A systematic review of clinical value of three-dimensional printing in renal disease

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Abstract: The aim of this systematic review is to analyse current literature related to the clinical value of three-dimensional (3D) printed models in renal disease. A literature search of PubMed and Scopus databases was performed to identify studies reporting the clinical application and usefulness of 3D printed models in renal disease. Fifteen studies were found to meet the selection criteria and were included in the analysis. Eight of them provided quantitative assessments with five studies focusing on dimensional accuracy of 3D printed models in replicating renal anatomy and tumour, and on measuring tumour volume between 3D printed models and original source images and surgical specimens, with mean difference less than 10%. The other three studies reported that the use of 3D printed models significantly enhanced medical students and specialists' ability to identify anatomical structures when compared to two-dimensional (2D) images alone; and significantly shortened intraoperative ultrasound duration compared to without use of 3D printed models. Seven studies provided qualitative assessments of the usefulness of 3D printed kidney models with findings showing that 3D printed models improved patient's understanding of renal anatomy and pathology; improved medical trainees' understanding of renal malignant tumours when compared to viewing medical images alone; and assisted surgical planning and simulation of renal surgical procedures with significant reductions of intraoperative complications. The cost and time associated with 3D printed kidney model production was reported in 10 studies, with costs ranging from USD\$100 to USD\$1,000, and duration of 3D printing production up to 31 h. The entire process of 3D printing could take up to a few days. This review shows that 3D printed kidney models are accurate in delineating renal anatomical structures and renal tumours with high accuracy. Patient-specific 3D printed models serve as a useful tool in preoperative planning and simulation of surgical procedures for treatment of renal tumours. Further studies with inclusion of more cases and with a focus on reducing the cost and 3D model production time deserve to be investigated.

Keywords: Model; renal tumour; simulation; surgical planning; three-dimensional printing (3D printing); threedimensional visualization (3D visualization)

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Introduction

Three-dimensional (3D) printing has been increasingly used in medicine with reports showing great promise in many areas ranging from applications in orthopaedics and maxillofacial reconstruction to cardiovascular and liver diseases (1-10). Patient-specific 3D printed models based on computed tomography (CT) or magnetic resonance imaging (MRI) data have been shown to accurately replicate complex anatomical structures and pathologies when compared to original source images (2-4,7). 3D printed models can also be used to assist pre-surgical planning and simulation, improve understanding of anatomy and individual lesions (8-10).

Although two-dimensional (2D) and 3D images are routinely used to diagnose and assess the renal tumours in relation to the surrounding structures prior to surgery (*Figure 1*), the complex relationships between the renal tumours and different renal structures cannot be fully appreciated on images alone. Further, treatment of renal lesions is moving from traditional open surgery to minimally invasive approach (11-14), thus, it is necessary for surgeons and trainees to obtain a tactile experience of renal tumour or renal system so that better surgical plans or treatment strategies can be determined. Use of 3D printed kidney models in clinical practice fulfils this goal.

Several studies demonstrate the application of 3D printed models in pre-surgical planning of complex renal tumours and renal transplantation (15-18). Despite promising results reported in these studies, there is no systematic review of the current literature about the accuracy of 3D printed models and their clinical applications in renal disease. Thus, the purpose of this review is to analyse current literature with regard to the clinical value of 3D printed models in renal disease, with a focus on application of 3D printed models in renal tumours. Types of 3D printers used for 3D printing, software tools for image post-processing and segmentation, and duration of 3D printing as well as cost associated with 3D printing are also analysed in the review.

Methods

Search strategy

This review was performed to comply with the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) guidelines (19). A search of two main databases PubMed and Scopus was conducted to identify studies reporting the usefulness of 3D printed models in renal

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disease. The following keywords were used to perform the search: renal tumours OR renal disease AND 3D printing; renal tumours or renal disease AND 3D printed models; renal tumours OR renal disease AND rapid prototyping; renal tumours OR renal disease AND additive manufacturing; renal tumours OR renal disease AND 3D simulations. The research was limited to peer-reviewed studies published in English within the last 10 years (last search: March, 2018) to ensure the relevance of these studies to the current practice in 3D printing.

