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Development and Validation of Augmented Reality Training Simulator for Ultrasound Guided Percutaneous Renal Access

Yanyu Mu, *The University of Western Ontario*

Supervisor: Peters, Terry. M., *The University of Western Ontario*

Co-Supervisor: Eagleson, Roy, *The University of Western Ontario*

A thesis submitted in partial fulfillment of the requirements for the Master of Engineering Science degree in Biomedical Engineering

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Abstract

Percutaneous renal access (PCA) is a critical step in needle-based renal procedures. Traditional PCA training relies on apprenticeship, which raises concerns about patient safety and limits training opportunities. In this thesis, we reviewed simulation-based training for PCA, described the development of a novel augmented reality (AR) simulator for ultrasound (US)-guided PCA, and evaluated its validity and efficacy as a teaching tool.

Our AR simulator allows the user to practice PCA on a silicone phantom using a tracked needle and US probe emulator under the guidance of simulated US on a tablet screen. 6 Expert and 24 novice participants were recruited to evaluate the efficacy of our simulator.

Experts highly rated the realism and usefulness of our simulator, reflected by the average face validity score of 4.39 and content validity score of 4.53 on a 5-point Likert scale. Comparisons with a Mann-Whitney U test revealed significant differences ($p < 0.05$) in performances between the experts and novices on 6 out of 7 evaluation metrics, demonstrating strong construct validity. Furthermore, a paired T-test indicated significant performance improvements ($p < 0.05$) of the novices in both objective and subjective evaluation after training with our simulator.

Our cost-effective, flexible, and easily customizable AR training simulator can provide opportunities for trainees to acquire basic skills of US-guided PCA in a safe and stress-free environment. The effectiveness of our simulator is demonstrated through strong face, content, and construct validity, indicating its value as a novel training tool.

Keywords: Augmented reality, Percutaneous renal access, Training simulator, Ultrasound-guided needle insertion

Summary of Lay Audience

Percutaneous renal access (PCA) is the initial step to gain access to the kidney for treating common kidney diseases such as kidney stones. At present, mastering of this technique relies on extensive clinical training. However, it is very challenging to keep up with the increasing training demand for many training centres. To lessen the burden of the clinical education and deliver safer patient care, training simulators were employed to provide supplementary training opportunities. This thesis reviewed the existing training simulators for PCA and found no augmented reality (AR)/ virtual reality (VR) simulator available for ultrasound (US)-guided PCA, which is a safer alternative to fluoroscopy (FL)-guided PCA. Therefore, the goal of this work was to develop and validate a training simulator for US-guided PCA.

Following a minimalism design approach, we integrated three-dimensional (3D) printed hardware components, an easy-to-make silicone phantom, and personal mobile device to build an low-cost training simulator for US-guided PCA. Since the surgical scene, including the the kidney and US images are simulated and visualized in AR, the tradition lab setting is no longer required. Trainees have the option to practice at home in a stress-free environment. In addition, this simulator provides performance feedback via direct visualization and data sheet, which facilitate deliberate practice without supervision. For educators, new training content, such as patient specific cases can be easily imported to this simulator without any hardware alteration.

A user study was conducted to validate some aspects of this simulator, and demonstrate that training using our simulator resulted in significant skill improvements. To incorporate this simulator into the training curriculum, more rigorous validation is required for future work.

Co-Authorship Statement

The following thesis contains one manuscript that has been published:

Mu, Y., Hocking, D., Wang, Z., Garvin, G.J., Eagleson, R. and Peters, T.M., Augmented reality simulator for ultrasound-guided percutaneous renal access. *International Journal of Computer Assisted Radiology and Surgery*, 15(5):749–757, 2020.

My contribution to this work including engineering design, programming, prototyping and testing, study design, data collection and analysis, drafting and revising the manuscript. Hocking, D. Wang, Z., and Garvin, G.J. provided clinical insight. Hocking and D. Wang Z. also aided with testing, user study and recruitment. Eagleson, R. and Peters, T.M. provided supervision on design and testing. All authors contributed editorial feedback during the manuscript review.

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List of Abbreviations, Symbols, and Nomenclature

Terminology

| | |
|------|------------------------------|
| 3D | Three-dimensional |
| PCA | Percutaneous renal access |
| AR | Augmented reality |
| VR | Virtual reality |
| OR | Operating room |
| US | Ultrasound |
| FL | Fluoroscopy |
| PCNL | Percutaneous nephrolithotomy |
| GRS | Global rating scale |
| HMD | Head-mounted display |
| CT | Computed tomography |
| MRI | Magnetic resonance imaging |
| EMT | Electromagnetic tracking |
| SDK | Software development kit |
| STL | Stereolithography |
| ROI | Region of interest |
| DLL | Dynamic-link library |
| SO | Shared object |

Chapter 1

Introduction

Percutaneous renal surgery is a standard minimally invasive procedure for diagnosis or treatment of a variety of renal pathologies such as kidney stones and tumors [84]. In most cases, the first step of percutaneous renal surgery is to create needle access from the skin into the kidney under ultrasound (US) or fluoroscopic (FL) guidance [75]. This initial needle access serves as a safe passage for larger surgical instruments, allowing surgeons to perform the surgery without opening layers of tissues. Compared to open renal surgeries, percutaneous procedures are less invasive and are associated with lower morbidity [43]. Nevertheless, access related complications, such as serious hemorrhage and pulmonary injuries, have been reported [47][68]. Given that the kidney is highly vascularized, and is surrounded by vital organs such as the lung and the colon, percutaneous renal access (PCA) is considered as the most challenging step during needle-based renal procedures. Thus, establishing a safe and efficient PCA is crucial to the success of percutaneous renal surgeries and the recovery of patients [46].

This chapter includes the steps taken to guide the design of a training simulator for PCA.

1. Provide a relevant clinical background of PCA, identify the technical challenges performing this procedure, and associated complications.
2. Review the current training method, identify the problem in training, and discuss potential solutions.

3. Summarize the characteristics of medical training simulators, review the existing training simulators for PCA, identify what is missing, and discuss feasible engineering solutions.

1.1 Clinical Background

1.1.1 The Early Development of PCA

The first documented PCA was performed by Thomas Hillier in 1864, to treat a 4-year-old boy with a severe hydronephrotic right kidney due to a congenital defect. Over a period of four and a half years, Hillier aspirated the boy's right kidney periodically for temporary drainage, and even attempted to establish a permanent fistula. Unfortunately, the fistula did not persist, and the boy later died of a stone that obstructed his healthy left kidney at the age of 8 [13]. As x-ray imaging and pyelography became available for visualizing the urinary system, Nils Alwall applied the liver biopsy technique on the kidney and performed the first image-guided percutaneous renal biopsy in 1944 [6][75]. Poul Iverson and Claus Brun [49] subsequently perfected this technique, and it was quickly adopted by physicians around the world for renal diagnosis. Henceforth, PCA techniques were adopted for the treatment of many urological diseases, such as kidney stones and tumors. Improvement in techniques and advancement in equipment resulted in percutaneous renal procedures, and because of its low complication rate and lower blood loss [15], percutaneous renal procedures became the preferred treatment over open renal surgeries .

1.1.2 Image-Guidance for PCA

During minimally invasive surgeries (MIS), image-guidance is crucial to surgeons, just as navigational technology is to a pilot when flying at night [70]. Since surgeons can no longer see or feel the pathology or anatomical relationships as they could during open surgeries, this information must be provided by intraoperative real-time imaging modalities. The most com-

monly used image-guidance modalities for PCA are fluoroscopy (FL) and ultrasound (US). In addition, computed tomography (CT)-guidance may be required for patients with abnormal morphology such as morbid obesity, retrorenal colon, enlarged spleen, etc. [46].

FL has been used as the primary imaging modality for PCA guidance, as it provides high-resolution imaging of the collecting system and good visibility of small stone fragments, needle, and guidewire [46][48]. However, cumulative radiation exposure to patients and providers during FL-guided PCA poses an increased health risk [97], while the long term effect of low-dose ionizing radiation is still being investigated [18][62].

Compared to FL, US has the advantage of imaging of the visceral organs surrounding the kidney in real time, eliminating the need for contrast material, and provides comparable surgical outcomes at a lower institutional cost. Above all, US-guidance is radiation free, hence ideal for pregnant women and children [34][97]. Disadvantage of US-guidance including the lack of anatomic details, limited visualization of the guidewire, and difficulty tracking of small stone fragments (Figure 1.1) [46][48].

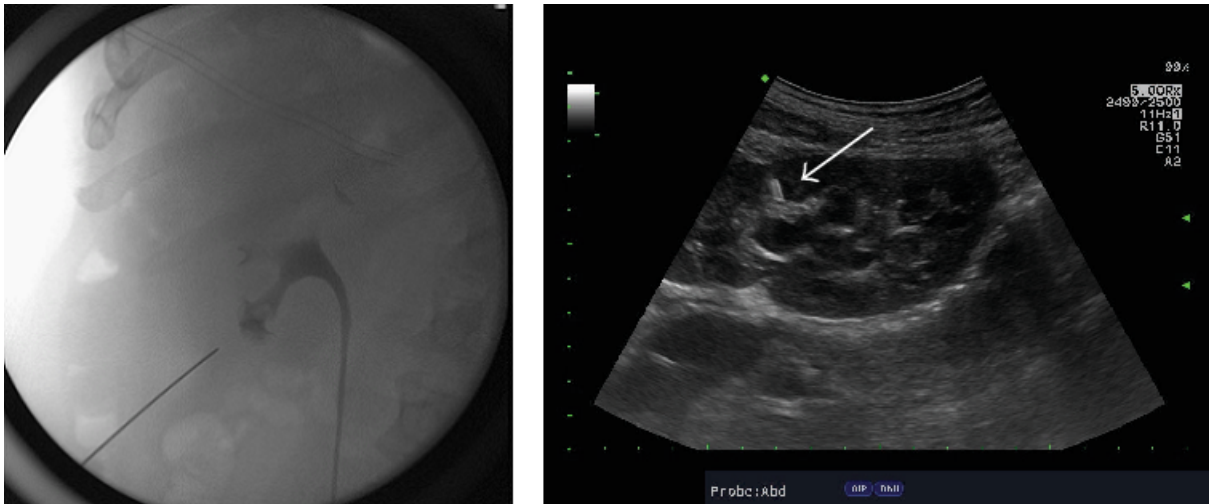


Figure 1.1: FL-guided PCA (Left) [86] vs US guided PCA(Right) [16]. Image used with the permission of Mary Ann Liebert Inc. and Hindawi.

A number of studies comparing clinical outcomes of percutaneous nephrolithotomy (PCNL) using US or FL guided access found no significant differences between the two groups in terms

of total operating time and success rate. Whereas the US groups had higher stone-free rate, shorter radiation exposure time, fewer access attempts, and lower incidence of hemorrhage (bleeding), the FL groups had shorter access time [2][7][12][25][38][42]. A Higher incidence of hemorrhage was found to be associated with the larger sheath size and a higher number of punctures performed during FL-guided procedures [7].

In conclusion, FL and US each have their advantages and disadvantages. While choosing an imaging modality for PCA guidance, the characteristics of each modality should be considered. Whereas US-guided PCA has gained popularity around world including China and the U.S., this technique is rarely practiced in Canada, mainly due to the lack of training [19]. As more Canadian institutions move towards adopting US-guided PCA, it is recommended to use US-guidance on patients with dilated collecting systems, and which are free of staghorn stones [12][53]. In addition, US can be used as an adjunct to FL as a strategy to reduce radiation exposure [2][110].

1.1.3 US-Guided PCA Techniques and Complications

Anatomical Considerations

While planning for the best approach to the collecting system, a solid understanding of the renal anatomy and the renal arterial system is of utmost importance for safe and efficient PCA. The kidneys are positioned between the abdominal lining and the back, defined as the retroperitoneal space. Nephrons are the filtering units of the kidney that filter the blood to regulate chemical concentration and produce urine. There are approximately one million nephrons found through out the medulla and the cortex, which is the pyramid shaped segments and the outer region of the kidney respectively. The renal pyramids project into funnel shaped chambers called calyces. The calyces are positioned radially around the renal pelvis, which is the innermost hollow centre of the kidney where urine collects. The kidneys are highly vascularized organs, with the renal artery branches into anterior and posterior divisions [39]. Between these two divisions, lies the avascular field, known as the Brödel's line (Figure 1.2) [39]. To

avoid vascular injury, the best point of entry is through the fornix into a posterior calyx, which usually traverses Brödel's line [39]. Direct puncture into the infundibulum or renal pelvis could result in significant hemorrhage. Moreover, the resulting tract would not be able to provide adequate stability for other surgical instruments such as a nephrostomy tube [53][77]. Inadvertent puncture of an anterior calyx can also lead to increased risk of bleeding and difficulty accessing the ureter [39][46].

