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Fast MRI registration leveraging k-space properties

Schnelle MRT-Registrierung mittels k -Raum Eigenschaften

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Eidesstattliche Erklärung

Ich erkläre hiermit an Eides statt, dass ich diese Arbeit selbständig verfasst und keine anderen als die angegebenen Quellen und Hilfsmittel benutzt habe.

Jan Meyer

Zusammenfassung

Irgendwas über MRT und Registration

Abstract

Something about MRI and registration

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1

Introduction

MRI is a commonly used medical imaging technique that is non-invasive, radiation-free and has great contrast for soft tissue. However, speed is its main weakness as full-body scan can take up to 30 minutes, which is a burden on patients as well as hindering efficiency. Thus, MRI acquisition is usually accelerated by subsampling the k-space in which the data is recorded. This can, however, lead to problematic image artifacts that hinder many processing algorithms. Another problem of the slow data acquisition are motion artifacts, which are especially common for organs like lung and heart. While breathing can be controlled, cardiac motion is involuntarily and should not be stopped. These challenges have been traditionally addressed with computationally intensive algorithms that usually iteratively solve an optimization problem. New approaches, however, are based around the deep neural networks, which have seen a rise in popularity in the recent years in many fields including image processing. In this thesis, the possibility of using an unsupervised deep learning approach to align subsampled MRI data as well its potential usage in a motion reconstruction pipeline will be investigated.

1.1 Contributions of the Thesis

We managed to implement an unsupervised approach to correct motion in subsampled MRI images and much more.

1.2 Related Work

There are many papers on image registration in general, however in the context of medical image registration with deep learning their number is reduced due to the specialized nature of the subject at hand. Yet, there are a couple of papers that give a good overview of the topic. They give a brief overview of registration methods, basics of deep learning with already existing networks for image registration as well as covering potential applications and challenges [1, 2, 3, 4, 5]. A lot of different approaches for medical image registration are based on *VoxelMorph* [6], such as both *Fourier-Net* [7] and its successor *Fourier-Net+* [8].

1.3 Structure of the Thesis

In chapter 2, the general basis of the thesis containing descriptions of MRI in section 2.1, Image registration in section 2.2 and deep learning architectures as well as their use in image registration in sections 2.3 and 2.4. A special emphasize is put on motion reconstruction in MRI as described in section ??.

The data used in this thesis is described in Chapter 3 with details concerning the *OASIS* and *CMRxRecon* datasets in sections 3.1 and 3.2 respectively.

In chapter 4, the networks used in this thesis, namely *Fourier-Net* and *Fourier-Net+*, are described in section 4.1 and 4.2.

The experiments conducted in the making of this thesis are explained in chapter 5 with results and discussion following in chapter 6.

Lastly, the thesis is briefly summarized in chapter 7.

2

Basics

In this chapter the basics of the thesis are explained.

2.1 Magnetic Resonance Imaging

Magnetic Resonance Imaging (MRI) is a non-invasive, radiation-free, tomographic imaging technology based on measurements of a magnetic field. An MRI machine comprises four main components as seen in Figure ?? . The first component is a strong magnet powerful enough to generate a static magnetic field B_0 that is required to induce nuclear proton polarization. The second is a radio frequency (RF) system which generates an alternating magnetic field B_1 at the resonant frequency f and detects the MR signal that is returned from the patient. The third component is the set of gradient systems (oriented orthogonally in X, Y and Z directions) that generates linear magnetic field variations, which are then superimposed upon B_0 and are used to spatially encode the MR signal. In clinical MR scanners, the three gradient sets and whole-body RF coils are typically concentrically positioned inside the bore of the magnet. The fourth component is a computer providing the user interface, generating images to be displayed and interpreted on the console [9].

