

## **Development of a D-SPECT myocardial perfusion phantom**

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Revision 0.2

Project plan



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## **Preface**

The project plan outlines an introduction and literature of the topic along with organisational information including a detailed planning.

Gijs de Vries  
Enschede, 3<sup>rd</sup> December 2018



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# 1 Introduction

[done] Read into background information on D-SPECT

[done] Write global background information

[done] Introduce the rest of the document

[done] Assignment was for dynamic SPECT scanning, but is that the same as using the D-SPECT? The D-SPECT can scan dynamically, and is available in ZGT

[done] Too much SPECT detail in introduction? Moved to literature

[todo] Give arguments why to choose SPECT

Myocardial Perfusion Imaging (MPI), or, simply put, the imaging of the blood flow in the heart muscle, plays an important role in diagnosing heart failure or detecting Coronary Artery Disease (CAD). Imaging systems like Computed Tomography (CT), Magnetic Resonance Imaging (MRI), Single-Photon Emission Computed Tomography (SPECT), or Positron Emission Tomography (PET) can visualise a (radioactive) contrast bolus in the supplying arteries and in underlying myocardial tissue, whose flow can give an indication of narrowed or blocked blood vessels.

Many variations in the visualisation process of myocardial perfusion, including variations in hard- and software, can (significantly) influence the outcome and in turn have consequences for patient treatment. These variations need to be validated against a well-known baseline.

## Document overview

[done] Update in correspondence with meeting december 10

The project plan consists of a literature review of existing myocardial perfusion phantoms, their comparison to human physiology, and a discussion between the different types of scanners. The literature is followed by the research methodology containing the research questions and goals of the project. The detailed planning is the last section of the project plan stating work-days and -weeks, off-days, deadlines, and meetings.

## Abbreviations

**AIF** Arterial Input Function

**CAD** Coronary Artery Disease

**CT** Computed Tomography

**CZT** Cadmium Zinc Telluride

**MPI** Myocardial Perfusion Imaging

**MRI** Magnetic Resonance Imaging

**PA** Pulmonary Artery

**PET** Positron Emission Tomography

**PET-MR** PET-Magnetic Resonance

**PMT** Photomultiplier Tube

**PV** Pulmonary Vein

**SPECT** Single-Photon Emission Computed Tomography

**VC** Vena Cava

## 2 Literature

[inpr] Read available literature

[inpr] Write literature review to more accurately define research questions

[inpr] D-SPECT literature?

[todo] Discuss division of work

- Dialysis tube mimics capillaries and not tissue?
- Mathys reported that no literature was found that manufacturers calibrate their scanners.

### 2.1 Phantoms

#### 2.1.1 Magnetic Resonance

[inpr] Shift the limitations to later section

Noguchi et al. (2007) developed a simple phantom that consists of a syringe, diameter of 40mm, beads, and tubes, diameter of 4mm. The perfusate, 0.1mM GD-DTPA doped 8L water solution, flows through the beads, to disturb the flow, and then perfuses through parallel tubes, to prevent cross-current. The perfusate is tagged at the beads while the images are taken at 5 parallel planes, perpendicular to the tubes. The actual measured flow is the average obtained from the parallel planes.

Ebrahimi et al. (2010) created a phantom using microfabrication to create a microvasculature on 4-inch ( $\approx 10$ cm) silicone wafers. The microvasculature is build up from four blocks, containing 4x2 (RxC) cells. Each cell is build up from 100 "features" separated by 25 $\mu$ m tracks. These tracks provide many different paths for perfusate to flow, effectively simulating capillaries in tissue. A 300 $\mu$ L bolus of a distilled water solution containing 25mM/L of Manganese.

Wang et al. (2010) used a haemodialysis filter connected to a nonpulsatile pump. A static water phantom was placed next to the haemodialysis filter to show that it cancels out between tag (with magnetic labelling) and control (no magnetic labelling) images.

