

Low cost fabrication of PVA based personalised vascular phantoms for in vitro haemodynamic studies: three applications

Giacomo Annio[†]

Dept. Medical Physics and Bioengineering
University College London
Torrington Place WC1E 7JE London, UK
Email: giacomo.annio.15@ucl.ac.uk

Gaia Franzetti[†]

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Email: gaia.franzetti.15@ucl.ac.uk

Mirko Bonfanti

Wellcome/EPSCRC Centre for
Interventional and Surgical Sciences
UCL Mechanical Engineering
University College London
Torrington Place WC1E 7JE London, UK
Email: m.bonfanti@ucl.ac.uk

Antonio Gallarelo

Dept. of Electronics, Information and Bioengineering
Politecnico di Milano
20133, Milano, Italy
Email: antonio.gallarelo@mail.polimi.it

Andrea Palombi

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Email: andrea.palombi.16@ucl.ac.uk

Elena De Momi

Dept. of Electronics, Information and Bioengineering
Politecnico di Milano
20133, Milano, Italy
Email: elena.demomi@polimi.it

Shervanthi Homer-Vanniasinkam

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Leeds Teaching Hospitals NHS Trust
Leeds, LS1 3EX, UK
Email: s.homer-v@ucl.ac.uk

Helge A. Wurdemann

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Email: h.wurdemann@ucl.ac.uk

Victor Tsang

Cardiothoracic Surgery Unit
Great Ormond Street Hospital for Children
Holborn
WC1N 3JH London, UK
Email: victor.tsang@gosh.nhs.uk

Vanessa Díaz-Zuccarini

Wellcome/EPSCRC Centre for
Interventional and Surgical Sciences
UCL Mechanical Engineering
University College London
Torrington Place WC1E 7JE London, UK
Email: v.diaz@ucl.ac.uk

Ryo Torii

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Email: r.torii@ucl.ac.uk

Stavroula Balabani^{*}

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Email: s.balabani@ucl.ac.uk

Gaetano Burriesci^{*}

UCL Mechanical Engineering
University College London
Torrington Place
WC1E 7JE London, UK
Ri.MED Foundation
Via Bandiera, 11
90133, Palermo, Italy
Email: g.burriesci@ucl.ac.uk

Vascular phantoms mimicking human vessels are commonly used to perform *in vitro* haemodynamic studies for a number of bioengineering applications, such as medical device testing, clinical simulators and medical imaging research. Simplified geometries are useful to perform parametric studies, but accurate representations of the complexity of the *in vivo* system are essential in several applications as personalised features have been found to play a crucial role in the management and treatment of many vascular pathologies. Despite numerous studies employing vascular phantoms produced through different manufacturing techniques, an economically viable technique, able to generate large complex patient-specific vascular anatomies, accessible to non-specialist laboratories, still needs to be identified. In this work, a manufacturing framework to create personalised and complex phantoms with easily accessible and affordable materials and equipment is presented. In particular, 3D printing with polyvinyl alcohol (PVA) is employed to create the mould, and lost core casting is performed to create the physical model. The applicability and flexibility of the proposed fabrication protocol is demonstrated through three phantom case studies - an idealised aortic arch, a patient-specific aortic arch, and a patient-specific aortic dissection model. The phantoms were successfully manufactured in a rigid silicone, a compliant silicone and a rigid epoxy resin, respectively; using two different 3D printers and two casting techniques, without the need of specialist equipment.

1 Introduction

In the past 30 years, fluid dynamic investigations have played an important role in the study of the cardiovascular system, providing insight into disease condition and progression and support for management optimisation. *In vitro* haemodynamic studies typically rely on a physical representation - *synthetic phantom* - of the vessels of interest to replicate physiology or disease in the lab. Vascular phantoms have thus been successfully employed in numerous applications, ranging from the study of healthy and diseased haemodynamics in regions of the human cardiovascular system [1], to the assessment of the accuracy of imaging modalities and computational techniques [2], evaluation of medical devices [3] and the planning of surgical interventions [4]. Their success is due to the advances in medical imaging modalities and the commercialisation of 3D printing devices. The former provided access to higher resolution images and noise reduction in the acquired data, thus enabling the retrieval of 3D accurate geometries. The latter allowed the manufacturing of cost-efficient 3D models of regions of the circulatory system based on patient-specific data.

A broad range of vascular phantoms has been reported in the literature, varying in complexity depending on the study aims and assumptions made. Reported phantoms differ primarily in geometry, material properties and fabrication methods. In terms of geometry, vascular phantoms tend to vary from idealised to patient-specific ones. Even though ide-

alised geometries can play a key role in parametric studies [5] and complex flow investigations [6], accurate anatomical descriptions are essential to reproduce patient-specific conditions and capture the *in vivo* haemodynamic features for a variety of applications; personalised features - morphology, flow patterns, pressures and velocities - have been shown to play an important role in several cardiovascular pathologies, and therefore diagnosis management and treatment are strictly patient-specific [7–9]. Experimental mock circulatory loop that can reproduce individualised conditions are being developed in this context [10], alongside personalised computational models [11], that also necessitate fabrication of accurate vascular phantoms.

Vascular phantoms can be manufactured to be either rigid or compliant, in order to approximate the elastic properties of the vasculature. The choice of material often depends on the imaging modality employed for the *in vitro* study. Magnetic Resonance Imaging (MRI) and Particle Image Velocimetry (PIV) are imaging techniques extensively used to perform *in vitro* haemodynamic studies in order to obtain flow field information in the vascular region of interest. For instance, MRI compatible phantoms, which only need to be non-magnetic, and have been shown to allow imaging protocol optimisation [12] or evaluation of different surgical outcomes [13]. On the other hand, PIV derived velocity information has been used as validation tool for computational haemodynamics studies [14], to investigate complex flow regimes [6] or to evaluate the consequences of medical device implantation [15], due to its high spatial and temporal resolution compared to MRI. To be PIV-compatible, phantoms need to be optically clear and manufactured in a material whose refractive index is matched with the working fluid. Rigid transparent models, manufactured mainly in resin [16] or glass [2], have commonly been employed in PIV studies due to optical access requirements, whereas compliant models have often been manufactured using a broad range of opaque materials such as rubber-like compounds [17], silicone [14], latex [18] and polyurethane [19], which are compatible with MRI.

Various phantom manufacturing methods have been reported in the literature. The most common ones are dipping [7], dripping [20], and the use of two different moulds, one for the inner and one for the outer part [21]. These approaches are invariably fraught with difficulties, especially when the geometry is complex. For example, dipping and dripping may require a large quantity of casting material (80–90%) to be discarded, leading to unnecessary costs [22], and do not allow control of the phantom wall thickness, which can be essential in *in vitro* studies with compliant geometries. To overcome these challenges, lost core casting based techniques have been employed and soluble moulds made in materials such as wax [23], chocolate [24] and isomaltose [25], that can be dissolved after the curing of the phantom material have been used. These moulds can achieve fine detail, but the shrinkage occurring during the cooling process can lead to inaccuracies in the final model. Moreover, the use of such materials for lost core casting requires one additional step in the manufacturing process (consisting in the creation of the

*Corresponding authors. † The authors equally contributed to this work.

negative mould for casting the lost core geometry), which leads to additional time and financial costs, as well as additional errors. Subtractive manufacturing of acrylic-based phantoms has been successfully implemented to manufacture PIV and MRI compatible phantoms [26]. However, this technique has two main drawbacks: first, it requires a CNC machine, which is not always readily available; and second, it may lead to issues regarding the fabrication of complex patient-specific geometries, especially for PIV compatible phantoms, arising from joining parts of the geometry that can cause laser optical distortion. Moreover, this methodology is unsuitable for the manufacturing of compliant phantoms, neglecting a factor which often contributes significantly to the flow dynamics.

