3D Printing in Congenital Heart Disease

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24.1 Introduction

Stereolithography, or as it is more commonly known, 3D printing, was first performed by Charles Hull in 1983 when he used the technique to produce a small plastic cup [1], and his first patent was issued in 1986 [2]. Since that time, there has been a rapid expansion of the use of 3D printing, initially in the fields of engineering and manufacturing, but more recently, in medicine. One field in which 3D printing has been embraced is congenital heart disease (CHD). Given the complex and variable nature of CHD, the ability to visualize and hold these hearts in your hand has been a game changer. In this chapter, we will explore some of the technical aspects of 3D printing, image acquisition in CHD, applications and limitations of 3D printing, and future uses of the technology.

24.2 The Process of 3D Printing

3D printing, or additive manufacturing, is the process of successive layering of a material to generate a structure with length, width, and height. The steps of digital model development and preparation are illustrated in Fig. 24.1. The first step is to acquire a digital 3D dataset, and for CHD, this is typically from computed tomography (CT) or magnetic resonance imaging (MRI). Once the dataset is acquired, the areas of interest for printing must then be identified and highlighted, a process called segmentation (Fig. 24.1), and usually converted to stereolithographic (STL) file formatting, the same that Charles Hull developed. The segmented data-

set is then sliced, or converted into a stack of cross sections, which will become the printed layers (Fig. 24.1). The sliced file also has instructions for the 3D printer to physically print the object. The process of segmentation and slicing can take minutes to hours, and the print time can be as long as 24 h, depending on the size and complexity of the structure to be printed.

The most common 3D printing technologies in use today are fused deposition modeling (FDM), selective laser sintering (SLS), stereolithography (SLA), PolyJet, and digital light processing (DLP). Each has its own advantages and disadvantages, and a full analysis is beyond the scope of this chapter, but some examples of the printing process and final products from FDM and DLP printers are shown in Fig. 24.2.

Some of the earliest descriptions of 3D printing in CHD came in 2006 when several groups reported printing models of various defects, including direct comparisons between the printed model, standard angiography, and direct visual inspection at the time of surgery [3, 4]. 3D datasets that have typically been used for CHD are CT or MRI, but they need to be high quality for accurate model generation. There is also a need for expertise in software and hardware manipulation [5, 6]. Several groups have quantitatively compared 3D-printed models to the original CT datasets and found high fidelity in the reproductions [7, 8].

While it is ideal to minimize exposure to ionizing radiation in patients with CHD, many have metallic implants (stents, occluder devices, and coils) which, even if safe for MR imaging, could cause enough image distortion to render MRI useless. In addition, image acquisition time is still longer for MRI such that infants need to be anesthetized to obtain diagnostic images. In response, cardiac CT scans with dose-reduction techniques have been used for infants with a variety of CHD and were found to generate excellent-quality images [9]. There have also been improvements in MR postprocessing techniques so 3D modeling from MRI may become more commonplace [10]. While CT typically provides higher resolution imaging for 3D model generation,



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Fig. 24.1 Process of 3D model generation for a neonate with pulmonary atresia, intact ventricular septum, and a tortuous patent ductus arteriosus. (a) Once a 3D dataset is obtained (typically from CT or MR imaging), the structures of interest are highlighted, a process known as segmentation. This shows axial, coronal, and sagittal projections with the segmented structures highlighted in orange and the 3D reconstruction in the upper right box. (b) Once the model has been cleaned and checked for errors, it is imported into slicing software which will generate support structures (gray) which are needed during the printing process and provide instructions for the movements for the specific 3D printed to be used



our group has been able to successfully use MRI to generate 3D models on newborn infants weighing less than 3 kg (Fig. 24.3).

Newer imaging modalities have allowed for acquisition of 3D datasets at the same time as other clinically indicated procedures. 3D rotational angiography (3DRA), which is increasingly performed as part of routine cardiac catheteriza-

tions, can produce printable 3D datasets [11, 12]. Due to the concern for increased radiation doses with 3DRA, Fetterly performed simulated radiation dose reductions for 3DRA imaging and found that even after reducing the radiation dose by 72%, there were no significant differences in the quality of the imaging or measurements of the structures of interest [13].