Anderson et al. (2011) extracted hollow fibres from haemodialysis filters to create their own single-fibre and multi-fibre phantom. Similar to some standard haemodialysis filters, their phantoms have access to both the extracellular space (i.e. the fluid outside of the fibres) and intracellular space (i.e. the fluid inside the fibres). As the name suggests, their single-fibre phantom consists of an individual fibre placed inside a capillary tube. The multi-fibre phantom consists of a variable amount of fibres that are placed in a heat-shrink tube. A four-way valve switched between the main perfusate and a aqueous solution of 0.2 mM GD-BOPTA.

Chiribiri et al. (2013) developed a four chambered anatomic phantom that resembles the heart of a 60kg person. The four chambers correspond to the four chambers of the heart, sized to match the physiological size. In addition, a Vena Cava (VC), Pulmonary Artery (PA)/Pulmonary Vein (PV) combination, and an aorta are present in the phantom. Contrast is injected in the same manner as is performed in patients; in a vein. In the phantom, the contrast is injected directly into the vena cava via a three-way stopcock. The contrast moves through the phantom's right atrium, right ventricle then via the PA/PV to the left chamber and finally to the left ventricle. The phantom is not capable of simulating the contrast's behaviour in the lungs since the PA is directly connected to the PV. Two myocardial chambers (one active and one control), the VC, PA, PV and the aorta are in the imaging plane where the proximal part of the aorta, where the aorta branches to the myocardium, is used for the Arterial Input Function (AIF).

Otton et al. (2013) used the same phantom to compare CT against MRI. In addition to the previous authors, O'Doherty et al. (2017a) used a water-filled torso phantom to ensure more anatomically correct image in PET-Magnetic Resonance (PET-MR).

### 2.1.2 Computed Tomography

Teslow and Robb (1991) developed a cylindrical perfusion phantom, shaped like the left ventricle of a dog. It has a 6.5cm outer diameter, 4.5cm inner diameter, and a length of 6.5cm. The authors specifically used methyl methacrylate plastic since it gives similar radiographic image as tissue and blood. Additionally, a solid plastic cylinder, placed in the centre, is used to attenuate x-ray beams similar to the attenuation of a blood-filled dog's ventricle. The capillaries are simulated by means of different sized nylon balls of 0.318, 0.476, and 0.635cm. The smallest balls are packed near the outlet, while the medium sized balls are packed at the inlet, and the largest balls are placed in between. The authors do not go into detail on the used contrast agent, other than 10ml of a radio-opaque indicator is injected for one second.

Klotz and König (1999)

Driscoll et al. (2011) developed a 10cm long, with a diameter of 5cm, phantom with two inputs, only one is used, and two outputs. The capillaries are simulated by means of a vinyl tube where mass can be exchanged with the main cylinder via sets of small holes on either side of the cylinder. The output of the vinyl tube and the output of the shell combine, and feeds back through the imaging area to validate outward going flow, which should be the same as the input flow. Their AIF, however, is created by means of a programmable pump which injects in the form of a typical clinical AIF rather than having a system-based AIF. The Visipaque<sup>TM</sup> (iodixanol) 270 mgI/ml is injected into a blood-mimicking fluid consisting of 40% glycerol and 60% water.

Ganguly et al. (2012) developed an interestingly, slightly different phantom than other; a linearly moving phantom. The phantom itself is a cylinder, 1.9cm inner diameter, with a length of 32.2cm. It contained 64 different compartments of 0.5cm in height separated by a 0.5mm thick carbon fibre wall. Every compartment had a single opening where contrast, 300mgI/Omnipaque, is injected. The concentration of contrast that is injected in subsequent compartments, resemble a sinusoidal signal. This phantom is placed on a linear motor which moves the phantom parallel to the patient's bed. The author's goal is to compare and determine the temporal accuracy of the imaging system. To simulate the attenuation of the head, a 15cm cylindrical, water-filled, phantom was placed around the perfusion phantom.

Mathys et al. (2012) developed a similar phantom that consists of two cylinders, a smaller (4cm diameter) inside a larger (11cm diameter) cylinder. Water flows via four tubes into the inner cylinder where it flows outward into the larger cylinder that contains 1.5mm polyoxymethylene globes as tissue replacement. The larger cylinder is drained by four holes. 2mL of contrast, Accupaque 300, is injected followed by 15 of saline at 5mL/s using a double-head injector.