Since research on 3D printing in renal disease is at its infancy and the number of studies is limited in the literature, both case reports and original research studies were included in this review to allow systematic analysis of the current research in this area.

Eligibility criteria and data extraction

Two assessors conducted the search of the studies by screening the title and abstract of all identified references independently. Eligible studies must meet the following criteria: either prospective or retrospective studies (case reports, case series or original studies) on patient-specific 3D printed kidney models reporting the clinical value or usefulness of 3D printing in renal anatomy or renal disease. Based on these criteria, review articles, phantom studies, editorials and conference abstracts were excluded. When multiple studies were published by the same research group, details of the study design and findings were checked and duplicate reports were excluded with the most recent ones included in the analysis.

Full texts of the eligible studies were then reviewed and assessed by two assessors independently with agreement reached during discussion. Data extraction includes: accuracy of 3D printed models in delineating renal anatomy and renal tumour, clinical value and usefulness of 3D printed kidney models in pre-surgical planning and simulation of renal procedures; and 3D printed kidney models in patient or medical education. Further, duration of image post-processing and segmentation, software tools used for image processing, types of 3D printers, 3D printing materials and associated costs, and duration of 3D printing were also analysed in the review.

Results

Literature search outcome

The initial search retrieved sixty-one articles. Forty articles



Figure 1 Contrast-enhanced CT images with 2D and 3D reconstructions showing malignant renal tumour. (A) 2D axial CT image shows a tumour with contrast enhancement at the lower and posterior region of left kidney (arrow); (B) coronal reformatted view shows the tumour's enhancement is heterogeneous (arrows), with low-attenuation areas within the lesion; (C) 3D volume rendering frontal view shows the tumour is located at the posterior aspect of left kidney (arrow); (D) 3D volume rendering posterior view shows the tumour location (arrows). CT, computed tomography; 2D, two-dimensional; 3D, three-dimensional.

were excluded due to their irrelevant to the topic. Of 21 full-text articles that were selected, two were excluded with one being review article and another one technical report. Of 19 eligible studies, 4 were further excluded because of duplicate publication in 2 studies by the same research group (20,21), while another 2 studies focused on 3D printed renal compartments for nuclear medicine imaging without reporting any clinical application of 3D printed models in renal anatomy or renal disease (22,23). Thus, a total of 15 articles were finally included in this review (15-18,24-34). *Figure 2* shows the search strategy to identify these eligible studies.

Table 1 is a summary of study characteristics of the use of 3D printed models in renal disease. Of 15 studies, isolated

case reports with creation of 1–3 patient-specific 3D printed models were seen in 6 studies, while generation of 5 to 10 3D printed models was reported in 9 studies.

CT is the most commonly used imaging modality for segmentation of anatomical structures, with 12 studies using CT data as the source of imaging data for 3D printing. MRI data were used in two studies, while in the remaining study, CT or MRI images were used for creating 3D printed models (*Table 1*).

Quantitative assessment of 3D printed kidney models

Of 15 studies, 8 (53%) provided quantitative assessments of 3D printed kidney models with reported applications



Figure 2 Flow chart shows the searching strategy to identify studies that are included in the analysis.

in different aspects (16,18,24,29,30,32-34). Comparison of dimensional accuracy or renal tumour volume between 3D printed models and original imaging data/or surgical results was reported in 5 studies (18,24,32-34). Results of these studies showed that 3D printed models are highly accurate in replicating anatomical structures or detecting tumour volumes with mean difference <10% in dimensional measurements between 3D printed models and original imaging data or surgically resected specimens (*Table 1*).