Fig. 24.2 The process of 3D model printing for a neonate with pulmonary atresia, intact ventricular septum, and a tortuous patent ductus arteriosus. (a-c) An example of fused deposition modelling (FDM) printing which heats and extrudes filament and draws the layers to create a 3D model. Once the model is complete, there are support structures which are needed during the print but are removed afterwards to

An exciting new development is the use of echocardiographic datasets to generate 3D-printed models without the use of any ionizing radiation. As detailed as CT and MRI are, thin structures, such as the atrioventricular valves, can be difficult to visualize, particularly in smaller patients. Several groups have utilized 3D datasets from either transthoracic or transesophageal echocardiograms to successfully print 3D models [14–17]. Additionally, others have been able to 3D print models of complex CHD with data obtained from fetal

produce the final model. $(\mathbf{d}-\mathbf{f})$ An example of digital light processing (DLP) printing which uses a light source to photocure resin material to generate the layers and create a 3D model. Once the model is complete, there are support structures which are needed during the print but are removed afterwards to produce the final model. Photos for $(\mathbf{d}-\mathbf{f})$ provided by Jason Pedersen

echocardiograms which have been particularly useful for prenatal counseling [18, 19].

Finally, combining all these modalities into hybrid model generation has allowed teams to use the best aspects of each technique to generate accurate 3D-printed models and optimize the clinical data obtained from the models. These models can be used for surgical planning, hemodynamic simulation, or education [14–16].



Fig. 24.3 Printed 3D model of a 2.7 kg neonatal with a hypoplastic transverse aortic arch (arrow) created from a cardiac MRI obtained on a 1.5 T scanner

24.3 Current Applications of 3D Printing for Congenital Heart Disease

The use of 3D-printed models in CHD is relatively new [3, 4], but there has been rapid adoption within the field for multiple uses, including procedural planning (surgical and transcatheter), hemodynamic simulations, and education [20]. To support these uses, several studies have demonstrated the accuracy of 3D-printed models within 1 mm of standard clinical imaging [21, 22].

24.3.1 Surgical Planning for Complex CHD Repair

Two of the most challenging forms of CHD to repair surgically are double outlet right ventricle (DORV) and complete atrioventricular septal defects (AVSD) as the complex intracardiac relationships can be quite challenging to visualize by standard transthoracic echo alone and the interventional decisions made can greatly alter the long term outcomes for these patients.

For DORV, the location of the aorta relative to the ventricular septal defect (VSD) determines whether a surgeon can create an intracardiac baffle from the left ventricle, through the VSD and out the aorta to establish biventricular circulation (Fig. 24.4). 3D-printed models have been used to guide the critical decision of either initiating single ventricle palliation and potentially later converting to biventricular circulation [23] or performing biventricular repair as the initial intervention in patients where standard imaging supported single ventricle palliation [24-29]. To further understand the potential clinical benefits of a pre-operative 3D-printed model, one team compared outcomes for patients with DORV between those with and those without 3D-printed models. Encouragingly, post-operative ventilator time and ICU stays were significantly shorter in the 3D model group [30].

For patients with an AVSD, particularly in the setting of heterotaxy syndrome, it can be difficult to determine patient suitability for single versus biventricular intracardiac repair. A review of the management of AVSD highlighted the beneficial addition of 3D echocardiography and 3D-printed models in helping to determine adequacy of biventricular repairs in borderline cases [31]. A series of patients with complex AVSD achieved conversion to biventricular circulation with the aid of 3D-printed models as part of the standard pre-operative evaluation [32].