Boese et al. (2013) developed a cylindrical phantom for brain perfusion measurements in a C-arm. Their phantom utilises a combination of large arteries, smaller arteries, and a sinter board for the capillaries. The main artery splits into smaller arteries which in turn splits into sixteen even smaller ones that connect to the sinter board. The upper two arteries have an inner diameter of 1.7mm representing the carotid arteries and the lower two arteries have an inner diameter of 1.0mm representing vertebral arteries. The authors used a very specific contrast injection protocol: relatively large pre-injection of 21mL NaCl, followed by a variable amount of Imeron 400, and ended by a 6mL post-injection of NaCl, all at 6mL/s.

Suzuki et al. (2017) designed a straight-forward CT phantom that uses a dry-type haemodialyser with a pressurised dialysate space to prevent the perfusate from leaving the hollow fibres. The authors varied the dose in order to determine the effects on the perfusion indices. They main-

tained a constant volumetric flow,  $Q$ , and concluded that the perfusion indices are susceptible to dose conditions.

Hashimoto et al. (2018) used the same phantom in combination with a commercially synthetic bone layer such that quantification software recognises the phantom as a human head. Instead of varying the dose, the contrast injection protocol and the scanning interval are varied based on their hypothesis that it would increase the quantitative accuracy. However, they concluded that they are independent factors when using the b-SVD algorithm.

### 2.1.3 Ultrasound

#### [done] Veltmann phantom

Veltmann et al. (2002) designed a flow phantom that consists of a high- and low flow circuit. The high flow circuit consists of a *heated* reservoir flowing into a haemodialysis cartridge, which filters any residue micro-bubbles (contrast agent) and removes air bubbles, before entering a second haemodialysis cartridge, the perfusion cartridge. Perfusate that does not enter the capillaries is returned to the reservoir passing a variable resistance. The perfusate that does enter the capillaries of the perfusion cartridge, is controlled by a gear pump, which simultaneously acts as a variable flow resistance for the low flow circuit. After the gear pump, a third haemodialysis filter filters the microbubbles from the perfusate. The authors performed two different experiments, one with an unmodified haemodialysis filter and one with a haemodialysis filter that has the majority of the lower capillaries glued shut. The contrast agent tends to float, especially in the low flow circuit. By decreasing the number of perfused capillaries, the flow is made more homogeneous and avoids attenuation in the lower areas. Both Sakano et al. (2015) and Lohmaier et al. (2004) use this phantom setup.

Kim et al. (2016) performs perfusion experiments using ultrasound without adding any contrast. Similar to the CT and MR phantoms, a dialysis tube is used to mimic human capillaries. The dialysis tube is submerged in water and part of the plastic case was removed, replaced by a latex foil as proposed by Veltmann et al. (2002), such that it creates an acoustic window. More interestingly, Kim et al. (2016) use a secondary, 1Hz, peristaltic pump to simulate cardiac motion. Gauthier et al. (2011) uses a peristaltic pump after a renal dialysis cartridge to create a pulsatile, but constant, flow. They do not use a secondary pump for the extracellular space.

### 2.1.4 Positron Emission Tomography / Single-Photon Emission Computed Tomography

Although the phantoms are not specifically designed for PET or SPECT scanning, the previously mentioned phantoms can be an inspirational source for PET/SPECT phantoms. The different technology requires a new approach to some (or many) of the materials used.

### 2.1.5 Phantom discussion

The phantom by Chiribiri et al. (2013) is unable to simulate the diffusion of contrast into heart tissue or the interstitial space, as admitted by the authors and confirmed by Otton et al. (2013); O'Doherty et al. (2017a). Furthermore, Chiribiri et al. (2013) mentioned that the blood flow resistance is lower than in patients due to its complexity.

The findings of Otton et al. (2013) are similar; the contrast curves represent those obtained from clinical trials and since the phantom can be used in a clinical MR scanner, the gap between phantom and clinical studies is decreased. Even with the addition of a water-filled torso phantom, it is still unable to mimic respiratory or cardiac motion (O'Doherty et al., 2017b).