A comprehensive review of different manufacturing materials and procedures, and their limitations can be found in the work of Yazdi et al. [22]. It is apparent that despite the number of studies reported in the literature, manufacturing patient-specific phantoms in the laboratory at accessible cost and without the need of specialised equipment (e.g. big volume vacuum chambers, water jets) and expertise remains an issue, especially in the case of complex geometries and large volumes.

In this paper, we present a simplified lost core methodology, modified to manufacture idealised and patient-specific vascular phantoms for haemodynamic studies and imaging applications in a low cost manner and without access to specialist equipment, addressing some of the limitations of previous works and allowing non-specialist laboratories to manufacture experimental phantoms. In particular, negative moulds are created through 3D printing in polyvinyl alcohol (PVA), a water soluble material, to develop both compliant and rigid phantoms through a lost core casting process. The methodology and associated costs and equipment requirements are demonstrated through three aortic phantoms with different geometrical and mechanical properties; these include a simplified aortic arch, a patient-specific aortic arch and a patient-specific aortic dissection model.

2 Material and Methods

In this section, the process of creating anatomical vascular phantoms using 3D printing in PVA is described.

2.1 Manufacturing process

The manufacturing process adopted in this study involves five stages, as illustrated in Fig. 1 and explained in detail in the following sections: (1) Creation of the geometry either via segmentation of patient-specific clinical images or creation of a CAD idealised geometry; (2) 3D post processing; (3) 3D printing in PVA; (4) Physical model refinement; (5) Casting and dissolving of PVA in water.

Clinical data and segmentation process

Volumetric medical images (e.g. computed tomography (CT), magnetic resonance imaging (MRI)) are used to reconstruct patient-specific vessels via the segmentation process.

Image segmentation is the process of partitioning a digital image into different regions containing pixels with similar attributes. It is used to identify and label regions of interest and can be used to create volumetric reconstructions of different organs and tissues. For vascular phantoms, thresholding is commonly employed to reconstruct the lumen volume (i.e. blood volume). It uses the background to foreground contrast ratio to isolate the region of interest. Manual refinement is lastly used to adjust the vessel area, especially in the presence of complex anatomies. Details regarding the segmentation procedure and the mesh refinement can be found in Bücking et al. [27].

This first step is not necessary in the case of idealised geometries and the process starts with the generation of the CAD geometry (Step 1b in Fig. 1).

3D geometry post processing

The image segmentation process described above yields a 3D model of the target area. A local smoothing approach of the surface is subsequently performed to compensate for errors resulting from the resolution of the original medical images and to facilitate the manufacturing of the building components [27]. Moreover, geometrical modifications can be made in this stage according to the purpose of the vascular phantom. For instance, for use with *in vitro* studies, inlet and outlet sections need to be designed to accommodate the phantom within the experimental system.

3D printing

The CAD model reconstructing the vessel lumen volume is 3D printed in PVA using a stereolithography (SLA) printer. PVA is commonly employed as a support material to create complex geometries in Polylactic acid (PLA), which is then dissolved in water to obtain the object of interest. However, in this work, it is used as the primary material to create a mould for the casting process (i.e. negative mould). Attention was paid to the choice of the infill, one of the most important factors for 3D printing (i.e. the structure printed inside the object). A higher infill will result in a stronger model, which is especially important in the case of a complex structure, but it will also lead to increased duration of the PVA dissolving process. To find the best compromise, different simplified test sections were 3D printed varying the infill parameter (10% - 60%) and the shell thickness (1 mm - 3 mm).

Physical model refinement

The 3D printed model, constituting the negative mould, undergoes further post processing before the casting procedure. The surface of the 3D printed material presents a layering effect, characteristic of the FDM printers, that can cause unacceptable surface roughness in the final phantom. In order to make the surface smoother, sanding and coating with liquid PVA glue are used to provide a good finish to the phantom. A transparent acrylic spray paint (PlastiKote, UK or Humbrol, UK) is lastly used to make the contact surface im-

permeable and hydrophobic, so as to reduce chemical interactions with the casting material, which may affect its optical properties.

Casting and dissolving procedure

The final stage in the manufacturing process is the casting. In this phase, the physical phantom is created around the PVA model of the vessel lumen, which is then dissolved, resulting in the hollow structure. Two casting options are available, as illustrated in Fig. 2: (i) to enclose the model in a rigid box and pour the casting material to fill the space, which will result in a box with a hollow structure reproducing the vessel lumen; or (ii) to design an external mould according to the vessel geometry to control the vessel wall thickness. In this scenario, the casting material is poured in-between the two, resulting in a physical phantom having the shape of the vessel of interest.

The casting material can be selected based on the design criteria. For instance, paying particular attention to the mechanical properties (e.g. Young's modulus) in case of flexible phantoms or to the optical characteristics (e.g. transparency and Refractive Index, RI) for compatibility with the imaging modalities employed (e.g. PIV). The curing time is adjusted accordingly, and subsequently the object is placed in a water bath to let the inner PVA mould dissolve.

2.2 Phantom 1: Simplified aortic arch model

Phantom 1 was manufactured to reproduce a simplified aortic arch without branch vessels, realised as a U bend (Fig. 3). It was designed to be rigid and compatible with both MRI and PIV imaging (which necessitates the use of an optically transparent material). The mould was designed with a CAD software (SolidWorks, Dassault Systèmes, Canada) and then 3D printed in PVA using a Delta WASP 2040 Turbo 2 (Wasp, Italy). Post processing with sanding and liquid PVA coating was performed to minimise the roughness of the model surface. For the casting process, an acrylic box with flat sides was manufactured to allow PIV laser illumination and pouring of the selected casting material. A clear, solvent free, low viscosity silicone elastomer (MED 6015, NuSil Technology, CA, USA) was chosen because it has a low refractive index ($RI = 1.4$) that could be matched by water-glycerol solutions. MED 6015 is supplied as two-part mix which have to be mixed thoroughly to ensure homogeneous optical properties. To remove bubbles entrained through this process, the casting material was degassed using a small vacuum chamber. The material is then poured in several batches to prevent air bubbles to be trapped in thick layers. After curing, the entire model was immersed in warm water and the inner mould and coating in PVA dissolved. No additional equipment was used at this stage (e.g. water jets, scrapers). The compatibility of Phantom 1 with PIV measurements was subsequently tested. Experiments were run connecting the phantom with a Harvard-Apparatus blood pump (Harvard Apparatus, USA) and using a fluid with the same refractive index (i.e. water-glycerine solution). The sagittal plane of the model was imaged using a TSI 2D-PIV (TSI, USA) system and the corre-

spondent 2D velocity field was reconstructed.