In addition to DORV and AVSD, researchers have utilized 3D-printed models for a variety of complex CHD cases, including variants of tetralogy of Fallot [33, 34], transposition of the great arteries [35], anomalous pulmonary venous connection [36], pulmonary artery pseudoaneurysm resection [37] and rerouting of a Fontan pathway [38]. It is felt that 3D-printed models used for surgical planning can reduce the exploration and surgical times, which in turn can improve clinical outcomes. The 3D model can also help in the understanding of complex patient anatomy and can help improve pre-operative communication between the surgeon and the patient and family [37] (Fig. 24.5).

Overall, 3D printing can be very useful in planning surgical techniques and in choosing between single ventricle, one and a half ventricle, and biventricular repair [39]. It can also improve multidisciplinary team decision-making, patient and family understanding, and education of medical professionals [39, 40].

24.3.2 Mechanical Cardiac Support and Heart Transplantation

It is inevitable that some patients with CHD will develop heart failure and require mechanical cardiac support and possible heart transplantation. Many of these patients will have unusual intracardiac and intrathoracic post-operative anat-

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Fig. 24.4 Utility of 3D models for surgical repair of double outlet right ventricle (DORV). (a, b) Two views of a virtual 3D reconstruction of an infant with DORV (right ventricular free wall removed) with the aorta highlighted in red, the pulmonary artery highlighted in blue, and the ventricular septal defect (VSD) outlined with the dashed line. (c, d)

omy, and 3D-printed models can be used to plan for ventricular assist device (VAD) implantation [41]. This can be particularly important for adults with CHD as the multiple prior surgeries can make re-intervention more challenging and higher risk. And while these patients could benefit from mechanical cardiac support, their anatomy and risk factors may preclude it. 3D-printed models can help identify the best approach for placing mechanical support devices, increasing surgeon confidence to undertake the procedure, and may lead to shorter procedure times as well [42].

For those who go on to heart transplantation, 3D-printed models can help surgeons anticipate and plan for problems that may arise during dissection or implantation of the donor

Two views of a 3D-printed model of the same patient with the red line representing the direction of flow from the left ventricle, through the VSD and out the aorta and the blue line representing flow from the right ventricle through the right ventricular outflow tract and out the pulmonary artery

heart. Models can be particularly useful in determining any special dimensions and requirements of the donor heart, such as extra donor venous tissue to "undo" Fontan procedures [43–45]. Better pre-operative planning can also reduce cross-clamp and total operative time, further reducing patient morbidity [44].

24.3.3 Cardiac Catheterization

The field of interventional congenital cardiology has always pushed the envelope to develop minimally-invasive approaches to minimize morbidities for this fragile popula420



Fig. 24.5 3D-printed model of a neonate with double aortic arch from anterior (**a**) and posterior (**b**) views. The double aortic arch (red) with a hypoplastic left segment (*) enwraps the trachea and esophagus (white).

These models can help families to understand the anatomy of their child's congenital heart defect and need for surgical intervention

tion. The complex anatomy can be challenging to understand with angiography alone, so 3D-printed models have been embraced by the interventional community to optimize the planning and safety of interventional catheterizations. There are almost as many uses of 3D-printed models to guide transcatheter interventions as there are CHDs. 3D-printed models have been used for coronary artery fistula closure [46, 47], ductus arteriosus stenting to maintain pulmonary blood flow [48], closure of an atrial baffle leak in a patient with a crisscross heart [49], device closure of a right ventricular outflow tract pseudoaneurysm [50], device closure of a ruptured sinus of Valsalva aneurysm [51], transcatheter Fontan completion [52], closure of an unroofed coronary sinus [53], recanalization of a chronically occluded branch pulmonary artery [54] and percutaneous edge-to-edge repair of systemic atrioventricular valve regurgitation [55]. For each of these reports, the ability to simulate and practice intervention, as well as improved communication with the care team and patients, were noted benefits of the 3D-printed models.