The other 3 studies reported quantitative assessments of 3D printed models in different perspectives (16,29,30). This includes the following findings: significant reduction in intraoperative ultrasound duration with use of 3D printed models when compared to that without using models (16); medical trainees' accuracy in nephrectomy score was significantly improved with reduced variability with assistance from 3D printed models in comparison with CT scans (29); 3D printed models assisted medical students, surgeons and radiologists to identify anatomical structures with highest percentage of correct answers, and shorter time spent on recognising anatomy when compared to 3D virtual reconstructions and 2D CT images (30).

Figures 3-5 are examples of showing patient-specific 3D printed kidney models based on CT or MRI data with excellent demonstration of renal anatomical structures and renal tumours.

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Qualitative assessment of 3D printed kidney models

There are 7 studies reporting the qualitative assessment of 3D printed models in renal disease (15,17,25-28,31). Three of them showed that patient's understanding of renal anatomy and pathology was improved with use of 3D printed models, with patient's satisfaction score more than 9 out of 10 (15,25,31). Another three studies demonstrated the clinical value of using 3D printed models in pre-operative planning or simulation of surgical procedures, which led to reduction of surgery duration and intraoperative complications, as well as assisting renal transplantation surgery (17,27,28). The remaining study by Dwivedi *et al.* presented their first report of correlating 3D printed renal molds with MRI tumour features, with the aim of providing opportunities for future studies of radiomic and radiogenomic analysis of renal tumours (26).

Cost and time associated with 3D printing in kidney models

Different software tools were used in performing image post-processing and segmentation of CT or MRI data, with Mimics and 3D Slicer being the most commonly used software, while details of software packages were not reported in two studies (*Table 1*). The time spent on image processing and segmentation was only reported in 2 studies as shown in the Table (18,33).

The cost of 3D printing is reported in 10 studies, ranging from USD\$100 to USD\$1,000, and this is highly dependent on the materials and the type of 3D printers used for 3D printing. In a recent study, Liu *et al.* compared the cost and accuracy of 3D printed kidney models using commercial 3D printer with those from home-made 3D printer (33). Authors reported that patient-specific 3D printed models with homemade 3D printer were accurate in demonstrating renal anatomical structures and renal tumours when compared to those from the commercial 3D printer, but at much lower cost (USD\$1 *vs.* USD\$ 200 per model).

This review shows a variety of different 3D printers used in printing 3D models, with Objet Connex (Stratasys) as the most common one, which was used in 5 studies. Time taken for 3D printing was available in 10 studies, ranging from 2 to 31 h. The entire process or duration of 3D printing ranged from 2 to 9 days according to some studies in this review (16,24,34).

Discussion

This systematic review analyses 15 studies with regard

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Table 1 Study characteristics of 3D printing in renal disease

12-14 h for printing a mold

with vith of	Study purpose 3D printing to	Imaging modalities used for 3D printing CT	Software used for image segmentation of segmentation NA	3D printer/printing materials/ costs/printing duration Fused Deposition Modelling	Key findings
olan for a rai somplicated surgery	Φ			Euro 100 (USD\$123) 22 h	 planning and identification of anatomical structures, thus improving safety and reducing the surgery duration Treatment decisions can be expedited with use of 3D physical models to facilitate communication between physicians from different disciplines
8D printed kidney model or training in aparoscopic procedures	S	Ъ	3D Slicer (Surgical Planning Lab, Harvard University, MA, USA) TinkerCAD (Autodesk, San Rafal, CA, USA)	MakerBot Replicator 2 (MakerBot Industries, LCC) Silicone material Elite Double 8 USD\$120 Printing kidney and turnours: 4 h and 15 min, 55 min, respectively Total time of production: 7–8 h	Simulation of surgical procedures based on 3D printed models improves actual laparoscopic surgery and shortens the duration of intraoperative renal ischemia
Juantitative and qualitative assessments is sessments a patient-specif 8D printed models on medical traine anderstanding enal tumours	e ic j of	с	N/A	Stereolithography (SLA) Translucent Resin	 3D printed models improved medical trainees' ratings (accuracy) with reduced variability when compared to CT scans with strong correlation using 3D model vs using the CT scans alone (ICC: 0.719 vs. 0.284) 3D printed models improved medical trainees' understanding of renal malignancy compared to CT (P<0.01)
HD navigation n minimally nvasive surgi procedures hrough 3D printed mode	ls cal	cJ	ZedView (LEXI Co, Ltd, Tokyo, Japan)/ Geomagic (Freeform, Geomagic, Rock Hill, SC)	Connex 500 (Stratasys Ltd., MN,USA)	 3D printed models assist surgeons to perform minimally invasive procedure of partial nephrectomy by significantly shortening intraoperative ultrasound duration (mean 3.3 vs. 6.3 min with and without using 3D model, P=0.021)
				Acrylic resin USD\$450-680 3D printing: 4–9 h Model completion: 3–9 davs	 3D printed models improve patient's understanding of their disease status when compared to viewing CT images alone
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Table 1 (continued)