2.3 Phantom 2: Patient-specific aortic model

Phantom 2 was manufactured from patient-specific data and designed to be flexible, in order to mimic the compliant behaviour of the vessel. The original dataset was obtained from a CT-angiography of a 76-year-old patient undergoing a TAVR (Transcatheter aortic valve replacement) procedure. The study was reviewed and approved by the internal review board of the research group, which was the relevant cross-institutional committee responsible for assessing the methodological appropriateness and ethics of the study design (AO San Camillo-Forlanini, Rome, Italy). The subject gave written informed consent in accordance with the Declaration of Helsinki and the data set was anonymised. Manual double-threshold segmentation was performed using 3DSlicer (Slicer, NIH) and part of the vessel – ascending aorta, aortic arch and the initial part of the abdominal aorta – was reconstructed (Fig. 4c).

The structure was post-processed using Geomagic Control (3D Systems, Canada) and MeshMixer (Autodesk, USA). Smoothing was performed to reduce the noise introduced by the thresholding method, in order to prepare the surface to the 3D printing process. The aortic lumen volume was 3D printed in PVA with a Delta WASP 2040 Turbo 2 (Wasp, Italy) (Fig. 4d). Since Phantom 2 was designed to reproduce the vessel wall thickness and mimic its compliance, a second, external mould was printed. An offset value of 2 mm was applied to the 3D aortic geometry and three pins were added to the internal geometry to allow a stable alignment between the internal and external moulds, hence maintaining the correct gap. The outer part of the mould was separated in multiple sections in order to simplify the extraction process of the phantom. These parts were kept together by mean of bolts and nuts to guarantee a perfect alignment. The external mould was printed using white resin with a Form 2 3D printer (FormLabs, USA).

A two-parts silicone (Smooth-On Ecoflex™ 00-30 silicone) was selected as phantom material and casting was performed by pouring the material in between the two moulds, in order to reproduce the desired vessel wall thickness. Prior to pouring, in order to remove air bubbles resulting from the mixing of the two parts, the casting material was degassed using a vacuum chamber. Once cured, the external part was removed and the inner PVA model dissolved in water. For ease of connection to the experimental mock loop and to avoid any unwanted movement of the central part of the phantom, the latter was placed in a custom made box and the inlet and outlet connected to rigid connectors. In order to assess the geometrical accuracy of the phantom, a MR (Philips Ingenia 3T) scan was carried out adopting a three-dimensional fluid attenuated inversion recovery in order to reduce the signal arising from the silicone.

2.4 Phantom 3: Patient-specific diseased aortic model

Phantom 3 was created to illustrate the manufacturing process of a patient-specific case of a complex pathological

vascular geometry, using Type-B Aortic Dissection as a case study. In this pathology, a tear in the aortic wall causes the creation of two separate lumina, the false (FL) and the true (TL) lumina, posing a major manufacturing challenge. Due to the relatively large dimensions of this geometry, it was essential in this case to minimise the costs and processing, by avoiding casting materials requiring degassing procedures (i.e. no equipment needed).

A patient was imaged with a CT scanner (Siemens AG, Munich, Germany; Field of View: 284 mm, slice thickness: 1 mm), thus obtaining 946 slices with an in plane resolution of 0.55 mm and inter-slice distance of 0.7 mm. The geometry of the aortic model was created (ethical approval study no 788/RADRES/16) with the software ScanIP (Synopsys, Mountain View, CA, USA) using a semi-automated segmentation tool based on thresholding operations, differentiating the vessel based on the grey-scale of the images. Smoothing was performed on the resulting masks to reduce the artefacts due to pixelation (Fig. 5). The final geometry file was exported into a CAD software in order to modify the model for compatibility purposes with the experimental rig. In particular, rigid regular connectors were added to the geometry to facilitate the connection with the experimental rig. The stl file was created and the lumen region of both the TL and FL was 3D printed in PVA using a SigmaX printer (BCN3D, Spain). To avoid problems during the printing phase and reduce the support material needed, the TL and FL were printed separately and connected later in the post processing phase (Fig. 6).

The physical model was post-processed in three steps. First, several coats of PVA liquid glue were added on the external surfaces to reduce the roughness due to the different printing layers. Second, the TL and FL were joined together employing the same dissoluble glue guided by the connecting tear. Third, a transparent spray acrylic paint was used to make the surface impermeable and hydrophobic. This was also performed to avoid possible absorption of the PVA from the casting material.

Similarly to Phantom 1, a custom made box was designed and manufactured in polypropylene to contain the model and allow the casting process. Particular attention was made to align the inlet and outlets of the geometry to avoid any misalignment between the TL and FL. A rigid clear epoxy resin (GlassCast 50, EasyComposites, UK) was used as casting material, selected to minimise cost and bypass the need of degassing. Different layers of up to 50 mm were poured sequentially to avoid possible exothermic reactions during the curing process. At the end of the curing phase, the external polypropylene box was mechanically removed and PVA dissolved in warm water.

The final geometry of the phantom was imaged with a CT scan (O-arm system Medtronic, USA; pixel size: 194 μm , inter-slice distance: 0.8 mm) in order to assess its anatomical accuracy. The inner lumen volume was reconstructed with the same semi-automated segmentation tools previously described and quantitative data was extracted. Specifically, the cross-sectional area of the phantom lumina (i.e. TL and FL) at ten axial planes along the longitudinal direction were mea-

sured and compared against the original clinical images.

3 Results and Discussion

Table 1 summarises the technical properties of the three different phantoms manufactured in this work, including material costs and necessary equipment.

Two different 3D printers, with negligible difference in purchase price, were used and both allowed the creation of a detailed and accurate geometry of the vessel lumen. Figure 7 shows the details of the complex curvature of the lumen for the pathological aorta reconstructed in Phantom 3 illustrating the ability of the printer to follow the intricacy of the structure. The printers were chosen because of (i) the large build volume, which allows to reproduce bigger geometries, and (ii) because they have two nozzles, so that one can be used only for PVA. Compared to other technologies, FDM was chosen to minimise the costs associated with both the initial purchase and the use.

After testing qualitatively different combinations of the 3D printing parameters, a 30% infill and 1.5 mm shell thickness was found to be a good compromise between model strength and ease of dissolution for Phantom 3, which has the most complex geometry. Lower values of the parameters were chosen for Phantoms 1 and 2, which have a simpler morphology and therefore need less structural support. Namely 15% infill and 1 mm shell thickness.

Differences were noted in the PVA filaments used: *eSun* 2.85 mm resulted to be more flexible and easier to dissolve in water compared to the *Formfutura* PVA 1.75 mm, resulting in a faster casting process (about 30% faster, using the simplified test sections manufactured to select the infill parameter).

The post-processing of the 3D printed physical model with PVA liquid glue was necessary to smooth the surface and reduce the roughness due to the layering printing process. The acrylic spray treatment avoided the interaction between the casting material and the PVA, which would have resulted in opaque/ slightly coloured final cast model. This aspect was of utmost importance for Phantom 1, designed to be optically clear in order to be compatible with PIV measurements.