- [Is contrast absorbed by tissue in the brain? Read something about the blood-brain barrier preventing such things.](#)

The straight forward phantom of Suzuki et al. (2017) does not resemble the human brain, which caused problems in certain programs, and the capillary possessions is much greater than in

clinical situations. This may ultimately compromise the reliability of the phantom to mimic clinical situations. Although Hashimoto et al. (2018) uses a commercially synthetic bone layer, the phantom does not simulate contrast uptake by surrounding tissue which does occur in myocardial perfusion measurements.

## 2.2 Physiology

## 2.3 Technology

### 2.3.1 SPECT

The imaging method in a typical SPECT scanner are scintillator-based gamma cameras, also known as Anger cameras. Gamma cameras use a scintillator to "transduce" gamma radiation, originating from an injected tracer, to photons. Part of these photons are directed towards a photocathode. If a quantum of light hits the photocathode, which is coated with a photosensitive coating, electrons are emitted due to the photoelectric effect. These electrons travel through a Photomultiplier Tubes (PMTs) and hitting series of dynodes, which in turn trigger secondary emission effectively multiplying the number of electrons travelling through the tube. Electrons hitting the last dynode, which is known as the anode, cause a current pulse which can be detected by measuring equipment. It is proportional to the amount of gamma ray photons entering the scintillator (GE Healthcare, 2009).

### 2.3.2 Digital SPECT

Developments in imaging systems gave rise to the digital SPECT scanner. In contrast to the analogue Anger cameras, the digital SPECT scanner utilises a direct conversion semiconductor: Cadmium Zinc Telluride (CZT). Wagenaar (2004) used CZT to develop pixelated detector units which could then be used for medical imaging. In a recent study, it is shown that a digital SPECT scanner, using multiple pixelated CZT detectors, showed significant improvements in image sharpness and contrast (Goshen et al., 2018). These detector units do not require any PMTs and thus allow for a more compact and flexible design (Erlandsson et al., 2009). The D-SPECT scanner, a digital SPECT scanner developed by Spectrum Dynamics<sup>1</sup>, offers improvements in sensitivity and energy resolution (Spectrum Dynamics, 2016) over Anger camera systems. However, these digital systems are relatively new and require proper validation to convince medical personnel of its value.

### 2.3.3 Scanner comparison

As is previously mentioned, there are various types of scanners that use different techniques, CT MRI, or Scintigraphy based (SPECT/PET) scanners. In cardiology, the SPECT scanner is widely employed for coronary and myocardial perfusion measurements (Rahmim and Zaidi, 2008). It is known that PET scans are generally more expensive (Hlatky et al., 2014; Goel et al., 2014). Hlatky et al. (2014) followed patients for two years, recording the costs and concluded that PET costs are 22% higher than the costs for SPECT for patients with suspected CAD.

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<sup>1</sup><https://www.spectrum-dynamics.com/>

## 3 Research methodology

[inpr] Define research questions

[inpr] Discuss main research question

[todo] Define research boundaries

[done] Define project goals

[todo] Implement scanner descision

[todo] Use limitations of phantoms to lead to research question

Different perfusion phantoms have been developed. Some are specifically designed to simulate the myocardial perfusion (Chiribiri et al., 2013; Otton et al., 2013; O'Doherty et al., 2017a,b; Teslow and Robb, 1991), others for brain perfusion (Hashimoto et al., 2018; Suzuki et al., 2017; Boese et al., 2013; Mathys et al., 2012; Ebrahimi et al., 2010; Wang et al., 2010; Noguchi et al., 2007; Ganguly et al., 2012; Klotz and König, 1999), or capillaries in general (Kim et al., 2016; Anderson et al., 2011; Driscoll et al., 2011; Gauthier et al., 2011; Veltmann et al., 2002; Lohmaier et al., 2004), and even phantoms to simulate perfusion in rheumatoid finger joints (Sakano et al., 2015) and skin (Kim et al., 2018). Most software packages are designed for human organs and look for recognition points such that it can perform the most optimal calculation. Part of these phantoms do not resemble the human anatomy of the parts they simulate, which will influence the reliability.

### 3.1 Research questions

The project is carried out using the V-Model (Osborne et al., 2005) approach, see figure 3.1. The research questions are defined based on the project definition phase.

#### Concept of operations

What must the myocardial perfusion phantom be able to simulate?