Physical Foundation

Individual hydrogen nuclei precess around the field B_0 at a resonance frequency known as the Larmor frequency ω_0 of the net magnetization vector, which generally occurs in the RF range of the electromagnetic spectrum and is related to the external magnetic field as:

$$\omega_0 = \gamma \cdot B_0, \quad (2.1)$$

with γ being the gyromagnetic ratio, which is $42.5 \frac{\text{MHz}}{\text{T}}$. When hydrogen nuclei are placed in the presence of a strong static magnetic field such as B_0 the nuclei split into two energy states, either aligned parallel to the magnetic field B_0 (called a spin-up state) or aligned anti-parallel to B_0 (called a spin-down state). The spin-up state has a slightly lower energy level as compared to the spin-down state and is therefore preferred. This slight difference in the spin states (0.001%) results in an overall net magnetization M aligned in the same

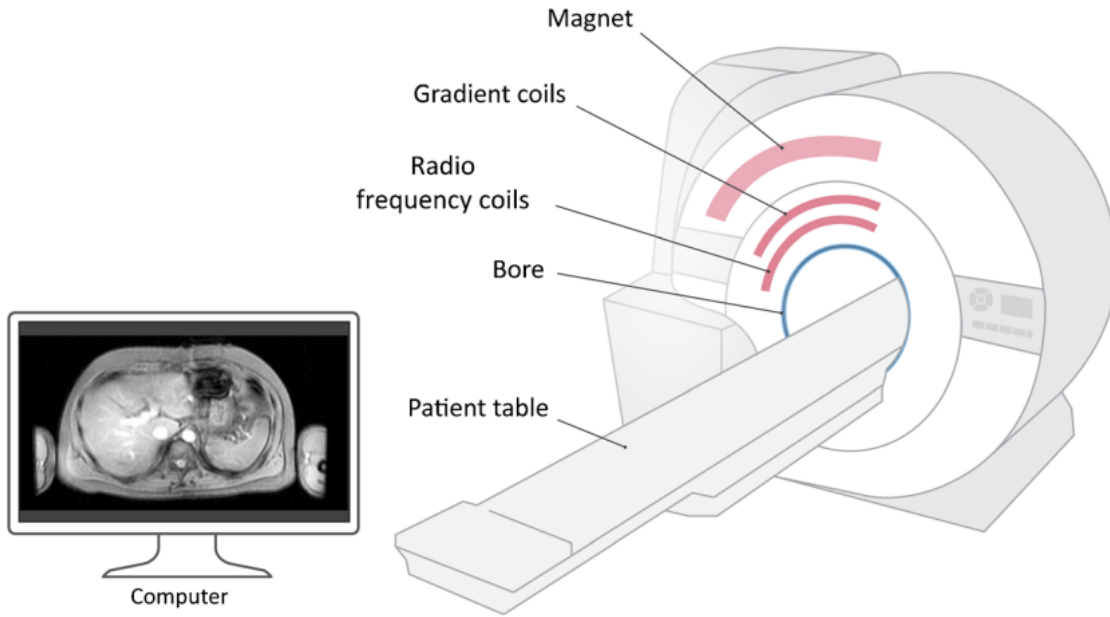


Figure 2.1: Schematic of an MRI scanner taken from [9].

direction as B_0 . To create an MR signal, the spins are excited out of their resting equilibrium, i.e. tipping M away from B_0 . To detect the signal from the hydrogen nuclei in the tissues an additional external field B_1 is introduced at the resonant Larmor frequency ω_0 that can affect magnetization vector, causing it to rotate into a plane orthogonal to its original orientation. The rotated vector continues to precess around the B_0 . The precession of the magnetization vector in the transverse plane can be detected by an RF coil tuned to the resonant frequency ω_0 . RF coils can be operated in a receive-only mode, in which case the inherent body coil is used as a transmitter; or the RF coils can be both transmit and receive. The purpose of the RF transmit coil is to create a time-varying B_1 field at right angles to B_0 that could be linearly or circularly polarized. The closer the receiving coil is to the source of the MR signal, the better the signal-to-noise ratio (SNR). Receiver coils generally comprise arrays of smaller individual coils or elements; however, each individual coil element has a limited depth penetration. Multiple arrays of coils, termed phased-array coils, can be used together to achieve a higher coverage. The multiple coils are electronically decoupled from one another so that they do not appear as just a single large coil. The images from individual coils are independently reconstructed and then grouped together to create the final image. An RF pulse of amplitude B_1 , called excitation pulse, is applied for a certain time duration to tip the magnetization at an angle away from the B_0 field. The precessing transverse magnetization induces a voltage in the receiver RF coil; this induced voltage is known as the free induction decay. After the pulse, the magnetization returns to thermal equilibrium by processes known as MR relaxation. To fully encode the spatial information within the field of view (FOV), pulse sequences must be iterated numerous times. The time between successive iterations of a pulse sequence is known as the repetition time (TR). The time between the application of the initial RF pulse and the middle of the detected echo is known as the echo time (TE).