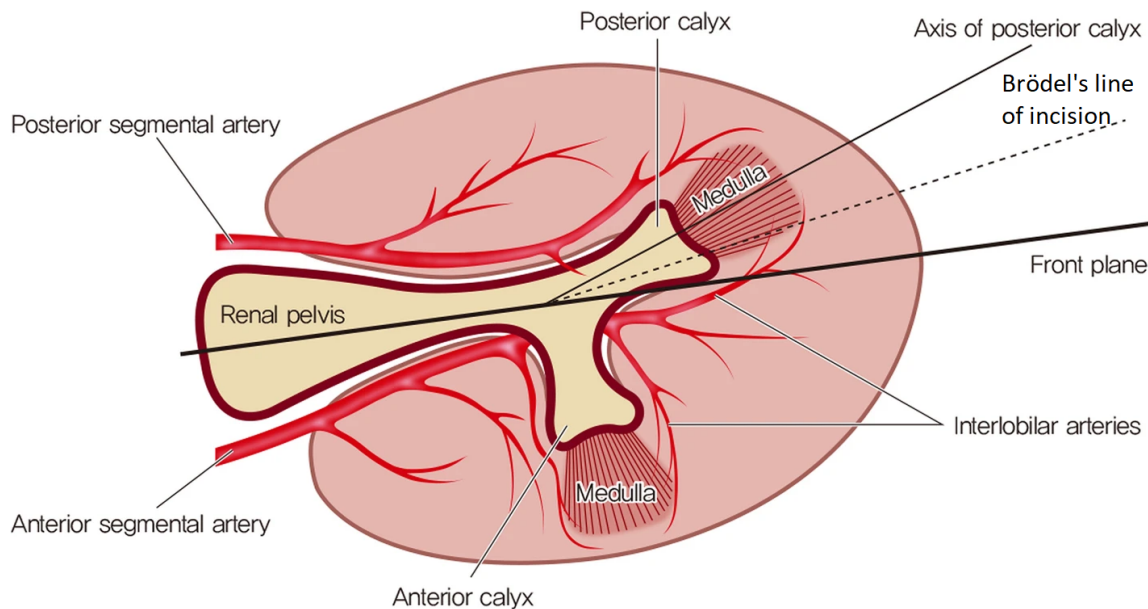


Figure 1.2: Access through a posterior calyx is preferred for lower risk of vascular injury [44]. Image used with the permission of National Center for Biotechnology Information.

Techniques of US-Guided Needle Insertion

US-guided PCA is often performed using a curvilinear ultrasound probe and an 18 gauge needle. During scanning, if the kidney is partially obscured by acoustic shadowing from the ribs, a 30-45 degree rotation can be applied to align the probe to the ribs [19]. Needle insertion can begin once a clear path to the target in the posterior calyx is planned on a longitudinal US view of the kidney [19]. During insertion, the goal is to maintain alignment between the needle and the US beam (needle-beam alignment), in order to visualize the needle in its entirety

(Figure 1.3). If misalignment happens, needle advancement should be stopped, and the US probe should be used to identify and redirect the needle [19]. Successful access is verified on observation or aspiration of urine after removing the stylet [19]. A needle guide would help with needle-beam alignment at the expense of the flexibility of the accessing angle [19].

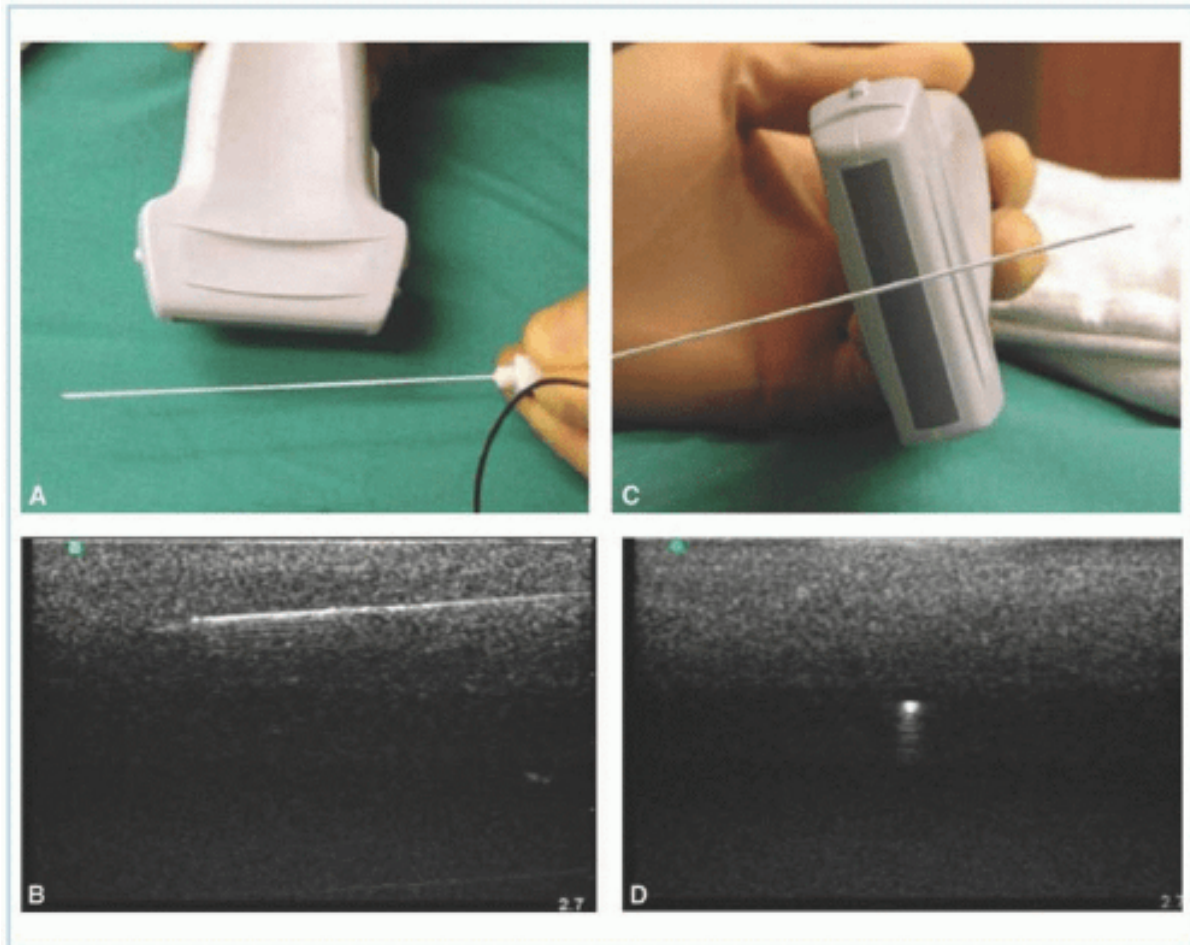


Figure 1.3: In-plane alignment technique (left), needle and US beam are in the same plane. Out-of-plane technique (right), needle is perpendicular to the plane of US beam ([94]. Image used with the permission of Wolters Kluwer Health Inc.

Access Related Complications

Hemorrhage is the most common complication of PCA, however, minor hemorrhage during PCA typically does not require intervention. Nevertheless, major hemorrhage happens to 1%-

15% of the patients, requiring blood transfusion [39][74][105]. In addition to renal hemorrhage, inadvertent needle puncture can cause complications in other organs and structures adjacent to the kidneys including thoracic injuries (4%-16%), and visceral organ injuries such as colon injuries (<1%) [39]. Liver and spleen injuries are very rare in the absence of anatomic abnormalities [39][92].

1.1.4 PCA Training Status Quo

In North America, PCA is often performed by interventional radiologists as a staged procedure before planned PCNL [46]. Recent studies have suggested that urologists can obtain PCA safely and effectively, transforming PCNL into a single-stage procedure, which eliminates the need to transfer patients between different departments [23][93]. In addition, having the ability to independently obtain the access provides better flexibility for the urologists, when selecting the optimum tract or making a secondary tract [46][53]. An American survey revealed that urologist-obtained PCA is associated with a higher stone-free rate (86% vs 61%) and lower number of complications (5 vs 15), in comparison with radiologist-obtained PCA. However, only 11% of urologists obtain access themselves, mainly due to the lack of training [101]. Despite previously reported benefits of urologist-obtained PCA, only 37.5% of Canadian residents train at centres where urologists obtain their own PCA, independent of radiologists [72].

At present, US-guided PCA is taught through the traditional apprenticeship approach, where trainees would perform this procedure on patients under the supervision of senior physicians, until the trainee is considered proficient to operate independently. Therefore, factors such as concerns for patient safety, and restricted training hours, limits the trainees' opportunity to gain experience with this procedure [76]. Moreover, achieving competency in PCA is challenging due to the steep learning curve. Two recent studies evaluated the learning curve for US-guided PCNL and reported that it would take 60 cases for novice trainees to gain surgical competency and 120 cases to achieve excellence [85][104]. For experienced surgeons, 20 cases are sufficient for the transition from FL-guided to US-guided PCNL [96]. Furthermore,

competency evaluation is usually conducted by the supervising physician, using checklists or in-training evaluation reports, which suffer from subjectivity [59][64].

1.1.5 Challenges and potential solutions

In summary, US-guided PCA is a challenging procedure associated with a steep learning curve. To achieve competency in US-guided PCA, a trainee should master technical skills, including a good understanding of renal anatomy, the ability to obtain and interpret US images, and proper skills to maintain needle-beam alignment during access [19]. However, the traditional training approach is not sufficient to meet the training demands except at some high-volume centres [46]. Beiko et al. [19] suggested the training be split into two different skill sets, diagnostic renal imaging, and needle control, for a more structured learning approach. More specifically, the authors advised trainees to practice renal US imaging on patients or abdominal phantoms whenever an opportunity arose, to shorten the learning curve [19]. Likewise, training simulators for percutaneous needle access can be a potential solution to the training demand for mastering needle control. To further improve the accessibility, simulated US could be used to eliminate the need to access US machines, providing flexibility in training time and location. In addition, training simulators could provide objective feedback to trainees, decreasing the demands for supervision by senior physicians.

1.2 Simulation-based Medical Education

Training simulators were first used in high-risk professions such as aviation and aerospace, where mistakes can be fatal [73]. Similarly, during a medical procedure, the smallest error could have dire consequences. Medical training simulators provide a stress-free and harm-free environment for health professionals to acquire knowledge and surgical skills through deliberate practice [82]. More importantly, simulation-based training can minimize risks as the trainees begin to operate on patients [5]. While simulators are no substitute for clinical training

in the operating room (OR), they can nevertheless help to shorten the learning curve [3][54]. A number of laparoscopic and endoscopic training simulators have demonstrated the transfer of skill from a virtual environment to the operating room [80]. Therefore, simulation-based training is now considered an essential adjunct to the traditional apprenticeship training, and it has been successfully adopted by some centres [4][81]. However, a recent survey from the United Kingdom indicated the lack of simulation-based training for many standard urology procedures including PCA, possibly due to the high cost of the existing simulators [64].

1.2.1 Simulator Fidelity

Existing medical training simulators can be categorized into low-fidelity (LF) and high-fidelity (HF) trainers based on their physical resemblance to the surgical procedure. Commonly used LF simulators including intravenous injection trainers, silicone pads for suture practice, box trainers for laparoscopic and endoscopic procedures, etc. A LF simulator typically provides training on a specific task that is designed to improve manual dexterity, hand-eye coordination, or tissue handling [78]. Even though these simulators have limited functionality, they are usually highly portable, easy to set up, and low maintenance [78]. Above all, they are very effective for novice trainees to acquire the basic surgical techniques at the initial stage of training. [78].

HF simulators are mostly computer-based applications, working in conjunction with hardware such as surgical tool emulators, haptic devices, tracking systems, phantoms, or manikins, for complex procedure simulation. One common approach to creating a training scenario that resembles the actual surgical procedure is to replicate it digitally, in a virtual reality (VR) environment. The other option is to artificially enhance the physical world with additional information in the form of a digital overlay, in other words, augmented reality (AR) [98]. In the field of urology, AR/VR simulators are widely used for endoscopic and laparoscopic training [31]. A significant advantage of AR/VR simulators is their ability to record training data, which could be used for progress monitoring or providing objective performance feedback. Compared to

the other types of simulators, the AR/VR technology offers the freedom to simulate various clinical scenarios, a range of tasks and difficulty levels, and patient-specific anatomies [64]. On the flip side, the cost of development and maintenance may be much higher. Live animal models are also considered high fidelity as they provide realistic haptic feedback, respiratory movement, and bleeding [4]. However, due to the limited supply, cost, and ethical considerations, live animal models are not suitable for repetitive practice [8].

It is commonly believed that the effectiveness of a simulator improves as its realism increases. Yet, this is only partially true. Beyond a certain level, the simulator performance reaches a plateau, regardless of the amount of money and effort invested [52]. A large body of studies has demonstrated that HF simulators are not necessarily more effective than LF, evidently suggesting that learning outcomes of training simulators are independent of their physical fidelity [20] [55] [61] [78]. Instead, more attention should be paid to the psychological process during a surgical procedure [57] [21]. Simulator design should follow a minimalist approach to direct the focus of the user to the critical tasks after establishing the objectives of training and intended trainee group [21] [61]. Any additional non-essential information could potentially cause distraction or cognitive overload [57] [21]. In addition, other aspects, including the accessibility, versatility, reproducibility, and maintenance requirements, should also be considered during the design process [20].

1.2.2 Review of Existing Training Simulators for PCA

Existing training simulators can be grouped into the following categories: live animal models, biological or non-biological bench models, and VR/AR-based simulators [64].

Live Animal

Two studies have reported the use of anesthetized live porcine models for PCA training, mainly because of the similarity of the pig and human renal anatomy [41] [58]. In addition, such models provide tactile feedback, respiratory movement, and bleeding that is superior to other

types of training simulators [58]. However, training on live animals requires ethics approval, and veterinary support, which also increases the training cost [58]. Therefore, the use of live porcine models for training is limited. Mishra et al. [58] concluded that live porcine models are more suitable for skill assessment.

