prostate (CaP) in terms of diagnosis, treatment decisions, and even predicting the disease-free survival. There is high observer-dependent variability in Gleason grading because of the subjective nature of the analysis of biopsy specimens. Considering this, Ström et al.^[41] developed an AI model for identification, Gleason grading, and localization of prostate cancer. The model was trained with 6682 digitized slides of 976 men and tested on 1631 biopsy specimens from 246 men. It achieved an accuracy of 0.997 (AUC) to differentiate between a malignant and a benign tumor. The results in terms of Gleason grading were also comparable to those achieved by the expert pathologists. In various studies, DL methods to calculate the Gleason Grading have been applied (Table 2).^[42]

Multiparametric imaging uses multiple modalities or techniques before making the ultimate diagnosis, and this adds to the burden of the radiologist. However, in the current era, computer-aided diagnosis is possible because of progress in AI, which eventually helps in making the diagnosis by image interpretation. This is particularly useful in situations where multiple modalities, parameters, or techniques are involved in diagnosing a condition.^[43]

Application of a Quadrant Discriminant Classifier to the radiomic features derived from T2-weighted MRI images for detection of CaP^[44] and application of ML-based random Forrest classification algorithm to localize CaP on transrectal ultrasonogram^[45] have been studied (Table 2).

In view of the toxic effects of docetaxel chemotherapy, 20% of the patients undergo therapeutic failure in metastatic castrate resistant CaP. Deng et al.^[46] developed an AI-based computational model that could differentiate patients in two groups, docetaxel-tolerable and docetaxel-intolerable, for better and individualized treatment for the patients in this category. Identification of the presence of biomarkers on tissue microarrays can predict the risk of recurrence and metastasis. Biomarker identification under IF microscope by the human eye is subjective as well as time-consuming. Hence, an automated method using DL algorithms was developed for analysis of biomarkers using 648 samples and IF staining with anti-Ki-67, ERG antibodies. The results were promising, with only 5% difference between manual and algorithm-based biomarker detection and 100% accuracy in identification of tumors positive for ERG.^[47]

Bibault et al.^[48] used data from the prospective clinical trial Prostate Lung Colorectal and Ovarian cancer screening, selected patients who were diagnosed with CaP during follow-up, and trained two models to predict ten-year cancer-specific survival (CSS) and OS. Of the 8776 patients diagnosed with PCa on follow-up, training of the models was done with 7021 and tested on dataset of 1755. It achieved an accuracy of 0.87 and 0.98 for OS and CSS, respectively. These models can be used online to provide predictions and support informed decision-making in CaP treatment.

Limitations

AI applications are gaining significant interest in urology, but their real-world implementation still faces an uphill task. There are limitations to some studies that use AI algorithms and its subsets in urological diseases. The key challenges that can be addressed before being integrated into the clinical setting are the incorporation of standardized criteria, the correction for system variation, and the data collection from multiple institutions in various geographical locations, so that the results can be generalized and applied to the real-world scenario.^[49]

Future Considerations

The President of the World Economic Forum, Klaus Schwabe, made the following announcements at the Davos Summit just a few years ago: "We stand on the brink of a technological revolution that will fundamentally alter the way we live, work, and relate to one another. In its scale, scope, and complexity, the transformation will be unlike anything human kind has experienced before".^[50]

Future work will concentrate on creating larger medical databases and expanding AI techniques further. The use of enhanced algorithms will take place on smartphones or can be accessed through the cloud. Applications for clinical decision-making and its use in the real world require appropriate permissions from the regulatory bodies. Issues exist concerning the reliability of a machine diagnosis and that prejudices of programming do not create hindrances in the diagnosis.

Conclusion

In the near future, we will see a shift in the clinical paradigm as AI applications will find their place in the guidelines and revolutionize the decision-making process. Having said that, human qualities of intelligence, adaptation, and sense of duty will prove to be important factors in further development of AI.

Peer-review: This manuscript was prepared by the invitation of the Editorial Board and its scientific evaluation was carried out by the Editorial Board.

Author Contributions: Concept - M.S., N.N., B.M.Z.H.; Design - N.N., B.M.Z.H.; Supervision - B.K.S.; Materials - M.S.; Data Collection and/or Processing - M.S.; Analysis and/or Interpretation - N.N., B.M.Z.H.; Literature Search - M.S., N.N., B.M.Z.H.; Writing Manuscript - M.S., N.N., B.M.Z.H.; Critical Review - B.K.S.