Due to this the overall time to acquire an MR image is quite long [9].

Once the RF pulse is turned off, M continues to precess as it returns to its thermal equilibrium state. During this time, two types of relaxation occur: T1 (longitudinal or spin lattice) and T2 (transverse or spin/spin). One attribute that makes T1 and T2 so valuable for determining the signal in MRI is their sensitivity to the presence and type of tissue. It is this tissue dependence property that gives MR its excellent soft-tissue contrast. T1 relaxation describes the recovery of the longitudinal magnetization back to thermal equilibrium following a perturbation by an RF pulse. The longitudinal component regrows along the Z direction with a time constant T1. In other words, after the RF pulse is turned off, the protons that were disturbed give their energy to the surrounding environment and return back to their original equilibrium state, realigning with B_0 . Hence T1 relaxation is also called longitudinal relaxation. Furthermore, because T1 relaxation involves the loss of energy that was put into the spin system by the RF pulse, it is also referred to as spin-lattice relaxation, the lattice consisting of surrounding macromolecules. This loss of energy is stimulated by the fluctuating magnetic fields associated with the dipole–dipole interactions of neighboring magnetic moments. T1 relaxation can only occur when these magnetic field fluctuations occur at the resonant frequency ω_0 . The rate at which the spin magnetization M_z recovers to M_0 at time t is called T1 relaxation time. It can be expressed as follows:

$$M_z = M_0 \cdot \left(1 - e^{-\frac{t}{T1}}\right). \quad (2.2)$$

A preferred method of measuring T1 relaxation is the Look-Locker method, which is based on the principle that one does not need to wait for the net magnetization vector to equilibrate in order to measure T1. Instead, an RF pulse is used with a small flip, which can be repeatedly applied. The acquisition pulse sequence is designed to generate a train of signals that gradually approaches a steady-state recovery. The recovery curve measured by this technique can be fitted to an exponential curve to provide an effective T1 measurement. This has become one of the most popular T1-mapping method for abdomen and cardiac imaging. The general principle of T1 mapping is to acquire multiple images with different T1 weightings and to fit the signal intensities of the images to the equation of T1 relaxation. T1 relaxation values for tissues can be estimated by fitting the data to the following equation:

$$S = S_0 \cdot \left(1 - A \cdot e^{-\frac{TI}{T1}}\right), \quad (2.3)$$

where S is the signal intensity measured at each inversion time value TI , S_0 is the initial signal intensity at time $t = 0$ and A is a constant.

T2 relaxation results in the loss of transverse magnetization caused by interactions between the magnetic fields of neighboring hydrogen nuclei. It is not an energy loss process like T1 but is a loss of phase coherence within the spin system. This process, also known as spin-spin relaxation, leads to the destruction of transverse magnetization and causes the magnetic moments of the tissue to dephase. T2 mapping is a method of measuring the T2 value of the tissue. T2 relaxation time can be calculated using a T2 sequence with different echo times (TEs). The most fundamental sequence for T2 mapping is signal measured

with spinecho techniques (multiple sequences with different TE values) [19]. Other 2-D sequences have been used, such as multi-echo spin echo (MSME) and fast spin echo (FSE) [19]. Synthetic MRI is another quantitative method in which a single saturation recovery turbo spin-echo sequence is used to estimate T2 transverse relaxation [20]. T2 relaxation values for tissues can be estimated by fitting the data to the following equation using a mono-exponential decay curve:

$$S(TE) = S_0 \cdot e^{-\frac{TE}{T_2}}, \quad (2.4)$$

where $S(TE)$ is the signal intensity measured at each TE and S_0 is the initial signal intensity at time $t = 0$.