For both the rigid phantoms, successive layers were poured during the casting process. For Phantom 1 this was necessary to minimise the layer thickness in order to avoid air bubbles remaining trapped in the material after the degassing procedure, whereas for Phantom 3 due to a specific layer thickness limit imposed to minimise the exothermic reaction of the resin.

This aspect may represent a limitation in specific applications, due to the layers becoming distinguishable. For instance, in the case of PIV experiments, the layers used in the casting sequence may pose a challenge in illuminating different planes in the flow; in order to overcome this limitation, in the present study extra care was taken to align the casting layers with the PIV imaging planes.

The choice of casting material when the final phantom has to be compatible with PIV is significantly limited by the requirement of transparency. In this work, Nusil MED

6015 was chosen for Phantom 1 (similar to Sylgard 184, which is often employed in the literature) and proved to be a good choice when the geometry has a contained volume (because of the higher cost of the material). Reproducing the domain of a full patient-specific aorta using the same methodology and material of Phantom 1 would lead to a significant increase in cost. In addition, to reproduce large volumes, since such materials require a degassing procedure after mixing and before casting, a larger vacuum chamber would be needed and further complications would arise due to layering of the material. To overcome these limitations in the manufacturing of Phantom 3, an epoxy resin was used, which facilitates the process overcoming the degassing procedure, reducing the costs and eliminating the need for access to specialist equipment. However, although the material appears to be suitable to manufacture rigid patient-specific phantoms, it is not sufficiently optically clear to be compatible with PIV. A possible solution would be to employ two moulds - a negative and a positive one - and manufacturing a thin wall phantom, such as Phantom 2, in order to minimise the costs of the material. Attention to the mechanical properties would be necessary as demonstrated by the fabrication of Phantom 2 using Smooth-On-Ecoflex 00-30 Silicon in order to reproduce the compliant behaviour of the vessel.

Once cured, for all phantoms, the dissolving of PVA was facilitated using warm and pressurised water and only the inlets and outlets of the geometries were used to allow dissolution (no raisers and runners have been used). The duration of this process was directly related to the complexity of the geometry, from approximately 1 day for Phantom 1 to 3 days for Phantom 3. This phase was also facilitated by mechanical friction inside the phantoms (e.g. using a thin plastic rod), exercising care not to scratch the inner surface of the model.

The anatomical accuracy of Phantom 2 geometry was qualitatively demonstrated through a comparison between the MR-scan and the original stereolithography file. More details about the mechanical characteristics of this phantom and its patient-specific features are described in [28]. Equally, good qualitative similarity was observed between the geometrical features of Phantom 3 CT-scan and the original clinical images (e.g. TL, FL and intimal flap shape, entry-tear dimension and location). Quantitatively, a cross-sectional area error of $7.83 \pm 4.29\%$ was found between the phantom geometry and the original stereolithography file. The comparison demonstrated the ability of the proposed methodology to accurately reproduce complex anatomical geometries.

Regarding imaging modalities, all the three phantoms were MRI compatible but, due to higher requirements, only Phantom 1 was optically clear for PIV. Refractive index matching with the chosen material was successfully realised using a 60%/40% (by weight) water-glycerol mixture (RI = 1.4) as working fluid in the experiments. Figure 8 shows PIV velocity contours measured on the sagittal plane of the geometry to illustrate its compatibility with this imaging modality.

4 Conclusions

Manufacturing of physical phantoms to reproduce vessel geometries is essential to perform *in vitro* haemodynamic studies of the vascular system, test medical devices and surgical procedures, validate computational models and imaging modalities, as well as serve as training tools. Despite the numerous studies described in the literature regarding different phantoms and manufacturing modalities, costs and necessary equipment still present a limitation, especially for laboratories and research groups without extensive expertise in the field.

The aim of this work was to present easier accessible manufacturing options to target this gap and allow non-specialised researchers to manufacture phantoms of different complexities and mechanical properties suitable for different investigation needs. A combination of affordable and accessible methodologies and materials is described for three different case studies of increasing complexity: an idealised aortic arch, a patient-specific aortic arch and a patient-specific diseased aorta. In particular, 3D printing with water-dissolvable PVA was employed to create the vessel - *negative* - mould and casting was performed to create the final physical model. The three cases vary in geometrical complexity and volume dimensions, material mechanical properties (i.e. rigid or flexible), costs involved, equipment needed and imaging modalities compatibility. The different 3D printers, PVA filaments and casting materials were compared and advantages and disadvantages of each discussed.

The described approaches represent cost-effective and highly accessible options to create physical vascular phantoms. Moreover, the case studies showcased a range of possibilities according to different design requirements, demonstrating the flexibility of the adopted methodologies. Further work is necessary to find an equivalent solution to manufacture larger volume patient-specific and ideally compliant phantoms that are PIV compatible.