Biological Bench Models

Bench models made of ex-vivo porcine or bovine kidneys were the most common type of PCA trainers from the 2000s to early 2010s [64]. The dissected kidneys were wrapped within various types of materials including foam [22], sponge [1], silicone [87], chicken carcass [32] [35] [100], as well as a combination of porcine skin flap [37], subcutaneous fascia and muscle [40] [107], and ribs (figure 1.4) [26] [88] [109]. These trainers are generally cheap and easy to make. Furthermore, artificial stones can be manually placed into the renal pelvis for PCNL training. Similar to the live animal model, biological bench models are designed to work with image-guidance equipment and thus require wet lab access. While all except for one of these trainers can be used for FL-guided PCA, four of them are not US compatible [22] [1] [32] [100]. In addition, each biological bench model can be used only for a limited number of times. All of these factors limit the accessibility, versatility, and reproducibility of these biological bench models.

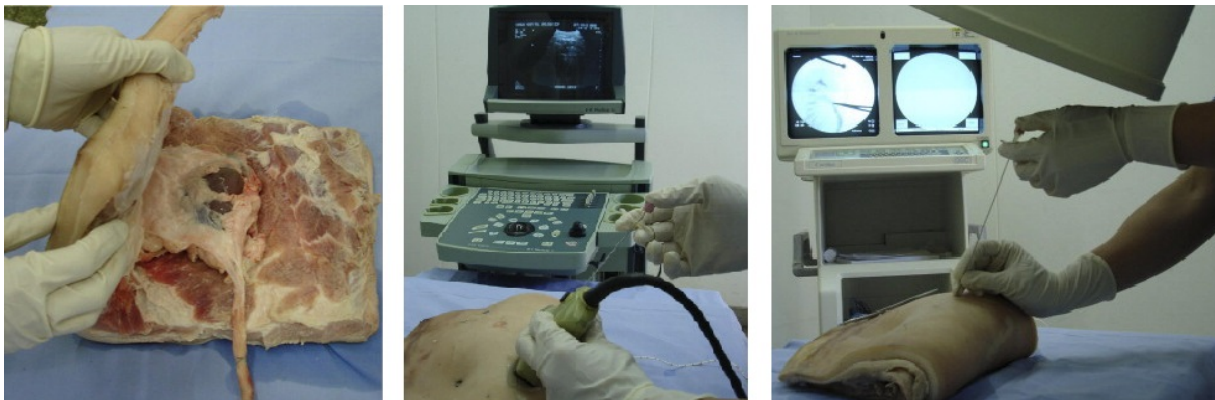


Figure 1.4: An example of ex-vivo porcine model for US/FL-guided PCA training [107]. Image used with the permission of Elsevier.

Non-Biological Bench Models

Since the mid 2010s, there has been an increase in research involving non-biological bench models, ranging from the low fidelity vegetable [83] or sponge [91] to anatomically correct silicone [106] or hydrogel [30] kidney models, to highly detailed three-dimensional (3D) printed silicone models based on patient CT scans [14] [95] [103]. All of the above models were design for FL-guided PCA with no US compatibility. Only three commercial US compatible bench models exist for full PCNL procedures: the PCNL trainer (Encoris), Perc Trainer (Mediskills), and PCNL trainer LS40 (Samed GmbH Dresden). These synthetic training models require a dry lab setting and access to C-arms or US machines.

To eliminate radiation exposure during FL-guided PCA training sessions, Veneziano et al. developed a 3D printed mini C-arm trainer (SimPORTAL), using two webcams to simulate FL images, and a silicone flank model for needle insertion (figure 1.5) [66] [99]. Images obtained from the cameras were filtered and fused together through the "chroma-key" technique [99]. This low cost replication of a C-arm provides a radiation-free training environment and dramatically improves the accessibility.

In summary, the artificial bench models are more accurate at replicating human anatomy, and more durable than the biological bench models for repetitive practice. However, their tactile feedback is inferior to that provided by biological tissues. One common drawback of these bench models is the lack of performance feedback to trainees. Hence the presence of an expert may be required to provide guidance [24].

VR/AR-based Simulators

The PERC Mentor (Symbionix) was the first VR simulator for FL-guided PCA, which has been thoroughly validated for training and assessment purposes. This simulator features several training scenarios of increasing complexity which were created in a virtual surgical environment, a virtual C-arm controlled by the touch screen or a foot pedal, a sensorized needle for location tracking, a torso mannequin that mimics various human tissues embedded with

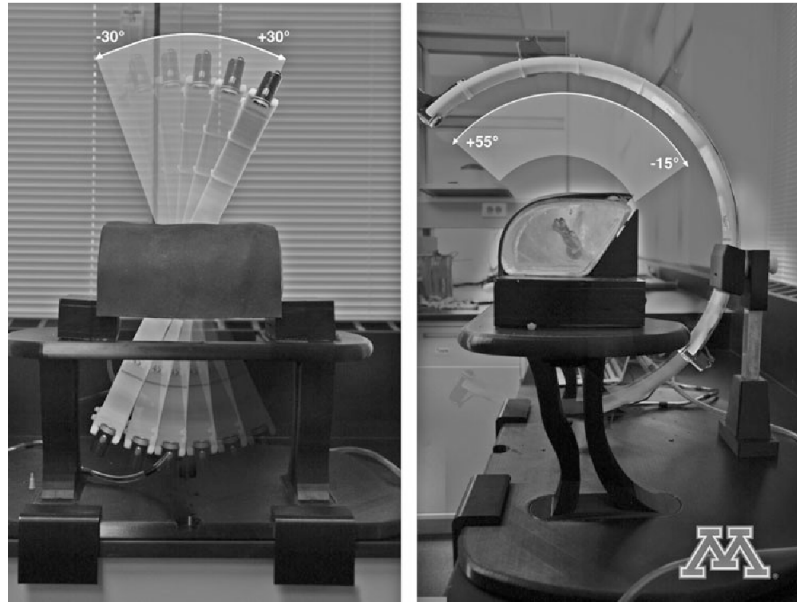


Figure 1.5: SimPORTAL-A mini C-arm simulator and a silicon flank model for radiation free PCA training [99]. Image used with the permission of Mary Ann Liebert Inc.

palpable ribs, the visualization of the virtual anatomy with respiratory movement, a virtual assistance providing warnings and directions, and performance reports of recorded training parameters such as operation time, FL time, number of attempts, rib collisions, injuries to adjacent organs, etc. [63]. A number of studies have reported its face [45] [60] [81], content [45] [58] [60] [81], construct [28] [45] [60] [81], and predictive validities [54] [60], as well as skill acquisition of trainees [28] [56] [69] [108]. Furthermore, two studies reported using the PERC Mentor for PCA assessment [63] [65]. However, the PERC Mentor hasn't been widely adopted by training centres mainly due to the high cost (\$100,000) [64].

In 2019, Tai et al. [90] published a study of a novel AR-based simulation platform —SimPCNL for FL-guided PCA training. SimPCNL consists of a PC, two PHANTOM Omni devices that provide realistic tactile sensation, and a Microsoft Hololens, which provides an AR view of the surgical scene [90]. The combination of visual and haptic simulation effectively replaces the traditional training phantoms. More importantly, the force feedback from the haptic devices is used for physics-based tissue mesh deformation in real-time [90]. These force and velocity

data are also recorded as valuable evaluation parameters [90]. In this study, the authors successfully demonstrated the face, content, and construct validities of this simulator [90]. Criterion validity was also established through a comparison with the PERC Mentor (figure 1.6) [90]. Even though the cost of the HoloLens and the PHANTOM Omnis is higher compared to the bench models, this simulator is much cheaper and more portable than the PERC Mentor .

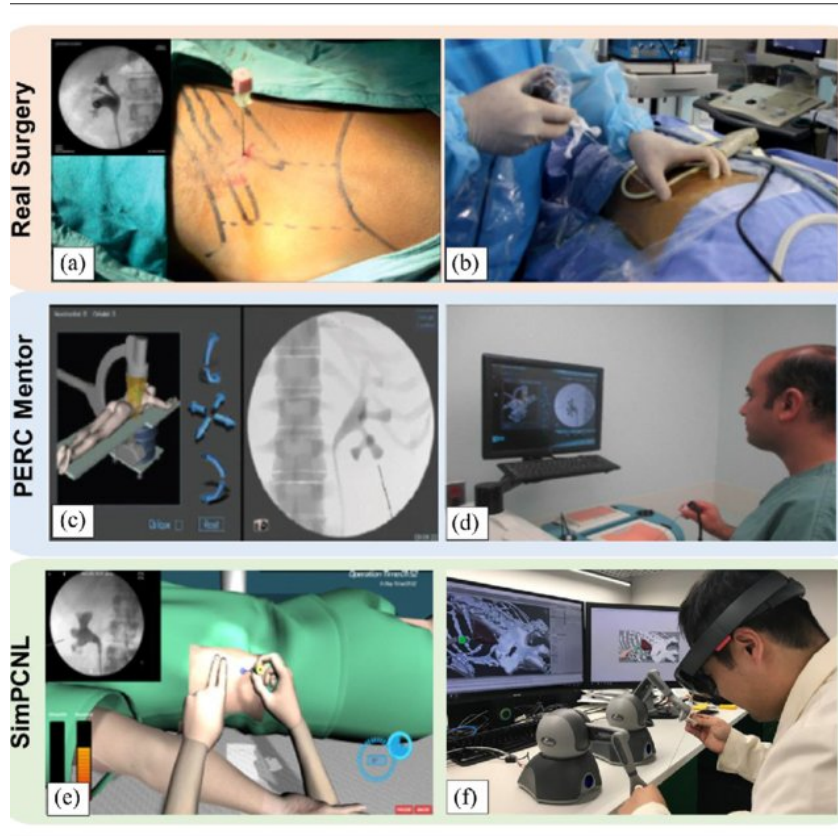


Figure 1.6: Comparison between real surgery, PERC Mentor, and SimPCNL. The left column displays the surgical interface. The right column displays the operation scene [90]. Image used with the permission of SAGE.

In conclusion, AR/VR training simulators can simulate various training scenarios, record objective performance parameters for skill assessment, and provide a radiation-free environment for repetitive, deliberate practice. Yet, both of these simulators are designed for FL-guided PCA. No AR/VR training simulator was reported for US-guided PCA.

1.2.3 Challenges and possible solutions

In a review study of 109 articles of simulation in medical education, Issenberg et al. [9] reported the top simulation features lead to effective learning are feedback (47%), repetitive practice (39%), curriculum integration (25%), and range of difficulty levels (14%). AR/VR training simulators have a clear advantage over bench models and live animal models in providing these features. The existing AR/VR simulators are validated but underutilized due to their high cost, highlighting the need for a low-cost simulator for US-guided PCA.

Visualization

Two types of AR/VR visualization devices are monoscopic devices, including cellphones, tablets, monitors etc, and stereoscopic head-mounted displays (HMD) such as the hololens, Vive (HTC), Google Cardboard, etc. The high-end HMDs are often equipped with multiple sensors and cameras for tracking the user's poses relative to the surrounding environment. Google Cardboard or similar passive devices can be used with cellphones to provide a stereo view. The simpler mobile devices merely serve as a display (VR) or capture video of the real world using a single camera (AR). Even though stereoscopic devices enhance depth perception, selection of an AR/VR device should be made according to the surgical scenario.

During an actual needle access procedure, the physician would be standing at a fixed location, operating at elbow height. His/her head motion would be limited, switching focus between the operating hands and the US display. Thus the viewpoint can be fixed once the procedure starts during training. The advanced HMDs provide no additional benefit over a simple mobile device for operator tracking.

US simulation

Since there is limited availability of US machines, US simulation would significantly improve the accessibility of a training simulator. There are two different approaches to simulate US images: real-time simulation based on mesh models [11], or manipulation of pre-recorded US

volume, each having its own advantages and disadvantages.

The pre-recorded US image approach would be more realistic than the use of simulated images, but it requires scanning of the real-patient for each training case. Additionally, the US effect of a needle needs to be added to the volume in real-time, which requires more programming effort, since the AR/VR development platform (Unity) has very limited capability for processing medical images. However, it is feasible to expand Unity's functionality via native plugins in order to achieve the desired result. However, this simulation approach heavily relies on the processing power of the mobile device.

On the other hand, the PLUS toolkit [11] can be used to simulate US images based on mesh models of the renal system and the needle. The acoustic properties of different tissues and material can be carefully customized for optimum results. This process can be performed on a PC/server to reduce the workload on a mobile device. Transmission between the PC/Server and the mobile device can be established via a simple communication plugin.

Tracking

Optical tracking and electromagnetic tracking (EMT) are the most commonly used systems for surgical navigation, with both systems providing sub-millimeter tracking accuracy. However, the optical systems require a direct and continuous line of sight between the sensor and the camera, and the EMT systems are prone to interference from metallic objects and electromagnetic fields [50]. Vuforia is a proprietary image processing software development kit (SDK) that uses feature detection algorithms to register and track image markers or 3D objects. Setting up the system only requires registering a unique marker to the Vuforia database, then attached it to an object which needs to be tracked. A camera is also required to capture the images, which is available on almost all mobile devices. The tracking accuracy of Vuforia varies, depending on the quality, size of the image, the camera resolution, and the lighting condition. With the new generation of mobile devices featuring depth-sensing cameras, the image tracking accuracy would be greatly improved.