Image Acquisition

Imaging Acceleration

To alleviate the slow image acquisition times of the MR imaging, different MRI acceleration techniques were used. Most of these included a subsampling of the k-space, meaning the omission of certain higher frequencies, from which a images is reconstructed. While there are different subsampling techniques, most involve fully sampling a center region containing the lower frequencies and dropping higher frequencies, which are deemed less important. However, all of these methods create artifacts during image reconstruction leading to different technologies being developed to minimize these effects such as compressed sensing (CS) and parallel imaging (PI).

2.2 Image Registration

Image registration is a challenging, yet important task for image processing. It can be described as the process of transforming different image datasets into one coordinate system with matched imaging contents [2]. In the medical field this can be used for clinical applications such as disease diagnosis and monitoring, image-guided treatment delivery, and post-operative assessment. Medical image registration is typically used to pre-process data for tasks like object detection (for e.g. tumor growth monitoring) and segmentation (for e.g. organ atlas creation) where variation in spatial resolution is common between modalities like CT and MRI and patients. Thus the performance of these methods is dependent on the quality of image registration [1].

Medical image registration was often done manually by clinicians, however, registration tasks are often challenging and the quality of manual alignments is dependent on the expertise of the user. These manual registrations are thus not only time consuming, but also hardly reproducible leading to high interobserver-variability. The need for automatic registration is very much apparent, but this task remained hard to solve for a long time, requiring a lot of computational power and time for computer algorithms to solve the problem. While neural networks also require a lot of computational power and time to train, they promise fast execution after training. With the rise of deep learning these network gained popularity and now pose a real alternative to conventional algorithms and

manual registration [2]. We will discuss these new approaches in the next section, but first we need to formally define our problem.

In pair-wise image registration two images (F and M) are to be aligned, with F denoting the fixed and M the moving image. T is the desired spatial transformation that aligns the two images. This can be posed as an optimization problem:

$$T' = \arg \max S(F, T(M)), \quad (2.5)$$

with T' being the best transformation that maximizes the similarity S between the two images. This process is done iteratively improving estimates for the desired T , such that the defined similarity in the cost function is maximized [1].

Transformations can be categorized as rigid, affine, and deformable. A rigid transformation consists of rotation and translation; an affine transformation includes translations, rotations, scaling, and sheering; the two kinds of transformations are described as a 2D single matrix. Unlike rigid and affine transformation, deformable transformation is a high-dimension problem that we need to formulate by a 3D matrix for 2D deformable registration i.e., a so-called deformation field. While rigid and affine registration algorithms have already achieved good performance in many applications, deformable registration is still a challenging task due to its intrinsic complexity, particularly when the deformation is large. However, these are also the transformations most likely encountered in clinical practice as it can be utilized to fuse information from different modalities such as MRI and CT [4]. Additionally, deformable image registration can also be utilized for various computer-assisted interventions like biopsy [10] and (MRI-guided) radiotherapy [11, 12].

Intuitively, deformable image registration is an ill-posed problem, making it fundamentally different from other computer vision tasks such as object localization, segmentation or classification. Given two images, deformable image registration aims to find a spatial transformation that warps the moving image to match the fixed image as closely as possible. However, there is no ground-truth available for the desired deformation field and without enforcing any constraints on the properties of the spatial transformation, the resulting cost function is ill-conditioned and highly non-convex. In order to address the latter and ensure tractability, all image registration algorithms regularize the estimated deformation field, based on some prior assumptions on the properties of the underlying unknown deformation [1].

Many methods have been proposed for medical image registration to deal with the complex challenges of this task. Popular conventional registration methods include optical flow [13], demons [14] and many more. However, most of these still lack accuracy and computation speed, which makes newer deep learning approaches all the more interesting [3].