#### Requirements and Architecture

What are the requirements for a myocardial perfusion phantom that can be used in combination with commonly used clinical software?

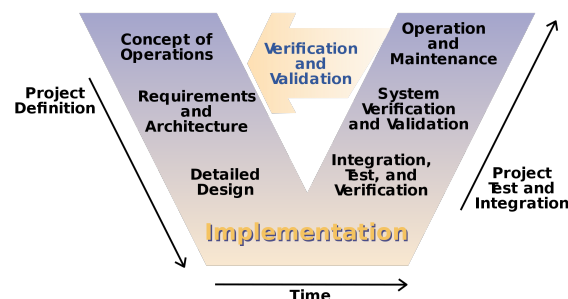


Figure 3.1: V-Model as proposed by Osborne et al. (2005)

## Detailed Design

How can the myocardial perfusion phantom meet the clinical requirements and mimic the perfusion of a human heart?

### 3.2 Project goal

The goal of the project is to develop a prototype myocardial perfusion phantom capable of repeated simulations of typical and cardiac defect situations using clinical software commonly used in myocardial perfusion scans. Most software packages require anatomical recognition points which imposes anatomical requirements on the phantom. In addition, the phantom can be used for educational and training purposes to demonstrate the impact of (poorly) chosen variables, e.g. pressure or flow, scanning parameters, cardiac defects, and so forth.

#### Project sub-goal

During the individual project, a calibration set-up for flow and pressure sensors has been developed. The prototype version showed that flow sensors can be calibrated using the "emptying tank" principle. Pressure sensors have not been implemented in the prototype. Furthermore, the calibration set-up relies too much on human interaction. A sub-goal of the project will be to improve the existing calibration set-up such that flow and pressure sensors can be calibrated which in turn will increase the reliability of the flow set-up.

#### 3.2.1 General concept

A general concept is shown in figure 3.2. Three main parts can be identified: flow set-up, phantom, and imaging device. The flow set-up consists of everything to produce and maintain pressures and/or flows and measure these variables. The user-interface is a computer or laptop. The phantom consists of everything needed to mimic cardiac defects and to provide a representative image for myocardial perfusion image processing software. The imaging set-up consists of the imaging device itself along with any contrast agents needed to properly image the phantom. Many imaging devices communicate with a dedicated workstation on which the image processing software runs.

### 3.3 Additional resources

During the individual project of Gijs de Vries, a prototype flow set-up, control module, and calibration set-up has been realised and can be used / re-cycled.

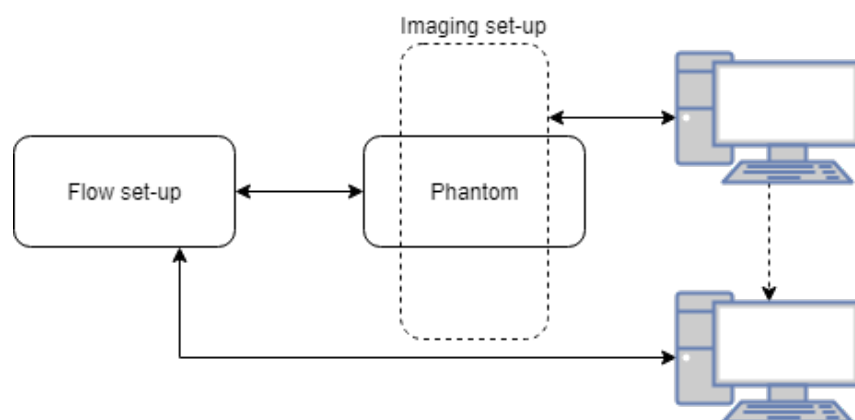


Figure 3.2: General concept, schematic

### 3.3.1 Flow set-up

The flow set-up uses simple, low costs pumps and available flow sensors. These might not be suitable for high precision flow systems. It is encouraged to look into alternatives.

### 3.3.2 Control module

The control module requires improvement when it is to be re-used. Two of the main improvements consist of improving the electro(magnetic) shielding to decrease the susceptibility to noise and to optimise the pump controllers.