The first reported use of a 3D-printed model to guide a catheterization was in 2014 when Olivieri and colleagues printed a 3D model of the heart of a patient with pulmonary venous baffle obstruction after a Mustard procedure to aid in planning successful transcatheter stent placement [56]. This was followed by reports of 3D-printed models to guide catheter-based interventions on complex aortic coarctation and arch hypoplasia [57, 58].

Device closure of atrial septal defects (ASDs) is one of the earliest transcatheter interventions, but the septal geometry can be complex and hard to assess by standard imaging techniques, such as echocardiography, such that some patients who could benefit from device closure instead undergo surgical closure. In particular, some patients appear to have insufficient tissue rims around the ASD on echocardiogram to allow for safe placement of standard occluder devices. Several groups have found that 3D-printed models can allow for more accurate visualization and assessment of the tissue rims [59] and for in vitro test closure of complex ASDs with standard and off-label closure devices [60, 61].

An exciting extension of 3D printing in planning transcatheter interventions is the treatment of superior sinus venosus ASDs and partial anomalous pulmonary venous return, which have historically required surgical closure. However, using 3D-printed models and bench simulations of covered stent placement has allowed planning to create the "missing" wall between the superior vena cava and the pulmonary veins (Fig. 24.6). The team at Evelina London Children's Hospital in the United Kingdom has pioneered this technique with good medium-term outcomes [62–64]. 3D model planning allows for selection of candidates with anatomy favorable for the transcatheter approach, and the procedure has successfully been replicated at other centers around the world [65].

3D-printed models have been used in electrophysiology catheter interventions as well. A number of adults with CHD will develop conduction abnormalities related to the underlying CHD or as a consequence of their prior interventions. Atypical anatomy is the norm for these patients and unusual approaches may be needed to successfully perform ablation procedures or place transvenous pacemaker leads (Fig. 24.7). Many patients with ventricular looping abnormalities can have unusual coronary sinus anatomy, [66] and it may even be absent [67], making pacemaker lead placement challeng-



Fig. 24.6 Planning for transcatheter correction of anomalous rightsided pulmonary venous return and a superior sinus venosus atrial septal defect. (**a**) 3D virtual model (anterior right atrial wall cut away) demonstrating the anomalous right-sided pulmonary veins entering the superior vena cava (*), the superior sinus venosus atrial septal defect (dotted line), and the normal entry of the azygous vein (†). (**b**) Virtual

implantation of a covered stent (white) to recreate the back wall of the superior vena cava, rerouting the anomalous pulmonary venous drainage and closing the atrial septal defect. (c) 3D-printed model for preprocedure simulation. Note that the azygous vein is not obstructed by the covered stent



Fig. 24.7 3D-printed model of a patient with transposition of the great arteries who has undergone a Mustard procedure and required transcatheter ablation for atrial flutter with a tricuspid annular focus from lateral (**a**) and anterior (**b**) views. Because of the complex surgical intraatrial venous rerouting, the inferior vena cava no longer communicated

with the tricuspid valve annulus (dotted line) and transbaffle puncture was needed to perform the intervention. The 3D-printed model was used to plan the two puncture sites (*) needed to position the interventional catheters (represented by the blue line) ing, but 3D-printed models have been used in such cases to plan and guide an alternative approach for successful lead placement.

24.3.4 Simulation

Aside from procedural planning, groups have also been exploring the utility of 3D-printed models for various simulations. The most common use has been simulated surgical intervention [20]. Trainees and junior surgeons can learn complex congenital surgical interventions on life-size 3D-printed models made from thin, flexible materials, such as polyurethane resin, before sewing a stitch in a human [68, 69]. Several authors have published small series and proofof-concept papers, but these works have lacked objective data to prove the utility of the models in improving surgical outcomes [70], aside from one objective assessment of successful simulated arterial switch procedures, to be able to better understand the utility of surgical simulation in improving outcomes [71]. Groups have also utilized 3D-printed models for simulated transcatheter interventions, such as intracardiac shunt occlusion [69] and interventions for pulmonary artery stenosis [72], but more assessment tools will need to be developed for other lesions to understand the potential benefit of 3D-printed models for clinical care.