2.3 Deep Learning Architectures

Neural networks, despite the theoretical concepts being around for decades, have seen a meteoric rise in popularity over the last few years as constraints on computational power

have been alleviated. Especially deep neural networks, which are often summarized under the term deep learning (DL). Recent years have witnessed an almost exponential growth in the development and use of DL algorithms, sustained thus far by rapid improvements in computational hardware (e.g. GPUs). Consequently, clinical applications requiring image classification, segmentation, registration, or object detection/localization, have witnessed significant improvements in algorithmic performance, in terms of accuracy and/or efficiency [1]. The following network architecture are widely used for different tasks including medical image registration.

Some basic stuff about network training, testing and different architectures that are relevant for the later Chapters.

Convolutional Neural Networks

Convolutional neural networks (CNNs) are a type of deep neural networks with regularized multilayer perceptron, which are mainly used for image processing. CNNs use convolution operations instead of general matrix multiplications in typical neural networks. These convolutional filters make CNNs very suitable for visual signal processing. Because of their excellent feature extraction ability, CNNs are some of the most successful models for image analysis. Different variants of CNN have been proposed and have achieved the-state-of-art performances in various image processing tasks. A typical CNN usually consists of multiple convolutional layers, max pooling layers, batch normalization layers, sometimes dropout layers, a sigmoid or softmax layer. In each convolutional layer, multiple channels of feature maps are extracted by sliding trainable convolutional kernels across the input feature maps. Hierarchical features with high-level abstraction are extracted using multiple convolutional layers. These feature maps usually go through multiple fully connected layer before reaching the final decision layer. Max pooling layers are often used to reduce the image sizes and to promote spatial invariance of the network. Batch normalization is used to reduce internal covariate shift among the training samples. Weight regularization and dropout layers are used to alleviate data overfitting [3]. The loss function is often defined as the difference between the predicted and the target output. CNNs are usually trained by minimizing the loss via gradient back propagation using optimization methods like Adam [15].

U-Net

The U-Net [16] architecture is an extension of the typical CNNs structure typically used for image segmentation, however it can also be used for image registration tasks. It adopts symmetrical contractive and expansive paths with skip connections between them. The encoding blocks on the left extract important features from the image using convolution layers and max pooling, which are then stored in the latent space in the middle. From there it is reconstructed using upsampling and convolutions in the decoding blocks on the right. Additionally, skip connections are used to improve the spatial resolution of the segmentation. This architecture allows effective feature learning from a small number of training datasets [3].

Autoencoders

An autoencoder (AE) is a type of CNNs that learns to reconstruct an image from its input without supervision. AEs usually consists of an encoder which extracts the input features, which are stored a low-dimensional latent state space, similar to a U-Net, and a decoder which restore the original input from the latent space. To prevent an AE from learning an identity function, regularized autoencoders were invented, which can be used for e.g. denoising AEs. Variational AEs (VAEs) are generative models that learn latent representation using a variational approach, which constrains the variability of the outputs. VAEs can be used for anomaly detection and image generation [3].

Generative Adversarial Networks

Generative adversarial networks (GANs) consist of two competing networks, a generator and a discriminator. The generator is trained to generate artificial data that approximate a target data distribution from a low-dimensional latent space similar to an AE. The discriminator is trained to distinguish the artificial data from actual data. The discriminator encourages the generator to predict realistic data by penalizing unrealistic predictions via learning. Therefore, the discriminative loss could be considered as a dynamic network-based loss term. The generator and discriminator both are getting better during training to reach Nash equilibrium. In medical imaging, GANs have been used to perform image synthesis for inter- or intra-modality, such as MRI to synthetic CT and vice versa. In medical image registration, GANs are usually used to either provide additional regularization or translate multi-modal registration to uni-modal registration [3].