## 4 Planning

[done] Create graphical planning

[done] Create workday overview

[done] Create week overview

[done] Define deadlines

[inpr] Define meetings: frequency, type, and already planned

This chapter details the planning for the 40ECTS final thesis, carried out under the Robotics and Mechatronics Chair of the University of Twente. The project will be carried out in two phases; proof-of-concept (phase 1) and definitive (phase 2). The Gantt planning for phase 1 and 2 can be found in appendix B in figures B.1 and B.2 respectively.

### 4.1 Workdays

The planning is based on 28 hours per European Credit as per Dutch standard. The final thesis is carried out full-time (40 hours per week). The overview of working hours is shown in table 4.1.

Day	Start time	End time	Productive hours
Monday	08 <sup>30</sup>	16 <sup>00</sup>	7
Tuesday	08 <sup>30</sup>	17 <sup>00</sup>	8
Wednesday	08 <sup>30</sup>	16 <sup>00</sup>	7
Thursday	08 <sup>30</sup>	17 <sup>00</sup>	8
Friday	08 <sup>30</sup>	17 <sup>00</sup>	8
Miscellaneous*			2
<b>Total:</b>			40

\* Miscellaneous hours are in evenings, weekends or during train rides.

**Table 4.1:** Workdays and -hours

### 4.2 Work weeks

[done] Discuss work days between christmas and new-year

[done] Discuss work days on holidays

The work weeks can be found in table A.1 in appendix A.

The project planning spans 35 weeks. Activities are planned from week 49 of 2018 up until, and including, week 28 of 2019 which spans a total of 32 weeks. Week 29 will be used to finalise practical aspects; handing in material and documentation, report printing, and so forth. The graduation presentation (and ceremony) will additionally take place in week 29. Weeks 30 and 31 of 2019 can serve as an extension if, and only if, approved by the assessment- and exam committee.

The planning takes into account one week around Christmas and new-years, one week spring break ("voorjaarsvakantie") in 2019, and a two week buffer. See section 4.3 for more details.

### 4.3 off-days

The University of Twente recognises three general holidays, New Year's day, King's Birthday and Liberation day, and six Christian holidays, Good Friday, Easter Monday, Ascension day, Whit Monday, Christmas day, and Boxing day<sup>1</sup>. Furthermore, the university recognises five bridging days in 2018 and four bridging days in 2019<sup>2</sup>.

Both the King's Birthday as well as Liberation day fall in weekends. The remainder of the holidays and bridging days are summarised in table 4.2.

Holiday	Date	Note
<b>Bridging day</b>	2018 December 24	<i>Collective closure<sup>2</sup></i>
<b>Christmas day</b>	2018 December 25	<i>Christian holiday<sup>1</sup></i>
<b>Boxing day</b>	2018 December 26	<i>Christian holiday<sup>1</sup></i>
<b>Bridging day</b>	2018 December 27	<i>Collective closure<sup>2</sup></i>
<b>Bridging day</b>	2018 December 28	<i>Collective closure<sup>2</sup></i>
<b>Bridging day</b>	2018 December 31	<i>Collective closure<sup>2</sup></i>
<b>New Year's day</b>	2019 January 1	General holidays <sup>1</sup>
<b>Good Friday</b>	2019 April 19	<i>Christian holiday<sup>1</sup></i>
<b>Easter Monday</b>	2019 April 22	<i>Christian holiday<sup>1</sup></i>
<b>Ascension Day</b>	2019 May 30	<i>Christian holiday<sup>1</sup></i>
<b>Bridging day</b>	2019 May 31	<i>Collective closure<sup>2</sup></i>
<b>Whit Monday</b>	2019 June 10	<i>Christian holiday<sup>1</sup></i>

**Table 4.2:** Off-days

Week 4 of 2019 is a planned vacation and no work will be done. This off-week spans from Monday 21<sup>st</sup> of January 2019 until, and including, Friday 25<sup>th</sup> of January 2019.

[done] Update time of lectures

Currently, three lectures are planned which will result in an absent from the workplace in order to follow these lectures. These lectures are summarised in table 4.3.