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References

- [1] Botnar, R., Rappitsch, G., Beat Scheidegger, M., Liepsch, D., Perktold, K., and Boesiger, P., 2000. "Hemodynamics in the carotid artery bifurcation.". *Journal of Biomechanics*, **33**(2), pp. 137–144.
- [2] van Ooij, P., Guédon, A., Poelma, C., Schneiders, J., Rutten, M. C. M., Marquering, H. A., Majoie, C. B., van Bavel, E., and Nederveen, A. J., 2012. "Complex flow patterns in a real-size intracranial aneurysm phantom: Phase contrast MRI compared with particle image velocimetry and computational fluid dynamics". *NMR in Biomedicine*, **25**(1), pp. 14–26.
- [3] Carey, R. F., Herman, B. A., Robinson, R. A., Stewart, H. F., Hoops, R. G., & Douglas, G. H., 1991. U.S. Patent No. 5,052,934.
- [4] Russ, M., OHara, R., Setlur Nagesh, S. V., Mokin, M., Jimenez, C., Siddiqui, A., Bednarek, D., Rudin, S., and Ionita, C., 2015. "Treatment Planning for Image-Guided Neuro-Vascular Interventions Using Patient-Specific 3D Printed Phantoms". pp. 219–227.
- [5] Tsai, T. T., Schlicht, M. S., Khanafer, K., Bull, J. L., Valassis, D. T., Williams, D. M., Berguer, R., and Eagle, K. A., 2008. "Tear size and location impacts false lumen pressure in an ex vivo model of chronic type B aortic dissection". *Journal of Vascular Surgery*, **47**(4), pp. 844–851.
- [6] Plesniak, M. W., and Bulusu, K. V., 2016. "Morphology of secondary flows in a curved pipe with pulsatile inflow". *Journal of Fluids Engineering, Transactions of the ASME*, **138**(10), pp. 1–18.
- [7] Rudenick, P. A., Bijnens, B. H., García-Dorado, D., and Evangelista, A., 2013. "An in vitro phantom study on the influence of tear size and configuration on the hemodynamics of the lumina in chronic type B aortic dissections". *Journal of Vascular Surgery*, **57**(2), pp. 464–474.e5.
- [8] Bonfanti, M., Balabani, S., Greenwood, J. P., Puppala, S., Homer-Vanniasinkam, S., and Díaz-Zuccarini, V., 2017. "Computational tools for clinical support: a multi-scale compliant model for haemodynamic simulations in an aortic dissection based on multi-modal imaging data". *Journal of the Royal Society, Interface*, **14**(136), p. 20170632.
- [9] Bonfanti, M., Balabani, S., Alimohammadi, M., Agu, O., Homer-vanniasinkam, S., and Díaz-zuccarini, V., 2018. "A simplified method to account for wall motion in patient-specific blood flow simulations of aortic dissection : Comparison with fluid-structure interaction". *Medical Engineering and Physics*, **58**, pp. 72–79.
- [10] Franzetti, G., Diaz-Zuccarini, V., and Balabani, S., 2019. "Design of an in vitro mock circulatory loop to reproduce patient-specific vascular conditions: towards precision medicine". *ASME JESMDT*, **2**(November).
- [11] Bonfanti, M., Franzetti, G., Maritati, G., Homer-Vanniasinkam, S., Balabani, S., and Diaz-Zuccarini, V., 2019. "Patient-specific haemodynamic simulations of complex aortic dissections informed by commonly available clinical datasets". *Medical Engineering & Physics*, **71**, pp. 45–55.
- [12] Qin, E. C., Sinkus, R., Geng, G., Cheng, S., Green, M., Rae, C. D., and Bilston, L. E., 2013. "Combining MR elastography and diffusion tensor imaging for the assessment of anisotropic mechanical properties: A phantom study". *Journal of Magnetic Resonance Imaging*, **37**(1), pp. 217–226.
- [13] Medero, R., García-Rodríguez, S., François, C. J., and Roldán-Alzate, A., 2017. "Patient-specific in vitro models for hemodynamic analysis of congenital heart disease – Additive manufacturing approach". *Journal of Biomechanics*, **54**, pp. 111–116.
- [14] Tango, A. M., Salmons-Smith, J., Ducci, A., and Burriesci, G., 2018. "Validation and Extension of a Fluid-Structure Interaction Model of the Healthy Aortic Valve". *Cardiovascular Engineering and Technology*, **9**(4), pp. 739–751.
- [15] Bulusu, K. V., and Plesniak, M. W., 2013. "Secondary flow morphologies due to model stent-induced perturbations in a 180 curved tube during systolic deceleration". *Experiments in Fluids*, **54**(3).
- [16] de Zélicourt, D., Kitajima, H., Yoganathan, A. P., Frakes, D., and Pekkan, K., 2005. "Single-Step Stereolithography of Complex Anatomical Models for Optical Flow Measurements". *Journal of Biomechanical Engineering*, **127**(1), p. 204.
- [17] Biglino, G., Verschuere, P., Zegels, R., Taylor, A. M., and Schievano, S., 2013. "Rapid prototyping compliant arterial phantoms for in-vitro studies and device testing.". *Journal of cardiovascular magnetic resonance : official journal of the Society for Cardiovascular Magnetic Resonance*, **15**(1), p. 2.
- [18] Tyszka, J. M., Laidlaw, D. H., and Asa, J. W., 2000. "Three-dimensional, time-resolved (4D) relative pressure mapping using magnetic resonance imaging - Tyszka - 2000 - Journal of Magnetic Resonance Imaging - Wiley Online Library". *Journal of Magnetic Resonance Imaging*, **329**, pp. 321–329.
- [19] Tai, N. R., Salacinski, H. J., Edwards, A., Hamilton, G., and Seifalian, A. M., 2000. "Compliance properties of conduits used in vascular reconstruction". *British Journal of Surgery*, **87**(11), pp. 1516–1524.
- [20] Tanné, D., Bertrand, E., Kadem, L., Pibarot, P., and Rieu, R., 2010. "Assessment of left heart and pulmonary circulation flow dynamics by a new pulsed mock circulatory system". *Experiments in Fluids*, **48**(5), pp. 837–850.
- [21] Cao, P., Duhamel, Y., Olympe, G., Ramond, B., and Langevin, F., 2013. "A new production method of elastic silicone carotid phantom based on MRI acquisition using rapid prototyping technique". In Proceedings of the Annual International Conference of the IEEE Engineering in Medicine and Biology Society, EMBS, IEEE, pp. 5331–5334.
- [22] Yazdi, S. G., Geoghegan, P. H. ., Docherty, P. D., Jermy, M. ., and Khanafer, A. ., 2018. "A Review of Arterial Phantom Fabrication Methods for Flow Measurement Using PIV Techniques". *Annals of Biomedical Engineering*

- neering, **46**(11), pp. 1697–1721.
- [23] Shmueli, K., Thomas, D. L., and Ordidge, R. J., 2007. “Design, construction and evaluation of an anthropomorphic head phantom with realistic susceptibility artifacts”. *Journal of Magnetic Resonance Imaging*, **26**(1), pp. 202–207.
- [24] Geoghegan, P. H., Buchmann, N. A., Spence, C. J. T., Moore, S., and Jermy, M., 2012. “Fabrication of rigid and flexible refractive-index-matched flow phantoms for flow visualisation and optical flow measurements”. *Experiments in Fluids*, **52**(5), pp. 1331–1347.
- [25] Allard, L., Soulez, G., Chayer, B., Treyve, F., Qin, Z., and Cloutier, G., 2009. “Multimodality vascular imaging phantoms: A new material for the fabrication of realistic 3D vessel geometries”. *Medical Physics*, **36**(8), pp. 3758–3763.
- [26] Bulusu, K. V., and Plesniak, M. W., 2018. “Insights on arterial secondary flow structures and vortex dynamics gained using the MRV technique”. *International Journal of Heat and Fluid Flow*, **73**(July), pp. 143–153.
- [27] Bücking, T., Hill, E., Robertson, J., Maneas, E., Plumb, A., and Nikitichev, D., 2017. “From medical imaging data to 3D printed anatomical models”. *PLoS ONE*, **12**(5), pp. 1–10.
- [28] Gallarello, A., Palombi, A., Annio, G., Momi, E. D., and Maritati, G., 2019. “Patient-Specific Aortic Phantom With Tunable Compliance”. *ASME JESMDT*, **2**(November), pp. 1–12.

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Table 1. Technical specifications and comparison of key manufacturing features among the three phantoms

	Phantom 1	Phantom 2	Phantom 3
Geometry	Simplified aortic arch	Patient-specific aortic arch	Patient-specific diseased aorta
Material	Rigid	Compliant	Rigid
Segmentation	/	3D Slicer	ScanIP
3D printer	Delta Wasp 2040 Turbo 2	Delta Wasp 2040 Turbo 2	BCN3D Sigmax
Build volume	∅ 200 mm x h 400 mm	∅ 200 mm x h 400 mm	420 mm x 297 mm x 210 mm
3D printing material	Formfutura PVA 1.75 mm	Formfutura PVA 1.75 mm	PVA eSun 2.85 mm
3D filament cost	\$45 / 300g	\$45 / 300g	\$38 / 500g
Casting material	Nusil MED 6015	Smooth-On Ecoflex™ 00-30 silicon	Epoxy resin
Casting material cost	\$154 / 600 ml	\$26 / 400 ml	\$89 / 5000 ml
Amount of casting material used	2000 ml	300 ml	5000 ml
Other equipment needed	Small vacuum chamber	External mould (\$254)	None
PIV compatibility	Yes (RI=1.4)	No	No
MRI compatibility	Yes	Yes	Yes