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First author and year of publication	Study sample size	Study purpose	Imaging modalities used for 3D printing	Software used for image segmentation/duration of segmentation	3D printer/printing materials/ costs/printing duration	Key findings
Kusaka <i>et al.</i> 2015 (17)	Case report with 1 case for donor graft kidney model	Clinical application of 3D kidney graft models and	ст	OsiriX (Pixmeo, Geneva, Switzerland)	Connex 500 (Stratasys Ltd., MN, USA)	 3D printed life-size models improve understanding of relationship among anatomical structures and may reduce intraoperative complications
	and 1 case for donor graft kidney and recipient pelvic cavity	pelvic cavity replicas for pre-surgical simulation in living kidney transplantation		Magics (Materialise, Leuven, Belgium)	VeroClear and TangoPlus for printing renal parenchyma, TangoPlus/VeroMagenta blend, TangoPlus/VeroMagenta/ VeroCyan blend for renal artery, renal artery vein and urinary tract structures, respectively	 3D printed models show beneficial roles in kidney transplantation surgery
Marconi et <i>al</i> 2017 (30)	. 15 cases scheduled for minimally invasive surgery with renal tumours	Objective assessment of 3D printed models in preoperative planning and surgical	cī	Paraview (http://www. paraview.org	USD\$200-240	 3D printed models led to highest percentage of correct answers in identifying anatomical structures (53.9%±4.6) compared to 3D virtual reconstructions (53.4%±4.6) and 2D CT images (45.5%±4.6)
	in 2 cases	training of renal disease by 30 participants (10 medical students, 10 general surgeons and 10 radiologists)			20-30 h	 Participants spent significantly shorter time on evaluating 3D printed models (60.67±25.5 s) than on 2D CT images (127.04±35.91 s), and shorter than on 3D virtual reconstruction images (70.8±28.18 s)
Porpiglia <i>et al.</i> 2018 (31)	18 cases with 8 undergoing robot-assisted radical prostatectomy	Usefulness of 3D printed models in pre-surgical planning, training and education	CT for renal tumours MRI for prostate cancer.	M3DISC	Selective laser sintering	 Overall median usefulness of 3D printed models and benefits compared to 3D virtual models as assessed by urologists and patients was ≥8 out of 10
	and 10 minimally invasive partial				Powder	 Accuracy of 3D printed models in replicating anatomical details was 10 out of 10
						 Improving understanding of surgical procedure and surgical planning was 8 out of 10
						 Patient's perception about usefulness of 3D printed models was 10 out of 10