Phantom

Haptic devices such as the PHANTOM Omni provide superior haptic feedback compared to that provided by simple artificial phantoms. Although it costs over \$1000 and the range of motion is limited. Poniowski et al. [71] studied the needle insertion forces for PCA on a human cadaver. While the peak of the force-depth representing skin puncture is evident, the peak representing renal collecting system puncture is non-uniform and indistinguishable at times. Therefore, confirmation of perforation of the renal collecting system should not be relied on using haptic feedback.

A wide variety of hydrogel or silicone materials is available for fabricating tissue mimicking phantoms. The hydrogel-based materials are often used to make US compatible phantoms, because their acoustic property is similar to that of human soft tissue [17]. However, some of these hydrogel materials degrade rapidly due to microbial invasion and dehydration [67]. Poly(vinyl alcohol) cryogel, PVA-C, is a magnetic resonance imaging (MRI) and US compatible material that can be preserved for several months [89]. But it cannot sustain repetitive needle puncture. While silicone materials are more durable for practicing needle insertions, their acoustic properties do not match those of human soft tissue. Maggi et al. [51] reported that the attenuation of silicone could be modified with additives if US compatibility is desired.

1.3 Objective

The objective of this thesis work was to develop a low-cost and accessible AR training simulator for US-guided PCA, utilizing personal mobile device and off-the-shelf components, to provide extra training opportunities to novice trainees, in addition to the traditional apprenticeship approach.

1.4 Thesis Outline

In response to the limitations of current systems outlined above, Chapter 2 of this thesis describes the development of a novel AR training simulator for US-guided PCA as well as the validation steps and results. Chapter 3 then discusses the advantages and limitation of the current work and proposed directions for future research.

Chapter 2

Development and validation of an augmented reality simulator for ultrasound-guided percutaneous renal access

This chapter is adapted from the following manuscript:

- Yanyu Mu, David Hocking, Zhan Tao Wang, Gregory J. Garvin, Roy Eagleson, and Terry M. Peters. "Augmented reality simulator for ultrasound-guided percutaneous renal access." *International Journal of Computer Assisted Radiology and Surgery*, 15(5):749–757, 2020.

2.1 Introduction

As reviewed previously, ultrasound (US)-guided percutaneous renal access (PCA) is gaining popularity over fluoroscopy (FL)-guided PCA for better procedure outcomes and reduced radiation in minimally invasive treatment of renal disease. However, the current apprenticeship

training cannot meet the growing training demand. As a supplement, simulator training can provide additional learning opportunities to novice trainees. Several augmented reality (AR)/virtual reality (VR) simulators for FL-guided PCA were reviewed in chapter 1. One of the main reasons that simulators are underutilized in training is their high cost. Therefore, we identify the need for a cost-effective, versatile and easily accessible simulator for US-guided PCA training.

In this chapter, we propose the development of a novel AR simulator, emphasizing the use of off-the-shelf components and incorporating a simple and easily made physical phantom for US-guided PCA, and evaluated its validity and efficacy as a teaching tool. The major contributions of our simulator are as follows:

1. The simulator was designed to exploit the popularity of modern portable devices and personal computers, which allows the trainees to use their personal devices for visualization and computation. This approach significantly reduces the cost and improves the accessibility of our simulator.
2. Through computer vision-based tracking of a silicone phantom, an US probe emulator, and a needle via a mobile device camera, we are able to simulate a virtual PCA procedure for the user to have a realistic training experience under AR without the need of any external tracking device. This implementation further reduces the cost.
3. Virtual anatomy models with different patient pathologies can be imported into our simulator as training cases, making it customizable for educators to create a variety of clinical scenarios ranging from common to extremely rare.

2.2 Material and methods

2.2.1 Phantom

The phantom consists of two layers of silicone, a 3mm deep skin mimicking layer on top of an 8.5cm deep soft tissue-mimicking layer. The skin mimicking layer was fabricated using a two-part silicone (Ecoflex 0010, Smooth-On inc., USA) with a Shore hardness of 00-10. Ecoflex[®] part A and part B were mixed with a 1:1 ratio by weight and cured at room temperature overnight. The cured surface was powdered with talc to create a dry and smooth surface for smooth movement of the US probe emulator. To achieve the softness of human soft tissue, a silicone tactile mutator Slacker[®] (Smooth-On inc., Macungie, PA, USA), Ecoflex[®] part A, and Ecoflex[®] part B were mixed at 2:1:1 ratio by weight. Although Slacker[®] provides many benefits such as increasing the softness, rebound, and self-healing properties of silicone [67], it also increases the tackiness which leads to greater friction during needle insertions. Consequently, we observed a small amount of silicone adhered to the needle shaft after each needle withdrawal. Glass microspheres have been used primarily as a filler material in the industry due to its low density. We introduced glass microspheres into the silicone mixture at a 4% weight ratio, which resulted in smoother needle insertion, and no silicone material adhesion to the needle shaft. We speculate that the ball-bearing property of glass microspheres effectively decreased the friction on the needle shaft, while maintaining the overall softness, rebound, and self-healing properties of the modified silicone.

2.2.2 Hardware

Initial Design

The initial design utilized Google Cardboard combining with a cellphone to provide better depth perception through immersive stereoscopic visualization. However splitting the cellphone display for stereoscopic view dramatically restricted the field of view (FOV). Moreover,

the generated stereoview was provided at the cost of a lower resolution, which was not suitable for inspecting details of the virtual renal anatomy at the operating distance. Lastly, the latency introduced by the software, that caused a perceptible lag between the time the US transducer or needle was moved, and the appearance of this motion on the display, induced motion sickness and eye fatigue while wearing a HMD [33]. Hence, a simple monoscopic mobile device was considered a more efficient and cost-effective option for visualization purposes.

Final Version

During the performance of a PCA, the physicians need to check their hand movement or the US machine display intermittently to confirm probe and needle position with reference to the US image. These head movements complicate the procedure, making it more challenging for novice trainees to master the basic techniques such as in-plane needle alignment. We eliminated the need for these head motions by combining the US image stream with the camera feed of the surgical site on the same tablet screen, therefore, providing a more intuitive user experience. The tablet screen was split into two sections: the top for the US image stream and the bottom for camera feed (Figure 2.1 Left). The tablet utilized in this project is a Samsung s5e (Samsung Electronics Co., LTD., Suwon, South Korea) with a 13-megapixel rear camera.

To track the poses of the US probe emulator, needle, and silicone phantom in real-time, unique image markers must be attached to each component securely. For this purpose, we designed the attachments for the needle and silicone phantom (Figure 2.1 Right) with SpaceClaim 19.1 (SpaceClaim Corp., Concord MA, USA). The stereolithography (.stl) file of an Ultrasonix C5- 2/60 curvilinear US probe model was downloaded from Plus toolkit printable three-dimensional (3D) models catalog (Perk Lab, Queen's University, Kingston, ON, Canada), then modified to incorporate an image marker. All components were then 3D printed using the Ultimaker 3 (Ultimaker B.V., Geldermalsen, Netherlands).

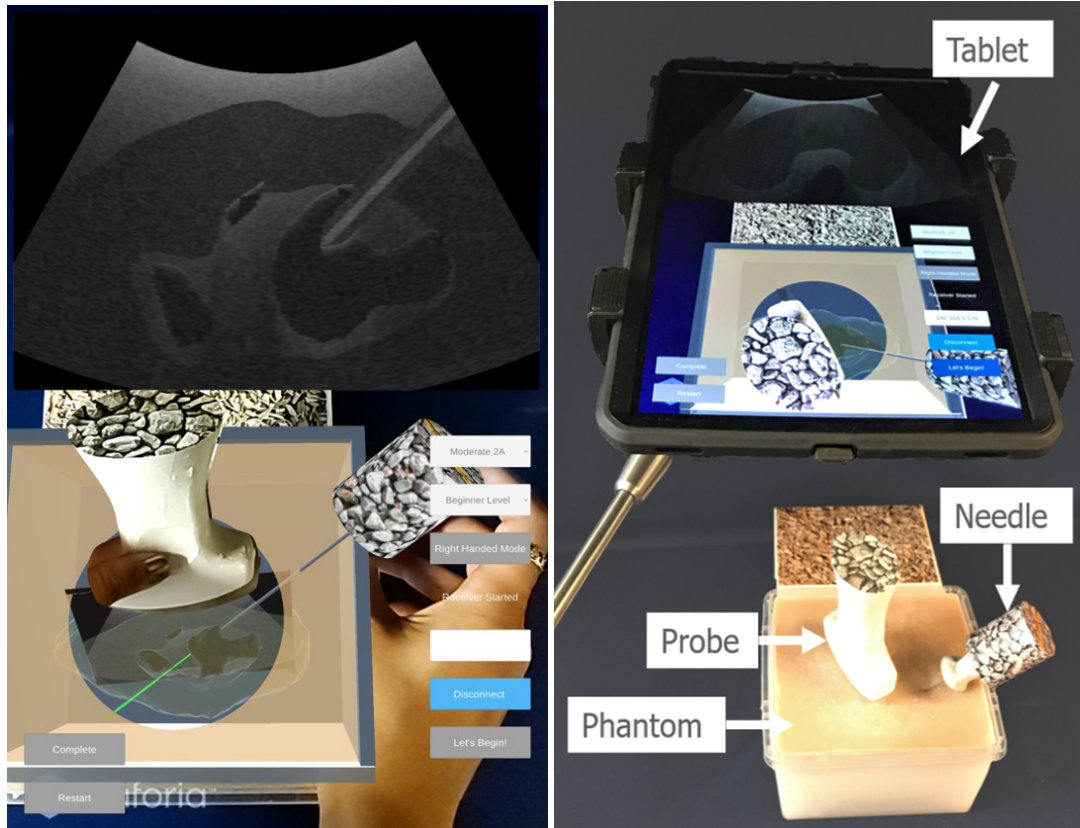


Figure 2.1: AR visualization and system hardware. (Left) The top section displays real-time 2D US simulation and the bottom section displays the 3D AR scene of PCA with full AR assistance. (Right) The hardware setup including a mounted tablet, a 3D printed probe emulator, a needle, and a silicone phantom. Image used with the permission of Springer Nature.

2.2.3 Software

Segmentation and three-dimensional (3D) model construction

The 3D kidney models were generated retrospectively from diagnostic nephrostomy computed tomography (CT) studies stored in a PACS database. Recent Nephrostomy tube insertion cases were reviewed by one of the interventional radiologists along with preceding CT studies. Five cases were selected based on hydronephrosis grade including one mild, two moderate, and two severe cases. Hydronephrosis is the swelling of a kidney caused by a blockage that doesn't allow proper urine drainage. The segmentation of the renal parenchyma and collecting system was performed by the same radiologist using TeraRecon Aquarius iNtuition ver.4.4.13.P2 (TeraRecon Inc., Foster City, CA, USA). The parapelvic fat, renal masses, and cysts were excluded from the segmentation and the proximal ureter was included only to the level of the lower pole of the kidney. The gross contour of the kidney was manually interpolated across these structures. Finally, the segmented volumes were converted to meshes and exported as .stl files.

Vuforia tracking

Infrared optical, electromagnetic, and GPS tracking systems are well-established for real-time tracking of surgical instruments in surgical navigation for their high accuracy. However, due to their complex setup and high cost, training simulators employing these tracking systems are not widely adopted in medical schools. As AR applications increase in popularity, computer vision-based camera tracking has become an active research area. Vuforia software development kit (SDK) (PTC Inc., USA) is licensed software that uses its proprietary computer vision-based image recognition algorithm to track 2D images or 3D objects in real-time by detecting sharp corners and edges from camera images. Based on detected features, the position and orientation of the marker in the virtual space are computed using affine transformation. Three unique, non-repetitive Vuforia markers were used to estimate the locations and orientations of

the needle, the US probe emulator, and the silicone phantom. To use Vuforia marker, a digital copy of the image needs to be uploaded to Vuforia server for feature detection (Figure 2.2). The digital markers were cropped and adjusted to ensure good visibility of texture details under the camera, which improved the performance of feature recognition and the robustness of object tracking. The physical dimensions of the printed markers were measured using a caliper then the processed marker data were imported into Unity (Unity Technologies, USA) at 1:1 scaling ratio. 3D virtual models of the needle, the US probe emulator and the silicone phantom were registered to the markers in the Unity. Once a marker was identified, the corresponding virtual object appears on the screen based on the pose of the marker. The dimensions of virtual objects are the same as the physical models. The data stream, referred to as the tracker stream in this chapter, containing positions and orientations of these virtual objects was packaged in Unity and then used for US simulation. The tracking precision of the needle marker was tested on a 20×20 cm grid placed 30 cm away from the tablet camera at 45° angle to simulate the tracking area during training. 225 tracking positions were recorded at various needle poses angled within 30° from the grid surface normal. The tracking error is consistently low over the entire tracking area with a mean and max error of 1.7 mm and 3.5 mm respectively.

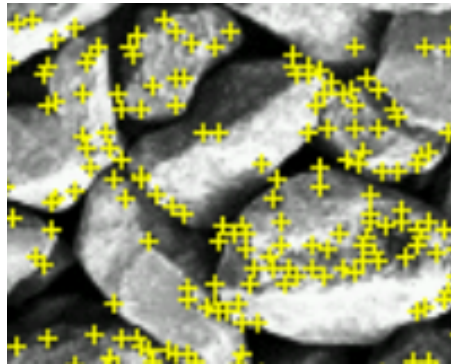


Figure 2.2: An simplified result of Vuforia feature detection.