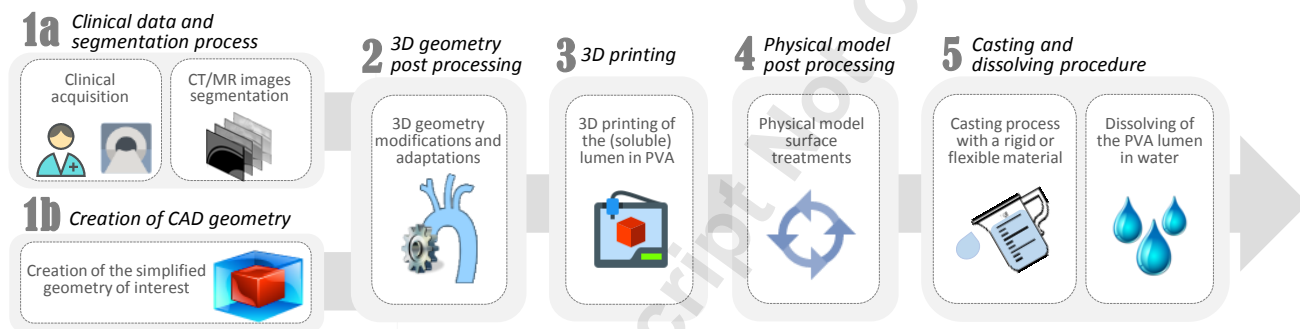


Fig. 1. The proposed PVA based manufacturing process: (1a) acquisition of patient-specific clinical images of the vessel of interest and segmentation process to reconstruct the geometry; (1b) creation of the CAD simplified geometry; (2) post-processing of the 3D geometry; (3) 3D printing of the vessel lumen in PVA; (4) Physical model refinement; (5) Casting and dissolving procedure of PVA in water

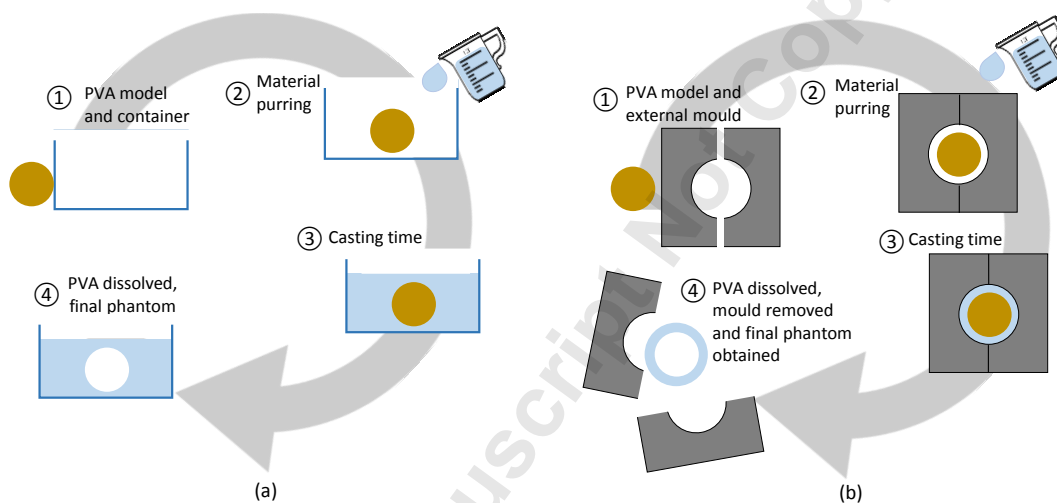


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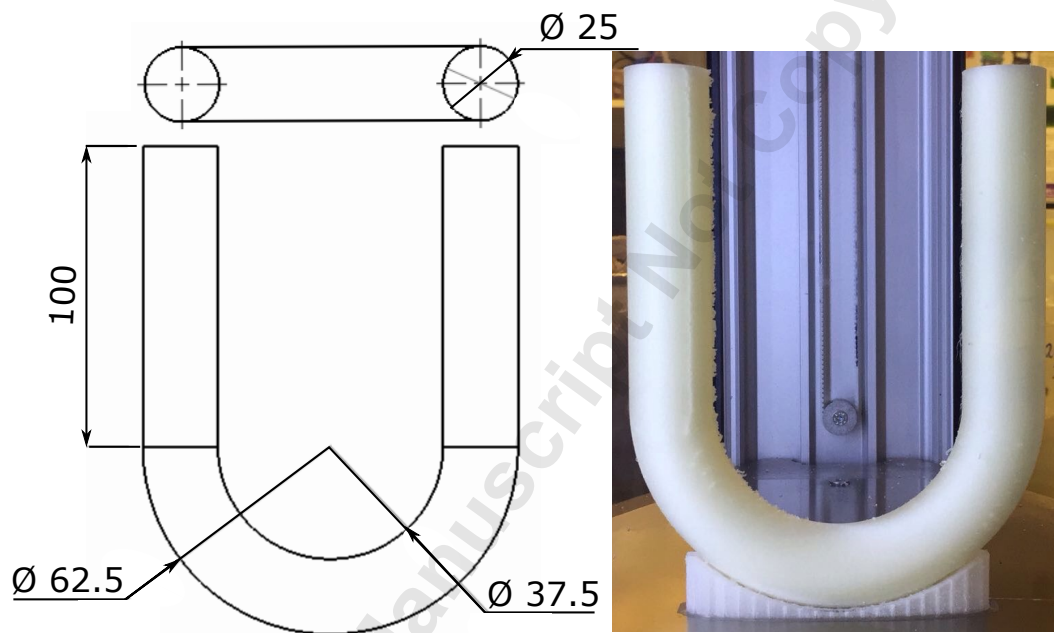


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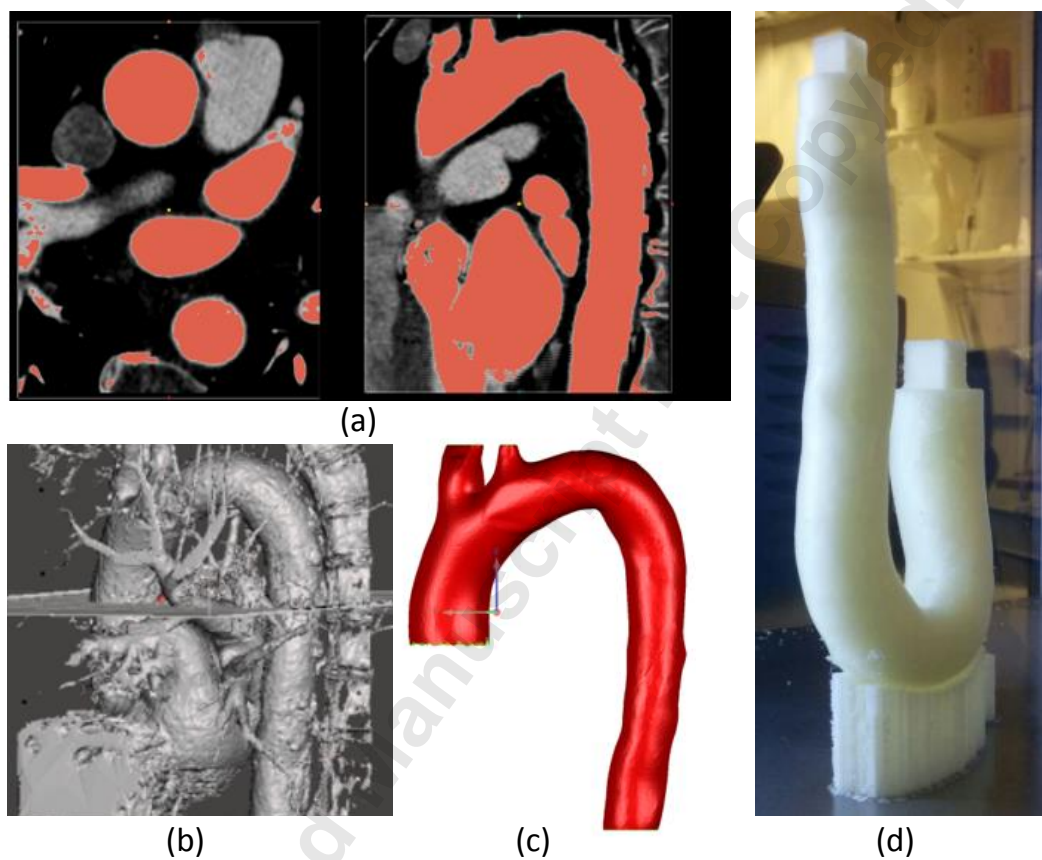