Table 1 (cont.	inued)					
First author and year of publication	Study sample size	Study purpose	Imaging modalities used for 3D printing	Software used for image segmentation/duration of segmentation	3D printer/printing materials/ costs/printing duration	Key findings
von Rundstedt <i>et al.</i> 2017 (32)	10 cases with renal tumours	Development of a novel protocol using 3D printed models for pre- surgical planning	CT or MRI	3D Slicer (https://www.slicer.org)	Lazarus 3D (Houston, TX, USA)	 No significance difference was found in total turmour volume between original 3D CT (38.88 mm³), resected pre-surgical model (38.50 mm³) and resected turmour specimen (41.79 mm³) (P=0.98)
		and simulation of robot-assisted laparoscopic partial nephrectomy			Silicone rubber	 No significance difference was found in resection times between 3D printed models and actual surgery (6.61 vs. 7.93 min, P=0.162)
						 Patient-specific 3D printed models assist surgical training and decision- making in patients scheduled for partial nephrectomy
Liu <i>et al.</i> 2018 (33)	2 cases of renal tumours	Comparison of 3D printed model accuracy	ट	Analyze 12.0 (AnalyzeDirect, Inc., Lexana, KS, USA)	Object 260 (Stratesys, EDEN 260 VS) vs. Home-made 3D printer	 Measurements on turmour diameters between commercial and home-made 3D printers were less than 1.0 mm
		between home- made 3D printer and commercial one		1.5 h	TangoPlus vs. Polylactide	 Good correlation between original 3D CT images and 3D printed models in measurements of tumour diameters (differences <1.0 mm)
					3.5 vs .4–5 h for each kidney USD\$200 vs. USD\$1	 Feasibility of producing 3D printed models with use of home-made 3D printer with similar accuracy but much lower cost than the commercial one
Wake <i>et al.</i> 2017 (18)	10 cases with renal tumours	Impact of 3D printed renal tumour models on surgical	MRI	Mimics 16.0 (Materialise, Belgium)	Connex 500 (Stratasys, MN, USA)	 Good correlation in tumour measurement between 2D images and 3D printed models with mean difference <1.5% (r=0.388, P<0.001)
		planning of renal cancer		7 h per kidney model	VeroCyan and VeroMagenta USD\$1,000	 Use of 3D printed models led to a change in planned approach in all cases with greatest impact noted on transperitoneal or retroperitoneal approach and clamping, with changed approaches in 30–50% cases
					10 h to print each model	 3D printed models improved understanding of anatomy, decisions on surgical approach and surgeons' confidence in procedure planning
Table 1 (cont.	inued)					

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First author and year of publication	Study sample size	Study purpose	Imaging modalities used for 3D printing	Software used for image segmentation/duration of segmentation	3D printer/printing materials/ costs/printing duration	Key findings
Zhang <i>et al.</i> 2016 (34)	10 cases with renal tumours	Impact of 3D printed models on surgical planning, training and patient's understanding of renal tumour surgery	ت	Medical Imaging TookKit (MITK) and 3DMed	LaserCore 5300 (Longyuan Rapid Prototyping Ltd., Beijing, China) Thermoplastic plastics USD\$150 3-4 days	 Overall usefulness of 3D printed models, assistance in surgical planning and training, and realism were rated with mean scores: 7.8, 8.0 and 6.0–7.8 out of 10, respectively Patient's overall satisfaction of the 3D models was ≥9.0 out of 10 Tumour diameter was measured on original CT images, 3D printed models and actual specimens: 39.3±11.4, 34.5±13.0, and 37.9±11.0 mm, with size deviation of 3.4±1.3 mm between 3D model and specimen
CT, compute	id tomography; IC	C, intraclass correla	tion coefficient; M	IRI, magnetic resonance im	aging; N/A, not available.	

to the clinical value of 3D printed kidney models, with a summary of the following key findings based on the current literature: first, 3D printed kidney models can be successfully generated with use of CT or MRI data, with high accuracy in reproducing renal anatomy and pathology according to quantitative assessments reported by some studies in the review. Second, 3D printed kidney models could serve as a useful tool in preoperative planning and simulation of complex surgical procedures, thus, reducing the operative duration and potential risks or complications associated with the surgery. Last, 3D printed kidney models improve medical trainees' understanding of complex renal anatomy and pathology, and patients' understanding of their disease condition. 3D printed models are also found useful by patients to assist doctor-patient discussion or consultation of clinical cases.