Simulation workflow

An overall deployment diagram is shown below in Figure [2.3](#). In this project, we used the previously validated public software library for US imaging research (PLUS) for real-time US image simulation [\[11\]](#). PLUS accepts the tracker stream and the mesh models of the kidney and the needle as inputs to compute individual US scanline based on the pose of the probe. Acoustic properties of each mesh model, including attenuation, absorption, and reflection, were modified in the configuration file (Appendix A.) to achieve the most realistic simulation effect. However, this algorithm does not simulate characteristics of real ultrasound images such as speckle and reverberation. It is important to set up a proper environment for the kidney model in the configuration file to simulate the US effect of kidney In Vivo. Therefore, layers of material were configured such as air, a working volume of gel, the kidney shell, and the collecting system. Similarly, a needle would appear as a thin line on simulated US. Adding a concentric tube over the needle model created a halo effect which is more realistic. To optimize the processing performance of the tablet, PLUS was configured to run on a separate PC.

Data transfer between the tablet and the PC was achieved through a communication plug-in built for Unity, which is a modified version of open image-guided therapy link (OpenIGTLink). Since the original OpenIGTLink was written in C/C++, while our Unity application was written in C#, marshaling methods (Appendix B.) were used to write the Unity implementation to transfer data between managed and unmanaged codes. The modified code (Appendix C. & D.) was compiled to a dynamic-link library (DLL) in Visual Studio. However, to create plug-in that works on Android devices, an extra step is required to cross-platform compile our code into shared object library (.SO) using Android Studio. Once the communication is established between the tablet and the PC, the tracker stream is transmitted to PLUS for US simulation, and the simulated images are sent back to our Unity application in raw data format.

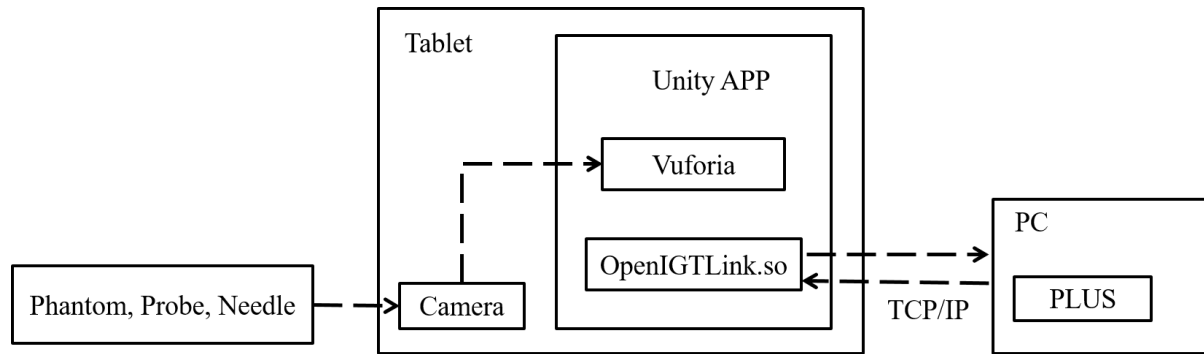


Figure 2.3: Simulation workflow diagram. Image used with the permission of Springer Nature.

AR visualization

To simulate the PCA training environment, we imported models of the 3D kidney, the US probe, and the virtual needle into a Unity scene. Nevertheless, when rendering an AR scene, virtual objects were always drawn on top of real objects, which caused the kidney to appear floating above the silicone phantom. To correct this depth perception problem, we placed the virtual kidney model inside a virtual box, that was manually registered to the phantom container. The illusion of a hollow box can be created by applying depth mask outside of the hollow area (colored in blue), to lower the render queue of the virtual walls (Figure 2.4). A virtual skin layer with a keyhole cutout was placed to help users focus on the region of interest (ROI) while providing additional depth cues. The virtual skin layer was manually registered to the surface of the phantom using the needle tip. A similar floating rendering effect also affects the US probe. While using the probe, the user's hand could be completely occluded by the virtual probe model, creating a disconnect between the real world and the virtual one. We applied the same depth mask technique to the virtual probe model to lower the render queue of the virtual probe model. As a result, the camera feed of the probe was drawn first, which allows the user to see part of the hand holding the probe.

Three training levels (beginner, intermediate and advanced) were designed for users with different skill levels. In the beginner level, a small US image plane was attached below the probe, which helps the user understand the 2D US image within the context of the 3D kidney

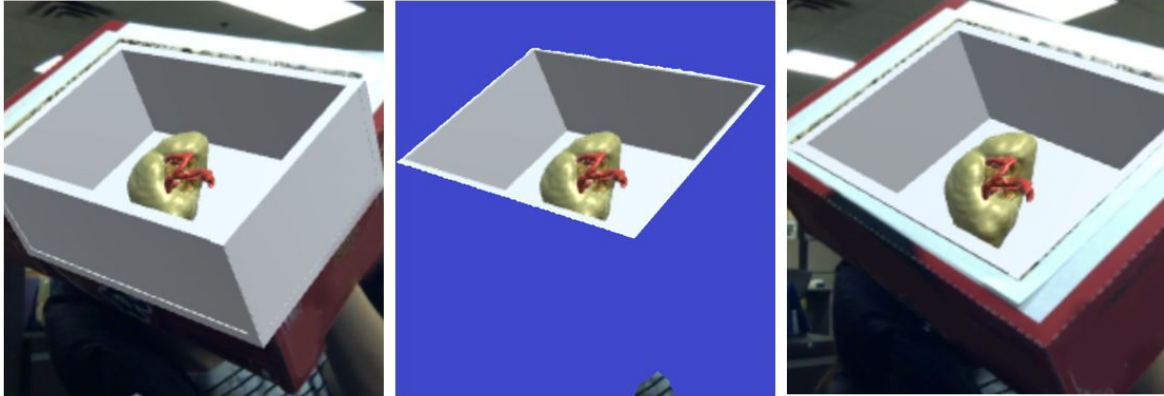


Figure 2.4: Demonstration of depth mask effect. Left: The wall of the hollow white box were visible. The white box appears to be floating on top of the red box from camera feed. Middle: Depth mask was applied to the blue area. Right: The walls of the red box from camera feed was drawn before the virtual white box.

model. In addition, the needle trajectory was also provided to the user, shown as a green projection line attached to the needle tip, serving as a visual aid for in-plane needle alignment (Figure 2.1 Left). Once the user had developed a good understanding of the anatomy and US images, the virtual kidney model was switched off in the intermediate level, while the augmented needle trajectory was still available. Finally, all the virtual aids, including the virtual kidney model, the augmented needle trajectory, and the US image plane below the probe were switched off at the advanced level. The users were expected to complete the task at the advanced level relying solely on the US display.

2.2.4 Simulator Training Procedure

To perform a PCA effectively, a user first inspects the US scans to identify the target inside the virtual kidney (the posterior calyx of the lower pole), under the simulated US guidance. Careful planning should be undertaken to achieve a direct needle access with the shortest path. Once the needle insertion path is planned, the user begins to insert an 18 gauge needle into the phantom, and guide it towards the target. As the needle advances inside the virtual kidney, it is crucial to maintain the in-plane view of the needle in the US image, in order to visualize the needle tip position. Losing track of the needle tip could cause severe complications by

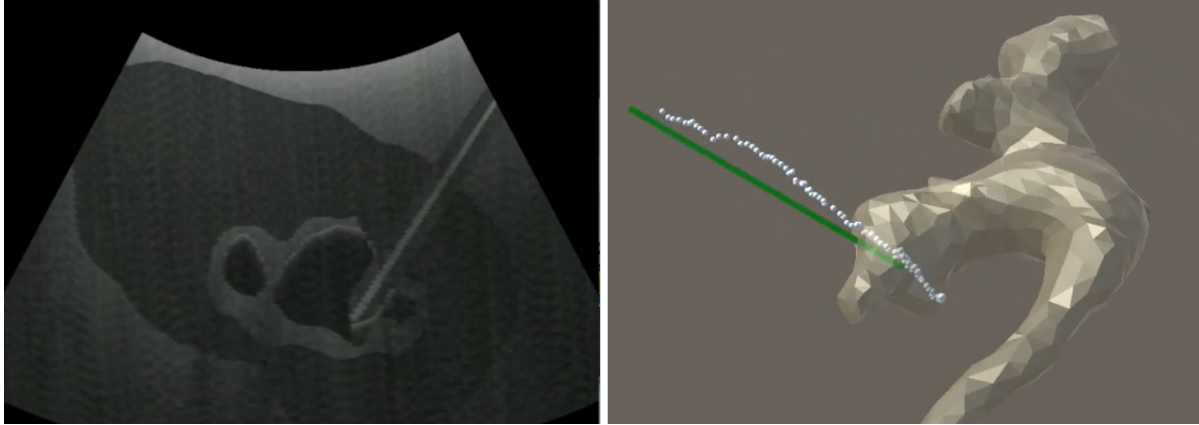


Figure 2.5: Screenshot of the result scene displaying the optimum access trajectory in green line and the user needle trajectory in white dotted line. Image used with the permission of Springer Nature.

inadvertently injuring critical tissue structures. Once the target area is reached, the user presses the ‘Complete’ button to terminate the simulation and review the needle access trajectory.

For user performance evaluation, a cone-shaped target region was manually registered to each virtual kidney, of which the central line was defined as the optimal needle access trajectory. This cone-shaped target region remains invisible to the users unless the ‘Show Hint’ button was pressed. The actual needle trajectory was recorded automatically once the needle tip pierces through the skin layer of the silicone phantom at a rate of 10 records per second. After the ‘Complete’ button is pressed, the application switches to the result scene where users receive intuitive visual feedback of their performance. An overlay of the optimal needle access trajectory (green line), the actual needle trajectory (white dots), and the transparent collecting system model are displayed in the result scene (Figure 2.5).

2.2.5 Evaluation Metrics

The following seven metrics are recorded by the simulator to evaluate user performance.

1. Final distance to target: the distance from the final needle tip position to the optimum target position.
2. Total path length: the total distance that the needle tip traveled in the phantom.

3. Total time: the total time from the moment that the needle tip entered the phantom until ‘Complete’ button was pressed
4. Overall adaptive distance: the weighted Euclidean distance from the actual needle path to the optimal trajectory (2.1), where N represents the total number of data points, w_i represents the adaptive weight based on the needle insertion depth: $1 - e^{-(depth/0.3)}$, and d represents the Euclidean distance between the needle tip position and the optimal trajectory.

$$WD = \frac{1}{N} \cdot \sum_{i=1}^N w_i \cdot d_i \quad (2.1)$$

5. Needle shaft visualization time: the total time that part of the needle shaft is visible on the US image
6. Needle tip visualization time: the total time that the needle tip is visible on the US image
7. Inside collecting system or not: a binary status indicating whether the final needle tip position is inside the targeted collecting system.

2.3 Validation

To integrate a simulator into the training curriculum, its validity must be proven through an established evaluation process including face, content, construct, concurrent, and predictive validity assessment [10][27][79]. Face validity examines the resemblance between a simulator and the actual procedure, and can be assessed using user survey. Content validity evaluates the educational content of a simulator, and it should be assessed by experts in the field. Construct validity examines whether the simulation performance of a user reflects the actual skill level, which can be assessed by comparing performance of an expert group with a non-expert group. Concurrent validity compares the simulation training effect to an established training method. Finally, predictive validity examines if performance in the simulated environment can

translates to the real world. This can be assessed by correlating simulation performance with the operating room performance. For this study, we focused on the evaluation of face, content, and construct validity.

2.3.1 Experimental Protocol

To validate the effectiveness of our training simulator, we performed a single-centre user study approved by the Health Sciences Review Ethics Board (114409) of Western University. For this study, we enrolled 24 novices and 6 experts on a voluntary basis. The novices consisted of 24 post-graduate medical students with no previous experience in PCA. The experts consisted of urologists and interventional radiologists with over five years of experience in PCA.

For the novices, we started with a five minutes introductory tutorial, which covered the rationale of this study, renal anatomy, targeting, basic US scanning techniques, and AR tracking. When the participants were comfortable with the simulator, the training and testing procedures began. The training and testing procedures were split into three phases including the pre-training testing, training, and post-training testing phases. During the pre- and post-training testing, the simulation was run at the advanced level, which meant no AR assistance was available for the participants. The procedure for pre- and post-training testing were identical. A single PCA procedure was performed by the novices on each of the three kidney models with different grades of hydronephrosis: one severe, one moderate, and one mild case. The evaluation metrics were recorded automatically by the application. In addition, each performance was video recorded and evaluated by two blinded experts using a modified global rating scale (GRS), which includes target identification, US needle tracking, economy of motion, and ability to perform needle access [82]. GRS is a survey tool which has been used to assess clinical competence [36]. During the training phase, the novices had the option to choose the level of AR assistance as they need. One moderate and one mild hydronephrosis cases were used for practicing, which were excluded from the testing cases. Participants were given 15 minutes to practice and ask questions.

Since the experts did not require training, they were given a short introduction of our simulator, followed by five minutes period to familiarize themselves with the system. The PCA procedures were performed by the experts following the same testing protocol for novices. After the session, a standard 5-point Likert scale (1 = strongly disagree and 5 = strongly agree) questionnaire was administered to the experts to assess the face and content validity.

2.3.2 Evaluation Method

First, the face and content validity was evaluated through questionnaires administered to the experts. Second, the construct validity was determined based on the distinguishability of a list of evaluation metrics collected from both novices and experts during needle insertion tasks, where the distinguishability is evaluated using the Mann-Whitney U Test. In addition, we also evaluated the training effectiveness of our simulator based on PCA skill acquisition. The PCA skill acquisition was determined by examining the change in performance of each novice participant based on the parameters collected before and after the training phase, with the paired T-Test being used to analyze the significance of this change. All the statistics were computed using Matlab R2018b.