What	Day	Date	When	Where
<b>CT lecture</b>	Thursday	2018 December 20	13 <sup>45</sup> - 17 <sup>30</sup>	Waaier 2
<b>PET lecture</b>	Thursday	2019 January 10	10 <sup>45</sup> - 12 <sup>30</sup>	HR C101
<b>PET/SPECT Radiology</b>	Monday	2019 January 14	08 <sup>45</sup> - 12 <sup>30</sup>	NH207

*\* Times will be updated when known*

**Table 4.3:** Planned lectures

<sup>1</sup> <https://www.utwente.nl/en/ces/planning-schedules/academic-calendar/holidays-closing-days/>

<sup>2</sup> <https://www.utwente.nl/en/hr/terms-of-employment/scope-of-employment/public-holidays-leave-days/#compulsory-leave-days>

## 4.4 Deadlines

The deadlines for phase 1 are shown in table 4.4 and those for phase 2 are shown in table 4.5.

What	R	Day	Date	Note
<b>Project plan</b>	0.1	Friday	2018 December 20	Before 2018 December 10
	0.2	Tuesday	2018 December 18	Before 2018 December 19
	1.0	Friday	2018 December 21	Before Christmas
<b>System</b>	0.1	Friday	2019 January 11	
<b>Requirements</b>	0.2	Friday	2019 February 1	
	1.0	Friday	2019 February 8	
<b>Design</b>	concept	Friday	2019 March 1	
	choice	Monday	2019 March 4	
	final	Friday	2019 March 15	Parallel development
<b>Realisation</b>		Friday	2019 March 29	Including testing

*Deadlines subject to change depending on weekly meetings*

**Table 4.4:** Deadlines phase 1

What	R	Day	Date	Note
<b>Project plan</b>	1.1	Friday	2019 April 5	
	2.0	Friday	2019 April 12	
<b>System</b>	1.1	Friday	2019 April 26	
<b>Requirements</b>	2.0	Friday	2019 May 3	
<b>Design</b>	concept	Friday	2019 May 24	
	choice	Monday	2019 May 27	
	final	Friday	2019 June 7	
<b>Realisation</b>		Friday	2019 July 12	Including testing
<b>Final report</b>	0.1	Friday	2019 June 14	
	0.2	Friday	2019 June 28	
	1.0	Friday	2019 July 12	

*Deadlines subject to change depending on weekly meetings*

**Table 4.5:** Deadlines phase 2

## 4.5 Meetings

- [Plan weekly progress meetings](#)

Two types of progress meetings will be regularly planned. Weekly progress meetings between Marije Kamphuis and Gijs de Vries, and progress meetings every four to six weeks where Kees Slump joins. The weekly progress meetings will, unless otherwise discussed, take place on ....

What	Day	When	Participants
<b>Progress meeting</b>	Monday	2018 December 10 14 <sup>00</sup>	Gijs de Vries, Marije Kamphuis, Kees Slump
<b>Progress meeting</b>	Wednesday	2018 December 19 15 <sup>30</sup>	Gijs de Vries, Marije Kamphuis, Kees Slump
<b>Orientation intro ZGT</b>	Wednesday	2018 January 2*	Gijs de Vries, Marije Kamphuis
<i>To be filled</i>			

*\* will be planned on this day.*

*Will be updated when new meetings are planned. Does not relate to weekly recurring meetings*

**Table 4.6:** Planned meetings

345

## A Appendix: Work weeks

Week	Monday	Working	Note
49	2018 December 3	Yes	
50	2018 December 10	Yes	
51	2018 December 17	Yes	
52	2018 December 24	Partly	<i>See off-days</i>
1	2018 December 31	Mostly	<i>See off-days</i>
2	2019 January 7	Mostly	<i>CT college</i>
3	2019 January 14	Mostly	<i>PET college</i>
4	2019 January 21	No	<i>Vacation</i>
5	2019 January 28	Yes	
6	2019 February 4	Yes	
7	2019 February 11	Yes	
8	2019 February 18	Yes	
9	2019 February 25	Yes	
10	2019 March 4	Yes	
11	2019 March 11	Yes	
12	2019 March 18	Yes	
13	2019 March 25	Yes	
14	2019 April 1	Yes	
15	2019 April 8	Yes	
16	2019 April 15	Mostly	<i>See off-days</i>
17	2019 April 22	Mostly	<i>See off-days</i>
18	2019 April 29	Yes	
19	2019 May 6	Yes	
20	2019 May 13	Yes	
21	2019 May 20	Yes	
22	2019 May 27	Mostly	<i>See off-days</i>
23	2019 June 3	Yes	
24	2019 June 10	Mostly	<i>See off-days</i>
25	2019 June 17	Yes	
26	2019 June 24	Yes	
27	2019 July 1	Yes	
28	2019 July 8	Yes	
29	2019 July 15	Yes	
30	2019 July 22	No	<i>Extension when needed</i>
31	2019 July 29	No	<i>Extension when needed</i>