Conflict of Interest: The authors have no conflicts of interest to declare.

Financial Disclosure: The authors declared that this study has received no financial support.

References

- Beam A, Kohane I. Big Data and Machine Learning in Health Care. JAMA 2018;319:1317. [Crossref]
- Grand View Research. Healthcare Predictive Analytics Market Analysis by Application (Operations Management, Financial, Population Health, Clinical), By End-Use (Payers, Providers), By Region (North America, Europe, Asia Pacific, Latin America, MEA) and Segment Forecasts, 2018-2025. 2016 November 2016.
- Drouin S, Yates D, Hupertan V, Cussenot O, Rouprêt M. A systematic review of the tools available for predicting survival and managing patients with urothelial carcinomas of the bladder and of the upper tract in a curative setting. World J Urol 2012;31:109-16. [Crossref]
- Parakh A, Lee H, Lee J, Eisner BH, Sahani DV, Do S. Urinary Stone Detection on CT Images Using Deep Convolutional Neural Networks: Evaluation of Model Performance and Generalization. Radiology: Artificial Intelligence 2019;1:e180066. [Crossref]
- Chiang D, Chiang H, Chen WC, Tsai FJ. Prediction of stone disease by discriminant analysis and artificial neural networks in genetic polymorphisms: a new method. BJU Int 2003;91:661-6.
 [Crossref]
- Kazemi Y, Mirroshandel S. A novel method for predicting kidney stone type using ensemble learning. Artif Intell Med 2018;84:117-26. [Crossref]
- Kriegshauser J, Silva A, Paden R, He M, Humphreys MR, Zell SI, et al. Ex Vivo Renal Stone Characterization with Single-Source Dual-Energy Computed Tomography: A Multiparametric Approach. Acad Radiol 2016;23:969-76. [Crossref]
- Eken C, Bilge U, Kartal M, Eray O. Artificial neural network, genetic algorithm, and logistic regression applications for predicting renal colic in emergency settings. Int J Emerg Med 2009;2:99-105. [Crossref]
- Dal Moro F, Abate A, Lanckriet G, Arandjelovic G, Gasparella P, Bassi P, et al. A novel approach for accurate prediction of spontaneous passage of ureteral stones: Support vector machines. Kidney Int 2006;69:157-60. [Crossref]
- Shabaniyan T, Parsaei H, Aminsharifi A, Movahedi MM, Jahromi AT, Pouyesh S, et al. An artificial intelligence-based clinical decision support system for large kidney stone treatment. Australas Phys Eng Sci Med 2019;42:771-9. [Crossref]
- Aminsharifi A, Irani D, Tayebi S, Kafash TJ, Shabanian T, Parsaei H. Predicting the Postoperative Outcome of Percutaneous Nephrolithotomy with Machine Learning System: Software Validation and Comparative Analysis with Guy's Stone Score and the CROES Nomogram. J Endourol 2020;DOI: 10.1089/end.2019.0475. [Crossref]
- Kadlec A, Ohlander S, Hotaling J, Hannick J, Niederberger C, Turk TM. Nonlinear logistic regression model for outcomes after endourologic procedures: a novel predictor. Urolithiasis 2014;42:323-7.
 [Crossref]