2.3.3 Results

Face and content validity

The mean scores of face validity statements are detailed in Table 1. In summary, all experts rated the simulator as a 4 or 5 in all categories including the overall realism of the US image, the identification process, and the needle insertion procedure, except for one 3 rating of the needle insertion procedure, achieving an overall average score of 4.39 out of 5 (Table 2.1). In terms of content validity, experts strongly agreed that our simulator is able to teach the basic techniques to perform PCA, such as understanding of the basic anatomy, US interpretation, needle insertion planning, and the in-plane needle alignment with US beam, achieving an overall average

Table 2.1: Mean scores of face and content validity (range 1-5)

| Face validity | Expert (n=6) |
|--|---------------------|
| Realism of the US images | 4.39 (4, 5) |
| Realism of the identification process | 4.33 (4, 5) |
| Realism and usefulness of the needle insertion procedure | 4.44 (3, 5) |
| Content validity | Expert (n=6) |
| Basic renal anatomy | 4.80 (4, 5) |
| Ultrasound images interpretation | 4.80 (4, 5) |
| Safe planning | 4.50 (4, 5) |
| In-plane needle-beam alignment | 4.50 (4, 5) |
| Perform PCA | 4.67 (4, 5) |
| Overall value of the simulator as a training tool | 4.16 (4, 5) |
| Overall value of the simulator as a assessment tool | 4.33 (4, 5) |
| Recommend this simulator to others | 4.67 (4, 5) |
| Use this simulator in your training program | 4.33 (4, 5) |

score of 4.65 out of 5. Furthermore, experts were extremely satisfied with the overall value of this simulator as a training and assessment tool. They are highly likely to recommend this system to others and use it in their training program.

Construct validity

As depicted in Table 2, experts significantly outperformed novices on 6 of 7 evaluation metrics on the simulator ($p < 0.05$). Compared to novices, experts were able to insert the needle closer to the optimum location and optimum trajectory, while achieving a shorter needle path length, a shorter completion time, more accurate needle tip tracking, and a 100 percent success rate to insert the needle inside the collecting system for all trials. However, there were no significant differences in needle shaft visualization time between experts and novices (0.89s vs 0.81s, respectively). We suspect that partially visualizing the needle shaft does not contribute to the overall performance of PCA.

Table 2.2: Construct validity

| Construct validity | Novice (n=24) | Expert (n=6) | p-value |
|-------------------------------------|----------------------|---------------------|----------------|
| Final distance to target (mm) | 18.5 ± 8.1 | 8.7 ± 3 | 0.005 |
| Total path length (mm) | 326.1 ± 166.6 | 175.1 ± 37.7 | 0.036 |
| Total time (s) | 36.7 ± 23.5 | 17.9 ± 5.8 | 0.036 |
| Overall adaptive distance (mm) | 15.6 ± 5.7 | 9.8 ± 3.2 | 0.031 |
| Needle shaft visualization time (%) | 78 ± 20 | 89 ± 9 | 0.154 |
| Needle tip visualization time (%) | 43 ± 19 | 66 ± 11 | 0.006 |
| Inside collecting system or not (%) | 60 ± 43 | 100 | 0.026 |

[†]Data in this table are represented in mean value ± standard deviation. Statistical differences were calculated with the Mann-Whitney U test. A p-value < 0.05 was considered significant.

Acquisition of US-guided PCA skills

The PCA performances of the novice group was assessed subjectively and objectively both before and after training on our simulator. The objective assessments were based on multiple elements such as operation time, distance, needle visualization, and needle access accuracy, as shown in Table 3. The novice participants demonstrated statistically significant improvements in most categories after training, except for the total time and needle shaft visualization, where the improvements were not significant. The subjective assessments of novice performance were made using a global rating scale (GRS) (range 1 - 5) in four categories, as shown in Table 4. Similar to the objective assessments, the novice participants demonstrated significant improvement in all categories after training.

2.4 Discussion

The introduction of medical simulators has brought promising opportunities for training medical professionals outside of operating room (OR), in a safe and stress-free environment [10]. To overcome the common challenges in terms of cost and accessibility, we developed an affordable and effective training simulator for PCA utilizing portable devices and AR. Using Vuforia, a computer vision-based camera tracking SDK, we are able to track the needle, the US probe

Table 2.3: Novice performance pre-training vs. post-training

| Novice (n=24) | Pre-training | Post-training | p-value |
|-------------------------------------|---------------------|----------------------|----------------|
| Final distance to target (mm) | 18.5 ± 8.1 | 8.7 ± 3.2 | <0.0001 |
| Total path length (mm) | 326.1 ± 166.6 | 263.9 ± 142.7 | 0.04 |
| Total time (s) | 36.7 ± 23.5 | 30 ± 14.5 | 0.11 |
| Overall adaptive distance (mm) | 15.6 ± 5.7 | 9 ± 2.5 | <0.0001 |
| Needle shaft visualization time (%) | 78 ± 20 | 81 ± 13 | 0.446 |
| Needle tip visualization time (%) | 43 ± 19 | 62 ± 15 | 0.0002 |
| Inside collecting system or not (%) | 60 ± 43 | 94 ± 13 | <0.0001 |

† Data in this table are represented in mean value ± standard deviation. Statistical differences were calculated with the paired T-test. A p-value<0.05 was considered significant.

Table 2.4: GRS (range 1-5) assessments of novice performance before and after training

| Novice (n=24) | Pre-training | Post-training | p-value |
|----------------------------------|---------------------|----------------------|----------------|
| Identify Target | 3.4 ± 1.2 | 4.5 ± 0.6 | <0.0001 |
| US Needle Tracking | 2.3 ± 1.2 | 3.6 ± 1 | <0.0001 |
| Economy of Motion | 3.1 ± 1.3 | 3.6 ± 1 | 0.041 |
| Ability to Perform Needle Access | 2.3 ± 1.2 | 3.7 ± 1 | <0.0001 |

† Statistical differences were calculated with Wilcoxon signed-rank test. A p-value<0.05 was considered significant.

emulator, and the silicone phantom through the camera video stream in real-time without the need for any external tracking hardware. As a result, the estimated cost of this simulator can be kept below \$100.

In general, the effectiveness of a training simulator is demonstrated through face, content, construct, concurrent, and predictive validities [10][27][79]. Following this convention, we first established face and content validity. The high face and content validity scores demonstrate that our simulator closely resembles an actual US-guided PCA procedure and that the skills required for an successful US-guided PCA using our simulator was similar to the skills required for an actual US-guided PCA. Next, construct validity was clearly indicated by the ability of our simulator to distinguish between novices and experts with a high degree of confidence for 6 out of 7 objective evaluation metrics except for the needle shaft visualization time. In addition, our experiment showed significant improvements in the US-guided PCA skills of the novice participants, which is demonstrated by the consistent increase in both objective evaluation metrics and subjective GRS in post-training testing. Compared with the objective evaluation metrics in pre-training testing, the final distance to the target, the overall adaptive distance, and the needle tip visualization time improved over 40% in post-training testing, which became very close to the experts' performance.

According to the user feedback, all novices agreed that our simulator is easy to use and useful to improve their PCA skills, and they would like to have the option to practice at home using our simulator. The expert group was satisfied with the overall value of this simulator as a training (4.16 out of 5) and assessment (4.33) tool, and they are highly likely to recommend this system to others (4.67) and use it in their training program (4.33), as shown in Table 1. The major area of improvement that the experts have suggested is the occasional tracking instability.

2.5 Conclusions

Our proposed AR training simulator can provide educators with an alternative to allow for additional training opportunities for trainees to acquire basic skills of US-guided PCA in a safe and stress-free environment. The effectiveness of our simulator is demonstrated through face, content, and construct validities. In addition, pre-post testing comparisons showed significant improvements in both the objective and subjective performances of our novice trainees. To further improve the objective evaluation of user performance, our next step is to design an overall scoring system combining the evaluation metrics, which will allow us to assess the concurrent validity of our simulator. In future iterations, with the advancement of mobile computing technology, we would like to incorporate realistic lighting and shadows, as well as tissue deformation characteristics, to further improve the depth perception and realism of our simulator.

Chapter 3

Conclusions and Future Work

3.1 Conclusions

Simulation-based training facilitate the acquisition and refinement of essential skills for percutaneous renal access (PCA) prior to clinical exposure by providing opportunities for deliberate and repetitive practice. The evaluation metrics from our simulator would provide more consistent objective feedback for targeted training of PCA skills and shorten the learning curve during early training, as demonstrated through the substantial performance improvements over a short amount of training time in the novice group. In addition, our simulator offers different levels of augmented reality (AR) assistance, which allows for more customizable, self-directed, and progressive training with increasing difficulty. Aside from skill enhancements, our simulator has a simple setup, which offers the flexibility to train at home. This not only protects patients from unnecessary harm but also provides trainees a stress-free environment for repetitive computer-guided practice. Since the visualization of patient pathology is implemented through virtual modeling, our simulator has the capacity to include a large dataset with a wide variety of clinical scenarios. In addition, the presented ultrasound (US)-guided needle insertions simulation system, utilizing the versatility of AR technology, can be extended to other clinical applications such as spinal needle insertion or central line insertion

The computer vision-based camera tracking for AR has distinct advantages in cost and portability. However, the performance can be sensitive to varying lighting conditions. During the experiment, we noticed mild tracking instability, which could be caused by the internal tracking algorithm and external lighting artifacts such as specular highlights [102]. Therefore, moderately bright and diffused lighting should be provided for the best user experience.

The haptic feedback can be another useful feature to improve the realism of our simulator. Nevertheless, studies have shown that haptic feedback is less reliable compared with visual feedback and may not be necessary for real-time image-guided needle insertion [29]. Hence, our goal is to focus on training the users to perform PCA solely relying on the US images without any AR assistance nor the sensation to confirm calyx puncture. In addition, our general purposed phantom provides us the freedom to simulate patient-specific scenarios without fabricating various physical phantoms, and thus improving the versatility and cost-effectiveness.

3.2 Future Work

As the presented training simulator is a preliminary design, more refinements are needed maximize its benefits. As we mentioned in section 1.1.5, PCA training can be split into two sections: diagnostic renal imaging and needle control. It would be beneficial to create two corresponding training modules with virtual guidance such as animations and prompts, as well as progress tracking, to provide a structured learning experience and the sense of accomplishment. Additionally, we can apply elements of gamification such as points and leaderboards to increase user engagement. Currently, user performance feedback is provided in the form of direct visualization of the needle path, and data sheets with recorded performance parameters. Thus, to calculate the total point, a marking scheme needs to be developed first to determine the weight of each performance parameter.

Another direction of improvement relies on technology advancement such as depth sensing cameras and the computing power of mobile devices. As mentioned previously, the newer gen-

eration of mobile devices feature depth sensing cameras which can be incorporated into camera tracking algorithms to further improve the tracking accuracy and stability. Improvements in visual fidelity including depth perception and real-time tissue deformation can be implemented, given a powerful mobile chipset.

In this preliminary study, we have successfully established face, content, and construct validity through a user study described in chapter two. The next step is to establish concurrent validity by comparing the performance against a gold standard. Lastly proof of correlation between simulator performance and operating room (OR) performance can be used to demonstrate the predictive validity of this approach.

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Appendix A. Example of US Simulation Configuration File

```
<PlusConfiguration version="2.3">

  <DataCollection StartupDelaySec="1.0" >
    <DeviceSet
      Name="PlusServer For Unity local Moderate 2"
      Description="Simulated Ultrasound"
    />
    <!-- Image marker tracker-->
    <Device
      Id="TrackerDevice"
      Type="OpenIGTLinkTracker"
      MessageType="TRANSFORM"
      ToolReferenceFrame="Tracker"
      ServerAddress="127.0.0.1"
      ServerPort="18946"
      AcquisitionRate="90"
      IgtlMessageCrcCheckEnabled="true"
      UseReceivedTimestamps="false"
      UseLastTransformsOnReceiveTimeout="true"
      LocalTimeOffsetSec ="0.0"
      ReconnectOnReceiveTimeout="true">
      <DataSources>
        <DataSource Type="Tool" Id="Probe" />
        <DataSource Type="Tool" Id="Needle" />
        <DataSource Type="Tool" Id="Kidney" />
        <DataSource Type="Tool" Id="Gelblock" />
      </DataSources>
      <OutputChannels>
        <OutputChannel Id="TrackerStream">
          <DataSource Id="Probe" />
          <DataSource Id="Needle" />
          <DataSource Id="Kidney" />
          <DataSource Id="Gelblock" />
        </OutputChannel>
      </OutputChannels>
    </Device>
    <!-- For getting images from the US simulator -->
    <Device
      Id="VideoDevice"
      Type="UsSimulator"
      LocalTimeOffsetSec="0.0"
```

```

AcquisitionRate="90" >
<DataSources>
  <DataSource Type="Video" Id="Video" PortUsImageOrientation="MF" />
</DataSources>
<InputChannels>
  <InputChannel Id="TrackerStream" />
</InputChannels>
<OutputChannels>
  <OutputChannel Id="VideoStream" VideoDataSourceId="Video" />
</OutputChannels>
<vtkPlusUsSimulatorAlgo
  ImageCoordinateFrame="Image"
  ReferenceCoordinateFrame="Reference"
  IncomingIntensityMwPerCm2="300"
  BrightnessConversionGamma="0.25"
  BrightnessConversionOffset="30"
  NumberOfScanlines="128"
  NumberOfSamplesPerScanline="1000"
  NoiseAmplitude="14.0"
  NoiseFrequency="2.5 3.5 1"
  NoisePhase="50 20 0"
  >
  <SpatialModel Name="Air"
    DensityKgPerM3="1.2"
    SoundVelocityMPerSec="343"
    AttenuationCoefficientDbPerCmMhz="100.0"
    BackscatterDiffuseReflectionCoefficient="0.1"
    SurfaceReflectionIntensityDecayDbPerMm="50"
  />
  <SpatialModel
    Name="Gelblock"
    Type="Model"
    ObjectCoordinateFrame="Gelblock"
    ModelFile="D:/AllModels/CubeModel_m.stl"
    DensityKgPerM3="910"
    SoundVelocityMPerSec="1540"
    AttenuationCoefficientDbPerCmMhz="3.0"
    BackscatterDiffuseReflectionCoefficient="0.1"
    SurfaceSpecularReflectionCoefficient="0.0"
    SurfaceDiffuseReflectionCoefficient="0.0"
    TransducerSpatialModelMaxOverlapMm="50"
  />
  <SpatialModel
    Name="Kidney"
    ObjectCoordinateFrame="Kidney"