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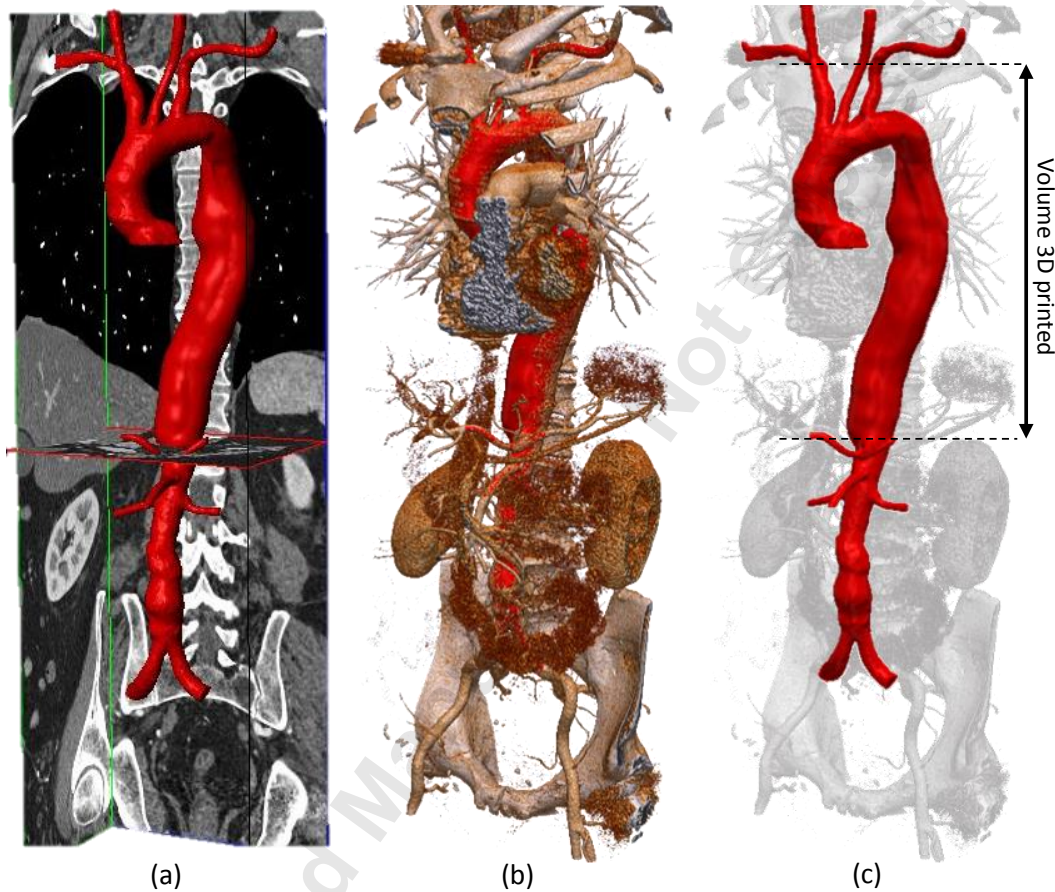


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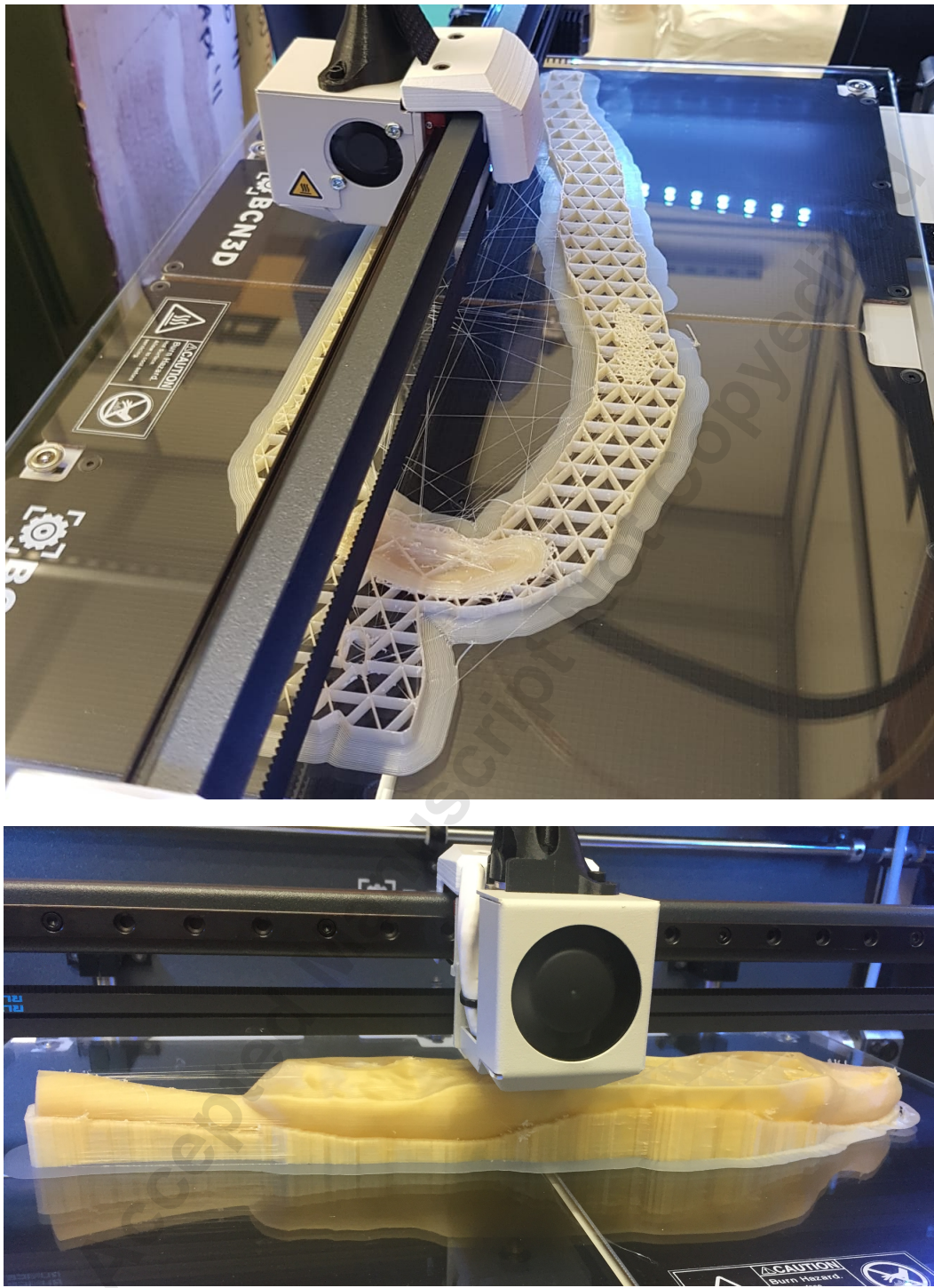