- Seckiner I, Seckiner S, Sen H, Bayrak O, Dogan K, Erturhan S. A neural network - based algorithm for predicting stone -free status after ESWL therapy. Int Braz J Urol 2017;43:1110-4. [Crossref]
- Torshizi A, Zarandi M, Torshizi G, Eghbali K. A hybrid fuzzyontology based intelligent system to determine level of severity and treatment recommendation for Benign Prostatic Hyperplasia. Comput Methods Programs Biomed 2014;113:301-13. [Crossref]
- Sonke GS, Heskes T, Verbeek AL, de la Rosette JJ, Kiemeney LA. Prediction of bladder outlet obstruction in men with lower urinary tract symptoms using artificial neural networks. J Urol 2000;163:300-5. [Crossref]
- Bagli D, Agarwal S, Venkateswaran S, Shuckett B, Khoury AE, Merguerian PA, et al. Artificial Neural Networks In Pediatric Urology: Prediction Of Sonographic Outcome Following Pyeloplasty. J Urol 1998;160:980-3. [Crossref]
- Logvinenko T, Chow J, Nelson C. Predictive value of specific ultrasound findings when used as a screening test for abnormalities on VCUG. J Pediatr Urol 2015;11:176.e1-176.e7. [Crossref]
- Blum ES, Porras AR, Biggs E, Tabrizi PR, Sussman RD, Sprague BM, et al. Early Detection of ureteropelvic junction obstruction using signal analysis and machine learning: A dynamic solution to a dynamic problem. J Urol 2018;199:847-52. [Crossref]
- Bentaleb J, Larouche M. Innovative use of artificial intelligence in urogynecology. Int Urogynecol J 2020; DOI: 10.1007/s00192-020-04243-2. [Crossref]
- Sabadell J, Salicrú S, Montero-Armengol A, Rodriguez-Mias N, Gil-Moreno A, Poza JL. External validation of de novo stress urinary incontinence prediction model after vaginal prolapse surgery. Int Urogynecol J 2019;30:1719-23. [Crossref]
- Jelovsek JE, Chagin K, Lukacz ES, Nolen TL, Shepherd JP, Barber MD, et al. Models for predicting recurrence, complications, and health status in women after pelvic organ prolapse surgery. Obstet Gynecol 2018;132:298-309. [Crossref]
- Atallah DM, Badawy M, El-Sayed A, Ghoneim MA. Predicting kidney transplantation outcome based on hybrid feature selection and KNN classifier. Multimed Tools Appl 2019;78:20383-407.
 [Crossref]
- Greco R, Papalia T, Lofaro D, Maestripieri S, Mancuso D, Bonofiglio R. Decisional Trees in Renal Transplant Follow-up. Transplant Proc 2010;42:1134-6. [Crossref]
- Goldfarb-Rumyantzev AS, Scandling JD, Pappas L, Smout RJ, Horn S. Prediction of 3-yr cadaveric graft survival based on pre-transplant variables in a large national dataset. Clin Transpl 2003;17:485-97. [Crossref]
- 25. Baessler B, Nestler T, Pinto dos Santos D, Paffenholz P, Zeuch V, Pfister D, et al. Radiomics allows for detection of benign and malignant histopathology in patients with metastatic testicular germ cell tumors prior to post-chemotherapy retroperitoneal lymph node dissection. Eur Radiol 2020;30:2334-45. [Crossref]
- Kocak B, Yardimci A, Bektas C, Turkcanoglu MH, Erdim C, Yucetas U, et al. Textural differences between renal cell carcinoma subtypes: Machine learning-based quantitative computed tomography texture analysis with independent external validation. Eur J Radiol 2018;107:149-57. [Crossref]
- 27. Feng Z, Rong P, Cao P, Zhou Q, Zhu W, Yan Z, et al. Machine learning-based quantitative texture analysis of CT images of small

renal masses: Differentiation of angiomyolipoma without visible fat from renal cell carcinoma. Eur Radiol 2017;28:1625-33. [Crossref]

- Coy H, Hsieh K, Wu W, Nagarajan MB, Young JR, Douek ML, et al. Deep learning and radiomics: the utility of Google Tensor-Flow[™] Inception in classifying clear cell renal cell carcinoma and oncocytoma on multiphasic CT. Abdom Radiol 2019;44:2009-20. [Crossref]
- Ding J, Xing Z, Jiang Z, Chen J, Pan L, Qiu J, et al. CT-based radiomic model predicts high grade of clear cell renal cell carcinoma. Eur J Radiol 2018;103:51-6. [Crossref]
- Li P, Ren H, Zhang Y, Zhou Z. Fifteen-gene expression based model predicts the survival of clear cell renal cell carcinoma. Medicine 2018;97:e11839. [Crossref]
- Kocak B, Durmaz E, Ates E, Baykara Ulusan M. Radiogenomics in Clear Cell Renal Cell Carcinoma: Machine Learning-Based High-Dimensional Quantitative CT Texture Analysis in Predicting PBRM1 Mutation Status. AJR Am J Roentgenol 2019;212:W55-W63. [Crossref]
- Ikeda A, Nosato H, Kochi Y, Kojima T, Kawai K, Sakanashi H, et al. Support System of Cystoscopic Diagnosis for Bladder Cancer Based on Artificial Intelligence. J Endourol 2020;34:352-8.
 [Crossref]
- Lorencin I, Anđelić N, Španjol J, Car Z. Using multi-layer perceptron with Laplacian edge detector for bladder cancer diagnosis. Artif Intell Med 2020;102:101746. [Crossref]
- Hashemia SMR, Hassanpourb H, Kozegarc E. Cystoscopic Image Classification Based on Combining MLP and GA. Int J Nonlinear Analys App 2020;11:93-105.
- Eminaga O, Semjonow A, Breil B. Diagnostic classification of cystoscopic images using deep convolutional neural networks. Eur Urol Supp 2018;17:e1232. [Crossref]
- Wang G, Zhang G, Choi K, Lam KM, Lu J. Output based transfer learning with least squares support vector machine and its application in bladder cancer prognosis. Neurocomputing 2020;387:279-92. [Crossref]
- Gavriel C, Dimitriou N, Brieu N, Nearchou IP, Arandjelovic O, Schmidt G, et al. Identification of Immunological Features Enables Survival Prediction of Muscle-Invasive Bladder Cancer Patients Using Machine Learning. BioRxiv 2020. DO: 10.1101/2020.02.24.963181 [Crossref]
- Hasnain Z, Mason J, Gill K, Miranda G, Gill IS, Kuhn P, et al. Machine learning models for predicting post-cystectomy recurrence and survival in bladder cancer patients. PLoS One 2019;14:e0210976. [Crossref]