Another innovative use of 3D-printed models is to simulate post-intervention physiology. One team 3D printed a series of different-sized simulated ASDs to allow for deployment of a transcatheter ASD occluder to better define and understand the deployed dimensions of different device sizes in different-sized defects [73]. Another group used 3D-printed models to predict adverse outcomes after surgical intervention on coronary arteries both for arterial switch procedures [74] and for repair of anomalous coronary artery origins [75]. 3D-printed models of right ventricular outflow anatomy (normal and tetralogy of Fallot after transannular patch repair) were used in hemodynamic simulators and 4D MRI modeling of flow to assess factors that can affect the longevity of implanted bioprosthetic valves [76]. Another group used 3D-printed mitral valves in hemodynamic simulators to predict hemodynamic changes after transcatheter edge-toedge valve repair [77], a technique that could potentially be used in single-ventricle patients. Another team compared 3D-printed models of Fontan conduits using SLA, SLS, and FDM printers and assessed flow dynamics with hemodynamic simulators. Overall, each made a reasonable representation, but the internal support structures required in FDM models produced less accurate flow modeling [78]. The attractive low-cost FDM models will need to be balanced against the anatomic and physiologic accuracy of the model.

24.3.5 Education

Just as surgeons can learn to perform an arterial switch operation on a 3D model, students can learn anatomy of CHD lesions using models. An open-access library of 3D models of CHD that would be freely available for downloading and printing has been proposed [79]. Others have used cardiac CT of fetal hearts [80] and digital manipulation of cardiac CT scans of normal hearts [81] to create libraries of CHD for education.

There have been numerous studies that assess the subjective and objective benefits of incorporating 3D-printed models of CHD into didactic sessions. These have included multiple levels of learners (medical students, nursing staff, cardiology fellows, attending cardiologists and surgeons) and covered a variety of CHD, such as VSDs and tetralogy of Fallot [82–90], extracardiac vascular defects [84, 85, 88–91], crisscross hearts [92], and complex single ventricle lesions [85, 93]. These studies have shown subjective improvements in learner understanding, but the objective improvements have been less consistent. One innovation that improves understanding is the use of different colored materials to distinguish anatomic structures [85]. There is also probably more benefit for higher complexity lesions (tetralogy of Fallot versus an isolated VSD) [87] and for more junior learners [89, 92].

3D-printed models of CHD have also been used for hands-on surgical training which can allow for practice on pathological hearts without patient risk or having to follow the traditional training model of waiting for potential opportunities to present themselves [68, 94-98]. Hearts can be 3D printed with flexible, tissue-like material to allow for cutting and suturing and have been used for transposition of the great arteries, tetralogy of Fallot, and hypoplastic left heart syndrome, among other defects. Earlier studies found the material to be too different from human myocardium to be useful [94], but there have been advances in material development [68]. Cardiac valves were poorly recreated in the earlier models [94], but a newer technique for valve simulation using 3D echocardiographic imaging to print negative molds and then cast silicone valves has created more tissuelike valves for surgical practice [96]. Hussein and colleagues have shown a reduction in simulated operative time for the arterial switch procedure with repeated practice using 3D-printed models [97]. The same group has also created an objective assessment tool to define successful simulated arterial switch procedures to better understand the utility of surgical simulation in improving clinical outcomes beyond just shorter operative time [71]. Scientific proof of the utility of 3D-printed models for surgical education is challenging due to many confounders that affect outcome [99], and more tools will need to be developed for other CHD to truly understand the potential benefit of 3D-printed models for clinical care.