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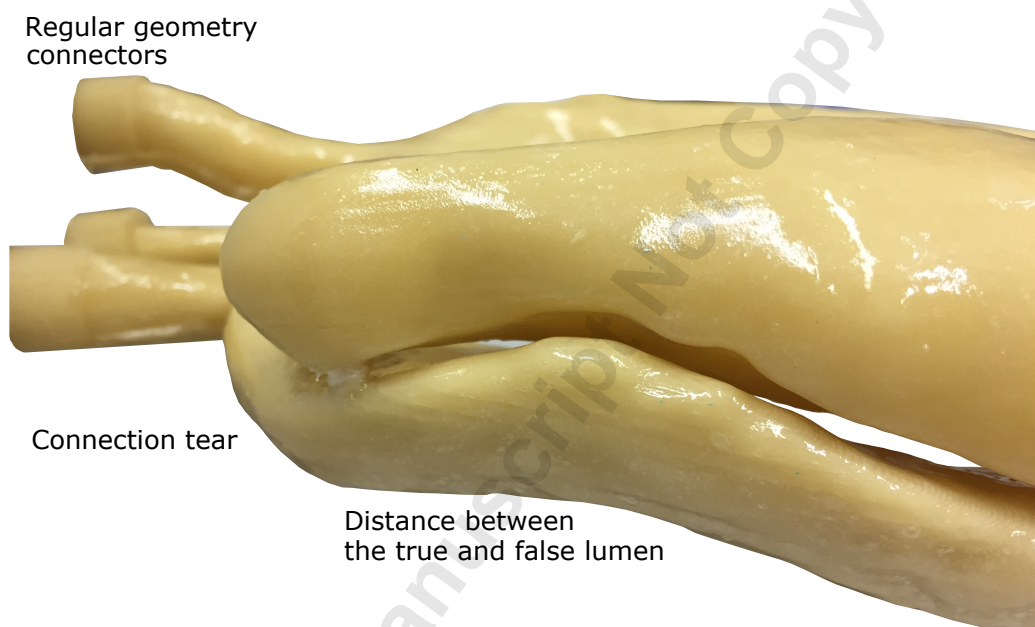


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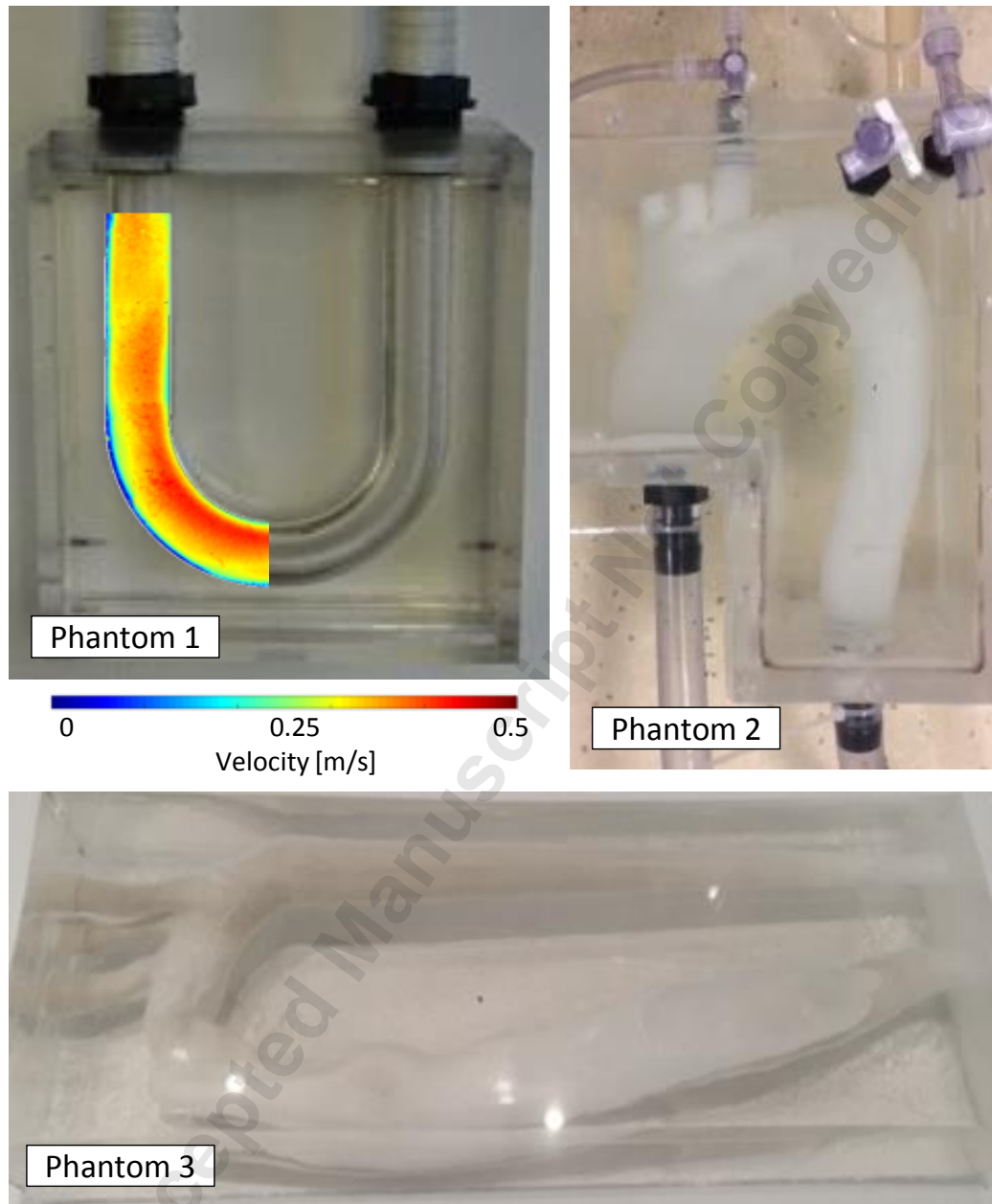


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