2D and 3D CT or MRI image visualizations are commonly used in the diagnosis of renal tumours, however, to fully understand the relationship between renal tumours and surrounding renal anatomical structures could be a challenging task in some cases. 3D printed physical models are reported to show increasing value in clinical practice which is demonstrated in different medical areas, ranging from accurate delineation of anatomical structures and pathologies to improved understanding of complex anatomy and pathology, assistance in preoperative planning and simulation of surgical procedures and medical education (4-10). Some recent systematic reviews have shown the value of 3D printing in cardiovascular and liver diseases. A systematic review of 48 studies has summarised the findings of using 3D printed models in cardiovascular and cerebrovascular diseases, with high accuracy of 3D printing in reproducing complex cardiac anatomy and pathologies, and usefulness in both education and surgical planning (8). Another recent systematic review of 28 studies focuses on the 3D printing in congenital heart disease (10). Through analysis of case reports and original studies the review further confirms the clinical value of patient-specific 3D printed heart models in preoperative planning and simulation of congenital heart disease treatment and improving medical education and doctor-patient communication.

Two systematic reviews on 3D printing in liver disease are available in the current literature with analysis of 6 and 14 studies, indicating the emerging area of using 3D printing in this field (35,36). Studies reporting the application of 3D printed models in renal disease are similar to those in liver disease. This is confirmed by this review with inclusion of 15 eligible studies. To the best of our

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Figure 3 Use of contrast-enhanced MR images for generating 3D printed model of renal tumour. (A) Axial, coronal, and sagittal views of MRI images with segmentation masks for one representative case. Kidney = teal, tumour = pink, artery = red, vein = blue, collecting system = green; (B) anterior and posterior 3D projections. Kidney = gray, tumour = pink, artery = red, vein = blue, ureter = green; (C) photographs of 3D printed model. Kidney = transparent, tumour = purple, artery = pink, vein = light blue, ureter = dark blue. Reprinted with permission from Wake *et al.* (18). MRI, magnetic resonance imaging; 3D, three-dimensional.

knowledge, this is the first systematic review of the clinical application of 3D printed kidney models. Despite limited number of studies reporting quantitative and qualitative assessments of 3D printing in renal disease, this review presents encouraging findings of 3D printed kidney models in demonstrating excellent 3D relationship between renal tumours and adjacent anatomical structures, which plays an important role in surgical planning and simulation, although further studies with inclusion of more cases are needed to confirm these results.

There is no doubt that 3D printed models serve as valuable tools in many applications, however there are

challenging issues related to 3D printing such as cost and time of production that need to be resolved. Image postprocessing and segmentation of medical imaging data could be time-consuming, especially when dealing with 3D printed heart or liver models due to complex structures in these regions. Further, a number of editing processes are applied to ensure successful 3D printing outcomes (10). For 3D printed heart and liver models, the time spent on image processing and segmentation could be up to 12 and 15 hours respectively, according to the systematic review and other reports (9,10). The whole process of creating a 3D printed liver model could be up to 100 hours, and this does not take



Figure 4 3D printed model for case 1 who is a 67-year-old male with renal tumour at the upper pole of left kidney. Comparative views of the CT scan at the nephrographic phase [(A) axial, (B) coronal and (C) sagittal planes] and corresponding views of the physical model [(D) superior and median view, (E) median and anterior view, (F) lateral view]. An inferior polar cyst is also displayed on this model (translucent yellow). The cubes show the 3D printed model orientation in space (I = inferior face, A = anterior face, L = lateral side, S = superior face, P = posterior face, M = median side). Case 1 underwent a left radical nephrectomy for a $65 \times 56 \times 42$ mm clear cell renal cell carcinoma, pT1bN0Mx, Fuhrman grade 3. The arterial tree is presented in opaque magenta, the collecting system in opaque yellow, and opaque orange for tumour display. The renal vein and renal parenchyma are kept translucent to allow the best visualization of the relationships between the renal tumour and surrounding structures. Reprinted with permission from Bernhard *et al.* (15). 3D, three-dimensional.