Besides medical professionals, 3D models can educate patients and their families about their cardiac defect. Children with CHD will require lifelong cardiology care, and a good understanding of the pathologic disease process is likely helpful in improving medical compliance. This can start as early as the initial diagnosis with fetal echocardiograms being used to 3D print models [18, 19]. Pre-operative 3D-printed models can also help families understand the indications for surgery and aid in operative consent [39, 100]. 3D-printed models have also been shown to help reduce patient and family anxiety before cardiac catheterizations [101]. Stressing the importance of lifelong cardiology follow-up, Liddle used virtual 3D models to teach adolescents with CHD about their disease and found improved understanding after the teaching sessions [102]. A metaanalysis found a small number of reports which suggest that 3D models are accurate and help with communication, but consistent with other studies, objective clinical utility has yet to be demonstrated [103].

24.4 Future of 3D Printing for Congenital Heart Disease

24.4.1 Machine Learning

There are several barriers to widespread adoption of 3D printing in CHD, and one of the most important is fast, accurate segmentation from the initial 3D dataset. This is currently mostly a manual, time-consuming process, and any inaccuracies that occur during segmentation will carry through the entire 3D printing process, which can lead to inappropriate clinical interpretations or plans [104]. There are only limited options for semi-automated segmentation of scans [105] and still no consistent method for model creation and printing across centers [106]. Machine learning has been proposed as a solution for improved automated segmentation [107]. The striking variation in CHD anatomy that occurs in a relatively small population of patients limits the datasets available for machine learning [107]. However, progress has been made with development of a machine learning technique to autosegment the left atrium and anomalous pulmonary veins for pre-operative planning in patients with total anomalous pulmonary venous return [108]. The authors report that using their method would decrease the radiologist's workload from several hours per scan to only 400 ms. Expansion of this technique to other CHD will be critical to widespread and clinically useful adoption of 3D modeling and printing.

24.4.2 Bioprinting

An exciting area of future development for 3D printing in CHD is bioprinting of customized implants [109]. Work has been done to create cardiac valve replacements [110], bioprinted scaffolds seeded with a patient's stem cells to create vascular grafts that can be used to replace stenotic or hypoplastic vessels [111], or implanted as Fontan conduits [112] that can grow with the patients. While tissue engineering has existed for some time, 3D printing can aid in more lifelike and anatomically accurate scaffold creation [113]. Another step toward growing implants is 4D bioprinting of materials that are designed to respond to specific stimuli, such as heat, to alter their shape which could allow for growth of implanted structures, such as intravascular stents [114]. In addition, implants printed from biodegradable material could grow with the patients, who would not need chronic anticoagulation typically used after implantation of metallic devices [115].

Currently, there is no ideal printer available, but industry has taken notice and is working to develop printers that can provide optimal resolution and speed with low costs and high viability of the printed cells [113].

24.5 Limitations of 3D Printing for Congenital Heart Disease

As exciting as 3D-printed models are for CHD, there are still several limitations that hinder widespread adoption. The first was discussed earlier and relates to the time-consuming nature of the process of model generation and the steep learning curve [104, 106, 116]. There have already been steps made at reducing segmentation time by utilizing machine learning [108] and these will only continue to improve as larger datasets of CHD anatomy are created. There is still a lack of consistency in methods for segmentation and model generation [105, 106], but future work will continue to refine the techniques to aid in universal adoption.

The costs of 3D printing cannot be ignored, and are often cited as a reason to reject adoption [37]. However, several studies have looked at this issue and found that highly accurate and clinically useful models can be printed using free and open source software and commercially available FDM desktop printers, often for average costs of less than €100 per model [117, 118].

Another limitation is the difficulty in quantifying the clinical benefits of 3D-printed models of CHD. There are several reports of subjective improvements in procedural planning, but objective measures are still lacking [6, 119]. Work that has already been undertaken to develop objective outcome measures for CHD surgery [97] will continue to provide data to "prove" the utility of 3D-printed models in improving care for patients with CHD.

24.6 Conclusions

Incorporation of 3D printing into clinical care for patients with CHD has rapidly evolved in the past two decades and has helped increase understanding of the anatomy and physiology and allowed for the development of novel treatment approaches. As technology continues to improve and adoption becomes more widespread, there will be further optimization of the outcomes for this complex patient population.

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