```

```

    ModelFile="D:/AllModels/ModerateKidney2_m.stl"
    DensityKgPerM3="1066"
    SoundVelocityMPerSec="1570"
    AttenuationCoefficientDbPerCmMhz="1.0"
    BackscatterDiffuseReflectionCoefficient="0.001"
    SurfaceSpecularReflectionCoefficient="0.0"
    SurfaceDiffuseReflectionCoefficient="0.0" />
<SpatialModel
  Name="Calyces"
  ObjectCoordinateFrame="Kidney"
  ModelFile="D:/AllModels/ModerateCollecting2_m.stl"
  DensityKgPerM3="1201"
  SoundVelocityMPerSec="2200"
  AttenuationCoefficientDbPerCmMhz="0"
  BackscatterDiffuseReflectionCoefficient="0.000"
  SurfaceSpecularReflectionCoefficient="0.0"
  SurfaceDiffuseReflectionCoefficient="0.3"
/>
<!--SpatialModel
  Name="Vessel"
  ObjectCoordinateFrame="Kidney"
  ModelFile="D:/Artery_m.stl"
  ModelToObjectTransform="
    1 0 0 0
    0 1 0 0
    0 0 1 0
    0 0 0 1"
  DensityKgPerM3="1102"
  SoundVelocityMPerSec="2000"
  AttenuationCoefficientDbPerCmMhz="0.01"
  BackscatterDiffuseReflectionCoefficient="0.001"
  SurfaceSpecularReflectionCoefficient="0.0"
  SurfaceDiffuseReflectionCoefficient="0.0"
/-->

<SpatialModel
  Name="Needle"
  ObjectCoordinateFrame="NeedleTip"
  ModelFile="D:/AllModels/NeedleModel.stl"
  DensityKgPerM3="2500"
  SoundVelocityMPerSec="5000"
  AttenuationCoefficientDbPerCmMhz="3.0"
  BackscatterDiffuseReflectionCoefficient="1"
  SurfaceSpecularReflectionCoefficient="0.0"
  SurfaceDiffuseReflectionCoefficient="0.3"

```

```

    SurfaceReflectionIntensityDecayDbPerMm="0.1"
  />
<SpatialModel
  Name="Needle"
  ObjectCoordinateFrame="NeedleTip"
  ModelFile="D:/AllModels/NeedleModel_Double_6mm.stl"
  DensityKgPerM3="1600"
  SoundVelocityMPerSec="5000"
  AttenuationCoefficientDbPerCmMhz="0.03"
  BackscatterDiffuseReflectionCoefficient="0.9"
  SurfaceSpecularReflectionCoefficient="0.0"
  SurfaceDiffuseReflectionCoefficient="0.0"
  SurfaceReflectionIntensityDecayDbPerMm="0"
/>
<RfProcessing>
  <ScanConversion
    TransducerName="Ultrasonix_C5-2/60"
    TransducerGeometry="CURVILINEAR"
    ModelToObjectTransform="
      1 0 0 0
      0 1 0 0
      0 0 1 0
      0 0 0 1"
    RadiusStartMm="60.0"
    RadiusStopMm="130.0"
    ThetaStartDeg="-28.0"
    ThetaStopDeg="28.0"
    TransducerCenterPixel="410 100"
    OutputImageSizePixel="820 616"
    OutputImageSpacingMmPerPixel="0.13 0.13" />
  </RfProcessing>
  <!--
- Image size pixel change background black size small: smaller black window -->
  <!--
- Image pixel spacing change ultrasound simulation size larger spacing: smaller
US image affects resolution -->
  </vtkPlusUsSimulatorAlgo>

</Device>

<Device
  Id="TrackedVideoDevice"
  Type="VirtualMixer" >
  <InputChannels>
    <InputChannel Id="TrackerStream" />

```

```

    <InputChannel Id="VideoStream" />
</InputChannels>

<OutputChannels>
    <OutputChannel Id="TrackedVideoStream"/>
</OutputChannels>
</Device>

<!--Device
    Id="CaptureDevice"
    Type="VirtualCapture"
    BaseFilename="RecordingTest.igs.mhd"
    EnableCapturingOnStart="FALSE" >
    <InputChannels>
        <InputChannel Id="TrackedVideoStream" />
    </InputChannels>
</Device-->
</DataCollection>

<PlusOpenIGTLinkServer
    MaxNumberOfIgtlMessagesToSend="10"
    MaxTimeSpentWithProcessingMs="50"
    ListeningPort="18944"
    SendValidTransformsOnly="true"
    OutputChannelId="TrackedVideoStream" >
<DefaultClientInfo>
    <MessageTypes>
        <Message Type="IMAGE" />
        <Message Type="TRANSFORM" />
    </MessageTypes>
    <TransformNames>
        <!--These transforms becomes active transforms in slicer-->
        <Transform Name="GelblockToReference" />
        <Transform Name="NeedleTipToReference" />
        <Transform Name="ProbeToReference" />
        <Transform Name="KidneyToReference" />
    </TransformNames>
    <ImageNames>
        <Image Name="Image" EmbeddedTransformToFrame="Reference" />
    </ImageNames>
</DefaultClientInfo>
</PlusOpenIGTLinkServer>

<CoordinateDefinitions>
<!--scaling change 1xX -->

```



```
<Transform From="Image" To="Probe"  
Matrix="  
  0.13 0 0 -53.3  
  0 0 0.13 0  
  0 -0.13 0 8  
  0 0 0 1" />
```

```
<Transform From="Image" To="TransducerOriginPixel"  
Matrix="  
  1 0 0 0  
  0 1 0 0  
  0 0 1 0  
  0 0 0 1" />
```

```
<Transform From="Tracker" To="Reference"  
Matrix="  
  1 0 0 0  
  0 1 0 0  
  0 0 1 0  
  0 0 0 1" />
```

```
<Transform From="NeedleTip" To="Needle"  
Matrix="  
  1 0 0 0  
  0 1 0 0  
  0 0 1 0  
  0 0 0 1" />
```

```
</CoordinateDefinitions>
```

```
</PlusConfiguration>
```

Appendix B. UNITY C# Marshalling

```
using System.Collections;
using System.Collections.Generic;
using UnityEngine;
using System.Runtime.InteropServices;
using UnityEngine.UI;
using System;
using System.IO;
using System.Net.Sockets;
using AOT;
using System.Threading;
using UnityEngine.SceneManagement;

public class UnityIGT : MonoBehaviour
{
    private delegate void ServerDelegate(int Svalue);
    [DllImport("OpenIGTLink")]
    static extern void ServerConnect(int serverport, double fps, ServerDelegate
ServerConnectionStatus);

    private delegate void ReceiverDelegate(int Rvalue);
    [DllImport("OpenIGTLink")]
    static extern void ReceiverConnect(string hostname, int clientport, ReceiverDelegate
ReceiverConnectionStatus);

    private delegate void PositionDelegate(float PositionX);
    [DllImport("OpenIGTLink")]
    static extern void GetData(float[] orientation, int Osize, float[] position, int
Psize, PositionDelegate SentPosition);

    private delegate void GetImageDelegate(IntPtr Address);
    [DllImport("OpenIGTLink")]
    static extern void GetImage(GetImageDelegate GetImageData);

    private delegate void CloseSendDelegate();
    [DllImport("OpenIGTLink")]
    static extern void CloseSendSocket();

    private delegate void CloseReceiveDelegate();
    [DllImport("OpenIGTLink")]
    static extern void CloseReceiveSocket();

    public GameObject[] TracketObject;
    float[] ObjectPosition = new float[12];
    float[] ObjectOrientation = new float[16];
    public static GameObject MySprite;
    public static GameObject MySpriteSM;
    public InputField EnterIP;
```

```

string IPAddress;
bool startR = false;
bool startS = false;
static bool Received = false;
public static byte[] ImageData;
public static Texture2D ImageTexture;
Thread ReceiveThread;
Thread SendThread;

public static Text Statustext; //disconnect/connect/receiver

void Start()
{
    MySprite = GameObject.FindWithTag("Image");
    MySpriteSM = GameObject.FindWithTag("SmallImage");
    Application.targetFrameRate = 60;
    ImageData = new byte[505120];
}

void Update()
{
    for (int i = 0; i < 4; i++)
    {
        if (TracketObject[i].activeSelf == true)
        {
            //ObjectStatustext.text = "Object Found";
            ObjectPosition[i * 3 + 0] = TracketObject[i].transform.position.x;
            ObjectPosition[i * 3 + 1] = TracketObject[i].transform.position.y;
            ObjectPosition[i * 3 + 2] = TracketObject[i].transform.position.z;
            ObjectOrientation[i * 4 + 0] = TracketObject[i].transform.rotation.x;
            ObjectOrientation[i * 4 + 1] = TracketObject[i].transform.rotation.y;
            ObjectOrientation[i * 4 + 2] = TracketObject[i].transform.rotation.z;
            ObjectOrientation[i * 4 + 3] = TracketObject[i].transform.rotation.w;
        }
        else
        {
            //ObjectStatustext.text = "Object Lost";
            Array.Clear(array: ObjectPosition, index: 0, length:
ObjectPosition.Length);
            Array.Clear(array: ObjectOrientation, index: 0, length:
ObjectOrientation.Length);
        }
    }
    Resources.UnloadUnusedAssets();
}

public void StartServer()
{
    ServerConnect(18946, 60, ServerConnectionStatus);
}
public void StartClient()
{
    IPAddress = EnterIP.text;
    ReceiverConnect(IPAddress, 18944, ReceiverConnectionStatus);
}

```

```

}
public void AutoSend()
{
    InvokeRepeating("StartSendData", 0f, 0.033333f);
}
public void StartSendData()
{
    GetData(ObjectOrientation, 4, ObjectPosition, 3, SentPosition);
}

public void StartReceiveThread()
{
    startR = true;
    ReceiveThread = new Thread(AutoReceive);
    ReceiveThread.Start();
}
public void AutoReceive()
{
    while (startR == true)
    {
        StartReceivingImage();
    }
}
public void StartReceivingImage()
{
    GetImage(GetImageDataAddress);
}
public void InvokeUpdate()
{
    InvokeRepeating("UpdateTexture", 0f, 0.0333333f);
}
public void UpdateTexture()
{
    if (Received == true)
    {
        ImageTexture = new Texture2D(820, 616, TextureFormat.Alpha8, false);
        ImageTexture.LoadRawTextureData(ImageData);
        ImageTexture.Apply();
        MySprite.GetComponent<RawImage>().texture = ImageTexture;
        MySpriteSM.GetComponent<RawImage>().texture = ImageTexture;
    }
}

[MonoPInvokeCallback(typeof(ServerDelegate))]
private static void ServerConnectionStatus(int Svalue)
{
    if (Svalue == 1)
    {
        Statustext = GameObject.FindWithTag("Status").GetComponent<Text>();
        Statustext.text = "Connected to Server";
    }
}

[MonoPInvokeCallback(typeof(ReceiverDelegate))]
private static void ReceiverConnectionStatus(int Rvalue)
{
    if (Rvalue == 1)

```

```

    {
        Statustext = GameObject.FindWithTag("Status").GetComponent<Text>();
        Statustext.text = "Receiver Started";
    }
}
[MonoPInvokeCallback(typeof(PositionDelegate))]
private static void SentPosition(float PositionX) //verify plus received position
{
}
[MonoPInvokeCallback(typeof(GetImageDelegate))]
private static void GetImageDataAddress(IntPtr Address)
{
    Marshal.Copy(Address, ImageData, 0, 505120);
    Received = true;
    //RC = RC + 1;
}
public void LetsGo()
{
    AutoSend();
    StartClient();
    StartReceiveThread();
    InvokeUpdate();
}
public void Restart()
{
    ReceiveThread.Abort();
    SceneManager.LoadScene("UltrasoundSimulator_Standing_Sep");
}

public void Disconnect()
{
    startR = false;
    CloseReceiveSocket();
    CloseSendSocket();
    ReceiveThread.Abort();
}
private void OnApplicationQuit()
{
    ReceiveThread.Abort();
}
}