- Bartsch G, Mitra A, Mitra S, Almal AA, Steven KE, Skinner DS, et al. Use of Artificial Intelligence and Machine Learning Algorithms with Gene Expression Profiling to Predict Recurrent Nonmuscle Invasive Urothelial Carcinoma of the Bladder. J Urol 2016;195:493-8. [Crossref]
- Wu E, Hadjiiski LM, Samala RK, Chan HP, Cha KH, Richter C, et al. Deep Learning Approach for Assessment of Bladder Cancer Treatment Response. Tomography 2019;5:201-8. [Crossref]
- Ström P, Kartasalo K, Olsson H, Solorzano L, Delahunt B, Berney DM, et al. Artificial intelligence for diagnosis and grading of prostate cancer in biopsies: a population-based, diagnostic study. Lancet Oncol 2020;21:222-32. [Crossref]
- Bulten W, Pinckaers H, van Boven H, Vink R, de Bel T, van Ginneken B, et al. Automated deep-learning system for Gleason grading of prostate cancer using biopsies: a diagnostic study. Lancet Oncol 2020;21:233-41. [Crossref]
- Wildeboer R, van Sloun R, Wijkstra H, Mischi M. Artificial intelligence in multiparametric prostate cancer imaging with focus on deep-learning methods. Comput Methods Programs Biomed 2020;189:105316. [Crossref]
- 44. Viswanath S, Chirra P, Yim M, Rofsky NM, Purysko AS, Rosen MA, et al. Comparing radiomic classifiers and classifier ensembles for detection of peripheral zone prostate tumors on T2-weighted MRI: a multi-site study. BMC Med Imaging 2019;19:22. [Crossref]
- 45. Wildeboer R, Mannaerts C, van Sloun R, Budäus L, Tilki D, Wijkstra H, et al. Automated multiparametric localization of prostate cancer based on B-mode, shear-wave elastography, and contrastenhanced ultrasound radiomics. Eur Radiol 2019;30:806-15. [Crossref]
- Deng K, Li H, Guan Y. Treatment Stratification of Patients with Metastatic Castration-Resistant Prostate Cancer by Machine Learning. iScience 2020;23:100804. [Crossref]
- de la Calle CM, Nguyen H, Hosseini-Asl E, So C, Socher R, Xiong C, et al. Artificial intelligence for streamlined immunofluorescence-based biomarker discovery in prostate cancer. J Clin Oncol 2020;38(Suppl 6):279. [Crossref]
- Bibault JE, Xing L. Predicting Survival in Prostate Cancer Patients with Interpretable Artificial Intelligence. Available at SSRN 3546050. 2020. [Crossref]
- Chen J, Remulla D, Nguyen J, Aastha D, Liu Y, Dasgupta P, et al. Current status of artificial intelligence applications in urology and their potential to influence clinical practice. BJU Int 2019;124:567-77. [Crossref]
- Schwab, K. The Fourth Industrial Revolution; World Economic Forum: Geneva, Switzerland, 2016; ISBN 9781944835002.