into account post-printing work, which could take up to 4–5 days as reported in some studies (37-39). Image postprocessing and segmentation of renal CT or MRI data for 3D printing is relatively easier when compared to those for heart or liver 3D printing since the contrast-enhanced renal parenchyma, renal vessels and renal tumours can be easily segmented through automatic or semi-automatic approach. In contrast, some manual editing is required for segmenting cardiac or hepatic structures which could be timeconsuming (9,10). In this review, only two studies reported the time spent on image processing and segmentation, however, most of the studies (10 out of 15) reported the time spent on 3D printing. With time spent on after 3D printing taken into account, the whole process could take up to a few days, thus, this long duration of 3D printing process needs to be addressed in future studies to make it a feasible and acceptable tool in clinical practice.

Cost of printing 3D kidney models is another challenge,

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Figure 5 3D printed model for case 7 who is a 53-year-old female with renal tumour at the interpolar region of left kidney. Comparative views of the CT scan at the nephrographic phase [(A) axial, (B) coronal and c sagittal planes] and corresponding views of the physical model [(D) superior view, (E) median view]. The cubes show the 3D printed model orientation in space (I = inferior face, A = anterior face, L = lateral side, S = superior face, P = posterior face, M = median side). Case 7 underwent a left partial nephrectomy for a $21 \times 15 \times 15$ mm angiomyolipoma. Description of colour corresponding to different renal structures and tumour is the same as in *Figure 4*. Reprinted with permission from Bernhard *et al.* (15). CT, computed tomography.

and this depends on the type of 3D printers and materials used for 3D printing. This review shows that with use of low-cost materials or home-made 3D printers, the cost associated with 3D printed kidney models is less than USD\$150, as shown in some studies (25,27,28,33). However, a high quality kidney model printed with flexible materials such as TangoPlus and with different colours to highlight renal tumours and vascular or renal collecting structures can cost up to USD\$1,000 (15,16,18,26). Therefore, future research is needed to reduce the costs associated with 3D printed kidney models. There are some limitations in this review. First, 3D printing in renal disease is an emerging area compared to other medical field such as orthopaedic or cardiovascular applications, thus, only a small number of studies are available in the literature. Of 15 studies included in the review, half of them provided quantitative assessments of 3D printed models in renal disease. Further, the quantitative findings provided by these studies are quite variable as shown in the analysis because they focus on different aspects such as dimensional accuracy and tumour volume measurements. Therefore, more studies are needed to provide further

evidence on the accuracy of 3D printed kidney models in replicating both anatomy and pathology. Second, this review shows that most of the studies did not provide information on the time spent on image processing and segmentation, most likely due to the use of different software tools (both commercial and open source software packages) for image processing and segmentation by different research groups. Further, as discussed previously, case reports or case series still dominate the current research of 3D printing in renal disease with 1-3 models reported in 6 studies and 5-10 models in 9 studies. There is a lack of original studies (either retrospectively or prospectively designed studies) looking at the clinical value of 3D printed kidney models, and this should be addressed by future studies. Finally, this review shows that it could take up to 30 h to print a 3D kidney model, and the up to a few days for the entire 3D printing process. Therefore, further improvement on production speed and cost reduction is necessary.

In conclusion, this systemic review analyses 15 studies reporting the clinical value or usefulness of 3D printed kidney models in renal disease with findings showing the feasibility and accuracy of 3D printed kidney models in delineating both anatomical renal structures and pathologies. 3D printed models are shown as useful tools in pre-surgical planning and simulation of renal tumours, in particular in minimally invasive renal nephrectomy of renal tumours. 3D printed models improve patient's understanding of their renal disease status, and enhance medical trainees' knowledge of renal malignancy when compared to viewing medical images alone. Future research is required to address the issued related to long duration and high cost associated with 3D printing process. Quantitative and qualitative assessments of clinical value of 3D printed kidney models based on a large cohort of patients are also needed to provide robust evidence on this rapidly growing technique.

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Footnote

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