**Table A.1:** Work weeks

## B Appendix: Gantt planning

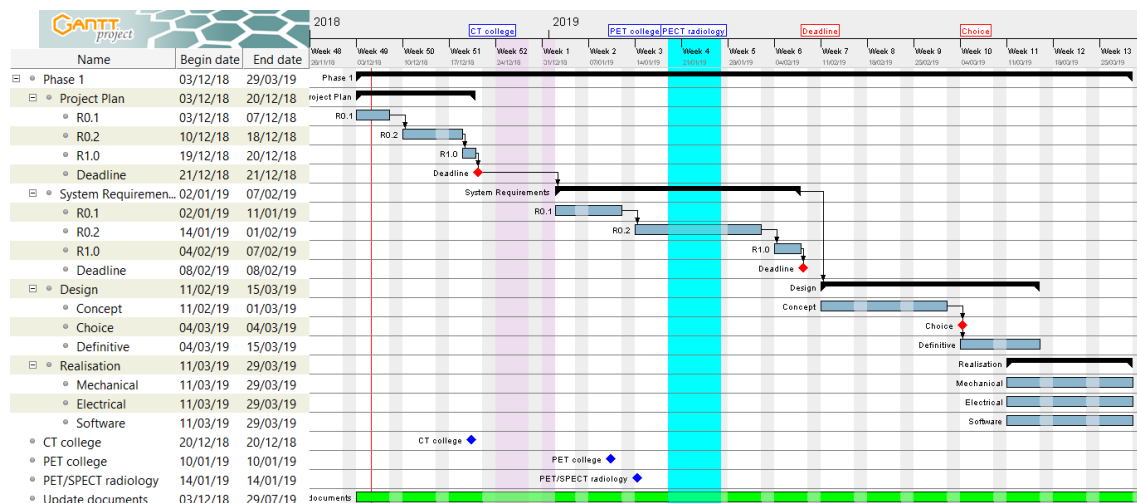


Figure B.1: Phase 1 project planning

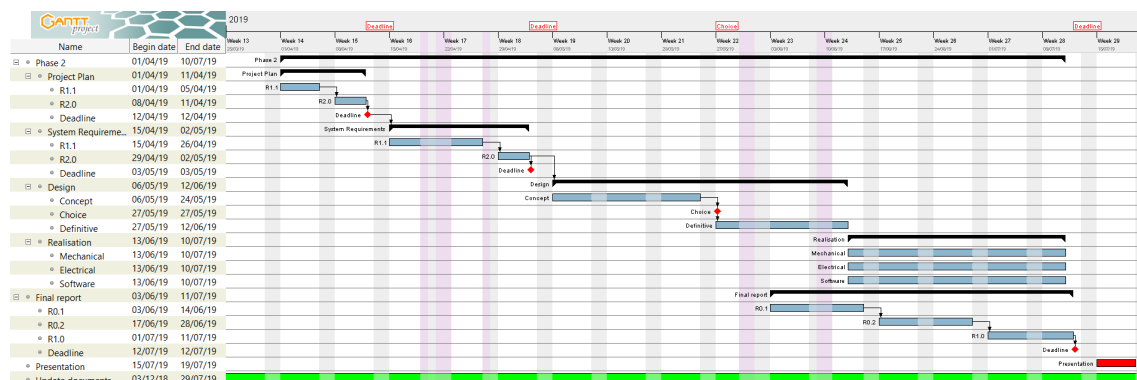


Figure B.2: Phase 2 project planning

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