```

Appendix C. OpenIGTLink Based Sender

```
#include <iomanip>
#include <iostream>
#include <math.h>
#include <cstdlib>
#include <cstdlib>

#include "igtlOSUtil.h"
#include "igtlMessageHeader.h"
#include "igtlTransformMessage.h"
#include "igtlServerSocket.h"
#include "igtlClientSocket.h"
#include "igtlTrackingDataMessage.h"
#include "igtl_tdata.h"

#define UnityIGTLinkSendAPI _declspec(dllexport)

igtl::ServerSocket::Pointer UnityServerSocket;
igtl::Socket::Pointer UnitySendSocket;
igtl::TimeStamp::Pointer ts;

extern "C" {
    typedef void(_stdcall* CallbackServer)(int Svalue);
    UnityIGTLinkSendAPI void _stdcall ServerConnect(int serverport, double fps,
    CallbackServer ServerConnectionStatus)
        //UnityIGTLinkSendAPI void ServerConnect(int serverport, double fps,
    void(*ServerConnectionStatus)(int Svalue))
    {

        //int interval = (int)(1000.0 / fps);

        // Establish Connection
        igtl::ServerSocket::Pointer serversocket;
        serversocket = igtl::ServerSocket::New();
        int r = serversocket->CreateServer(serverport);
        UnityServerSocket = serversocket;
        UnitySendSocket = serversocket->WaitForConnection(10000);

        if (UnitySendSocket.IsNotNull())
        {
            ServerConnectionStatus(1);
        }

    }
    //-----
    -----

    void SendProbeToReference(igtl::Matrix4x4& Pmatrix, igtl::ServerSocket::Pointer
    serversocket, igtl::Socket::Pointer clientsocket)
    {
        // Allocate Transform Message Class
        igtl::TransformMessage::Pointer PTransformMessage =
    igtl::TransformMessage::New();

        //Package Data then send
    }
}
```

```

PTransformMessage->SetHeaderVersion(IGTL_HEADER_VERSION_1);
PTransformMessage->InitPack();
PTransformMessage->SetDeviceName("ProbeToTracker");
PTransformMessage->SetMatrix(Pmatrix);
ts = igt1::TimeStamp::New();
ts->GetTime();
PTransformMessage->SetTimeStamp(ts);
PTransformMessage->Pack();
UnitySendSocket->Send(PTransformMessage->GetPackPointer(),
PTransformMessage->GetPackSize());
    //igt1::Sleep(1); // wait
    // Close connection
    //socket->CloseSocket();
}
UnityIGTLLinkSendAPI void _stdcall CloseSendSocket()
{
    UnityServerSocket->CloseSocket();
    UnitySendSocket->CloseSocket();
}
//-----
void SendNeedleToReference(igt1::Matrix4x4& Nmatrix, igt1::ServerSocket::Pointer
serversocket, igt1::Socket::Pointer clientsocket)
{
    // Allocate Transform Message Class
    igt1::TransformMessage::Pointer NTransformMessage =
igt1::TransformMessage::New();
    //Package Data then send
    NTransformMessage->SetHeaderVersion(IGTL_HEADER_VERSION_1);
    NTransformMessage->InitPack();
    NTransformMessage->SetDeviceName("NeedleToTracker");
    NTransformMessage->SetMatrix(Nmatrix);
    ts = igt1::TimeStamp::New();
    ts->GetTime();
    NTransformMessage->SetTimeStamp(ts);
    NTransformMessage->Pack();
    UnitySendSocket->Send(NTransformMessage->GetPackPointer(),
NTransformMessage->GetPackSize());
    //igt1::Sleep(1); // wait
    // Close connection
    //socket->CloseSocket();
}
//-----
void SendKidneyToReference(igt1::Matrix4x4& Kmatrix, igt1::ServerSocket::Pointer
serversocket, igt1::Socket::Pointer clientsocket)
{
    // Allocate Transform Message Class
    igt1::TransformMessage::Pointer KTransformMessage =
igt1::TransformMessage::New();
    //Package Data then send
    KTransformMessage->SetHeaderVersion(IGTL_HEADER_VERSION_1);
    KTransformMessage->InitPack();
    KTransformMessage->SetDeviceName("KidneyToTracker");
    KTransformMessage->SetMatrix(Kmatrix);
    ts = igt1::TimeStamp::New();
    ts->GetTime();
    KTransformMessage->SetTimeStamp(ts);
    KTransformMessage->Pack();
}

```

```

        UnitySendSocket->Send(KTransformMessage->GetPackPointer(),
KTransformMessage->GetPackSize());
        //igt1::Sleep(1); // wait
        // Close connection
        //socket->CloseSocket();
    }
    void SendGelblockToReference(igt1::Matrix4x4& Gmatrix, igt1::ServerSocket::Pointer
serversocket, igt1::Socket::Pointer clientsocket)
    {
        // Allocate Transform Message Class
        igt1::TransformMessage::Pointer GTransformMessage =
igt1::TransformMessage::New();
        //Package Data then send
        GTransformMessage->SetHeaderVersion(IGTL_HEADER_VERSION_1);
        GTransformMessage->InitPack();
        GTransformMessage->SetDeviceName("GelblockToTracker");
        GTransformMessage->SetMatrix(Gmatrix);
        ts = igt1::TimeStamp::New();
        ts->GetTime();
        GTransformMessage->SetTimeStamp(ts);
        GTransformMessage->Pack();
        UnitySendSocket->Send(GTransformMessage->GetPackPointer(),
GTransformMessage->GetPackSize());
        //igt1::Sleep(1); // wait
        // Close connection
        //socket->CloseSocket();
    }
    //-----
    typedef void(_stdcall* CallbackPosition)(float ReceivedPosition);
    UnityIGTLLinkSendAPI void _stdcall GetData(float *Rorientation, int Osize, float
*Rposition, int Psize, CallbackPosition CheckPosition)
    //UnityIGTLLinkSendAPI void GetData(float *Rorientation, int Osize, float
*Rposition, int Psize, void(*CheckPosition)(float ReceivedPosition))
    {

        // build matrix
        igt1::Matrix4x4 Pmatrix;
        float Porientation[] =
{ Rorientation[0],Rorientation[1],Rorientation[2],Rorientation[3] };
        igt1::QuaternionToMatrix(Porientation, Pmatrix);
        Pmatrix[0][3] = Rposition[0];
        Pmatrix[1][3] = Rposition[1];
        Pmatrix[2][3] = Rposition[2];

        igt1::Matrix4x4 Nmatrix;
        float Norientation[] =
{ Rorientation[4],Rorientation[5],Rorientation[6],Rorientation[7] };
        igt1::QuaternionToMatrix(Norientation, Nmatrix);
        Nmatrix[0][3] = Rposition[3];
        Nmatrix[1][3] = Rposition[4];
        Nmatrix[2][3] = Rposition[5];

        igt1::Matrix4x4 Kmatrix;
        float Korientation[] =
{ Rorientation[8],Rorientation[9],Rorientation[10],Rorientation[11] };
        igt1::QuaternionToMatrix(Korientation, Kmatrix);
        Kmatrix[0][3] = Rposition[6];

```



```

    Kmatrix[1][3] = Rposition[7];
    Kmatrix[2][3] = Rposition[8];

    igtl::Matrix4x4 Gmatrix;
    float Orientation[] =
{ Rorientation[12],Rorientation[13],Rorientation[14],Rorientation[15] };
    igtl::QuaternionToMatrix(Gorientation, Gmatrix);
    Gmatrix[0][3] = Rposition[9];
    Gmatrix[1][3] = Rposition[10];
    Gmatrix[2][3] = Rposition[11];

    float receivedX = Nmatrix[0][3];
    CheckPosition(receivedX);

    SendProbeToReference(Pmatrix, UnityServerSocket, UnitySendSocket);
    SendNeedleToReference(Nmatrix, UnityServerSocket, UnitySendSocket);
    SendKidneyToReference(Kmatrix, UnityServerSocket, UnitySendSocket);
    SendGelblockToReference(Gmatrix, UnityServerSocket, UnitySendSocket);
}

//-----
-----

}

```

Appendix D. OpenIGTLink Based Receiver

```
#include <iomanip>
#include <iostream>
#include <math.h>
#include <cstdlib>

#include "igtlOSUtil.h"
#include "igtlMessageHeader.h"
#include "igtlTransformMessage.h"
#include "igtlServerSocket.h"
#include "igtlClientSocket.h"
#include "igtlTrackingDataMessage.h"
#include "igtl_tdata.h"
#include "igtlImageMessage.h"
#include "igtlStatusMessage.h"

#define UnityIGTLinkReceiveAPI _declspec(dllexport)

igtl::ImageMessage::Pointer SendingMsg;
igtl::ClientSocket::Pointer Unityclientsocket;

extern "C" {
    typedef void(_stdcall* CallbackImage)(void* ImageDataPointer);
    UnityIGTLinkReceiveAPI void _stdcall GetImage(CallbackImage GetImageData)
        //-----
    --
    {
        // Create a message buffer to receive header
        igtl::MessageHeader::Pointer headerMsg;
        headerMsg = igtl::MessageHeader::New();
        //-----
        // Allocate a time stamp
        igtl::TimeStamp::Pointer ts;
        ts = igtl::TimeStamp::New();
        //-----
        //for (int i = 0; i < 100; i++)
        //{
        // Initialize receive buffer
        headerMsg->InitPack();
        // Receive generic header from the socket
        int r = Unityclientsocket->Receive(headerMsg->GetPackPointer(),
headerMsg->GetPackSize());
        headerMsg->Unpack();
        //-----
        // Get time stamp
        igtlUInt32 sec;
        igtlUInt32 nanosec;
        headerMsg->GetTimeStamp(ts);
        ts->GetTimeStamp(&sec, &nanosec);

        // Check data type and receive data body
        if (strcmp(headerMsg->GetDeviceType(), "IMAGE") == 0)
        {

            igtl::ImageMessage::Pointer imgMsg;
```

```

        imgMsg = igt1::ImageMessage::New();
        imgMsg->SetMessageHeader(headerMsg);
        imgMsg->AllocatePack();
        // Receive transform data from the socket
        Unityclientsocket->Receive(imgMsg->GetPackBodyPointer(),
imgMsg->GetPackBodySize());
        // Deserialize the transform data
        // If you want to skip CRC check, call Unpack() without argument.
        int c = imgMsg->Unpack(0);
        if (c & igt1::MessageHeader::UNPACK_BODY) // if CRC check is OK
        {
            // Retrieve the image data
            int size[3]; // image dimension
            float spacing[3]; // spacing (mm/pixel)
            int svsize[3]; // sub-volume size
            int svoffset[3]; // sub-volume offset
            int scalarType; // scalar type
            int endian; // endian

            scalarType = imgMsg->GetScalarType();
            endian = imgMsg->GetEndian();
            imgMsg->GetDimensions(size);
            imgMsg->GetSpacing(spacing);
            imgMsg->GetSubVolume(svsize, svoffset);
            imgMsg->AllocateScalars();
            imgMsg->GetScalarPointer();
            GetImageData(imgMsg->GetScalarPointer());
        }
    }
    else
    {
        Unityclientsocket->Skip(headerMsg->GetBodySizeToRead(), 0);
    }
}
typedef void(_stdcall* CallbackReceiver)(int Cvalue);
UnityIGTLinkReceiveAPI void _stdcall ReceiverConnect(char* hostname, int
clientport, CallbackReceiver ReceiverConnectionStatus)
//UnityIGTLinkReceiveAPI void ReceiverConnect(char* hostname, int
clientport, void(*ReceiverConnectionStatus)(int Cvalue))
{
    igt1::ClientSocket::Pointer clientsocket;
    clientsocket = igt1::ClientSocket::New();
    int j = clientsocket->ConnectToServer(hostname, clientport);
    Unityclientsocket = clientsocket;

    if (j == 0)
    {
        ReceiverConnectionStatus(1);
    }

    //-----
    // Close connection (The example code never reaches this section ...)
    // clientsocket->CloseSocket();
}
UnityIGTLinkReceiveAPI void _stdcall CloseReceiveSocket()
{
    Unityclientsocket->CloseSocket();
}

```

}

}

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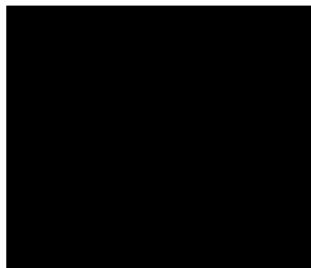
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Curriculum Vitae

Name: Yanyu Mu

**Post-Secondary
Education and
Degrees:** MEng, Biomedical Engineering
Western University, London, Ontario
2019 - Present

BEng, Electrical Engineering - Biomedical signals and systems
Western University, London, Ontario
2018

**Related Work
Experience:** Teaching Assistant
The University of Western Ontario
2018 - 2020

Research Assistant
The University of Western Ontario
2017 - 2018

Service Technician
Canadian Measurement Metrology
2014 - 2017

Service Technician
Industrial Drive Service Inc.
2013 - 2014

Journal Manuscript:

Mu, Y., Hocking, D., Wang, Z., Garvin, G.J., Eagleson, R. and Peters, T.M., Augmented reality simulator for ultrasound-guided percutaneous renal access. *International Journal of Computer Assisted Radiology and Surgery*, 15(5):749–757, 2020.

Abstracts:

Rivero, M., **Mu, Y.**, Roth, J., Wilson, T., Eagleson, R., and Sandig, M., Studying histology in 3d: Development and evaluation of an interactive virtual histology learning tool using a 3d model of the renal corpuscle. *The FASEB Journal*, 32(S1):25.5–25.5, 2018.

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