2.4 Deep Learning for Image Registration

Recently, there has been a surge in the use of deep learning based approaches for medical image registration. Their success is largely due to their ability to perform fast inference, and the flexibility to leverage auxiliary information such as anatomical masks as part of the training process. The most effective methods, such as *VoxelMorph* [6], typically employ a U-Net style architecture to estimate dense spatial deformation fields. These methods require only one forward pass during inference, making them orders of magnitude faster than traditional iterative methods. Following the success of *VoxelMorph*, numerous deep neural networks have been proposed for various registration tasks [8]. Other approaches also utilize CNNs, AEs and GANs. Typical strategies are discussed in more detail in the following sections.

Supervised Registration

Supervised registration describes training a network with a ground truth displacement field that is either real (created by hand) or synthetic (generated via traditional iterative registration algorithms). Thus the loss can easily be calculated as the difference in the displacement fields of the network prediction and the ground truth (see Figure 2.2 for a visual overview). These methods have achieved notable results with real displacement

fields as supervision. However, this approach is very limited by the size and the diversity of the dataset. As the displacement fields are often calculated by conventional algorithms their effectiveness might be limited for difficult problems with which the traditional algorithms struggle. Fully supervised methods are widely studied and have notable results, but the generation of real or synthetic displacement fields is hard, and these displacement fields might be different from the real ground truth, which can impact the accuracy and efficiency of these kinds of methods [4]. Notable approaches include *BIRNet* [17] and *LAPNet* [18], however the latter works in the k -space domain.

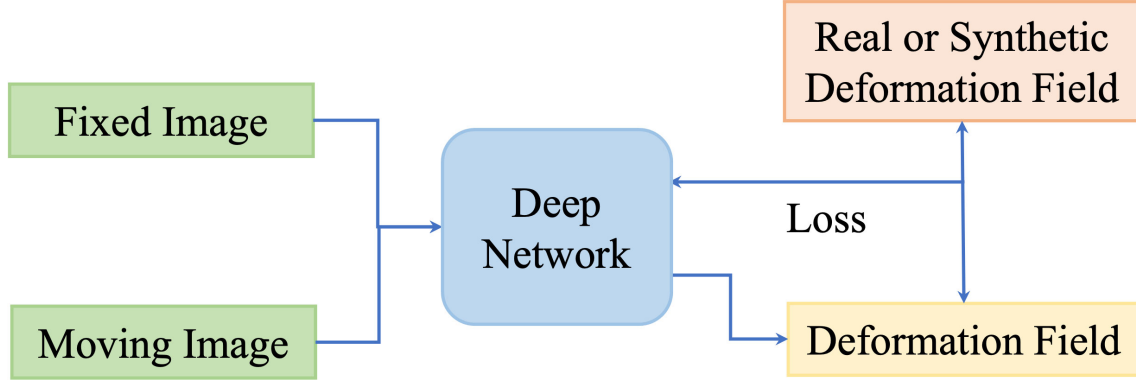


Figure 2.2: Example graph illustrating the training process of a supervised network, taken from [4].

Unsupervised Registration

As the preparation of the ground truth displacement field for supervised methods is inconvenient, limitations in generalizing results in different domains and various registration tasks are inevitable. Thus, unsupervised registration has a more convenient training process with paired images as inputs, but without a ground truth. Generally, unsupervised learning consists of similarity-based (see Figure 2.3) and GAN-based methods (see Figure 2.4), where the loss function computes the similarity between the aligned images and the smoothness of the displacement field, rather than the difference to a ground truth [4]. Well known examples are *IC-Net* [19], *VoxelMorph* [6], *TransMorph* [20] and *SYM-Net* [21].

Evaluation Metrics

Different metrics can be used for evaluation of registration performance including similarity metrics and other stuff. The mean squared error (MSE) is calculated between a fixed (ground truth) image and the warped image giving a pixel-wise comparison:

$$\text{MSE} = \frac{1}{N} \sum_{i=1}^N |F(x, y) - W(x, y)|^2, \quad (2.6)$$

where F is the fixed image and W is the warped image with N being the number of pixels in the images. The lower the MSE the higher the similarity with 0 being a perfect match

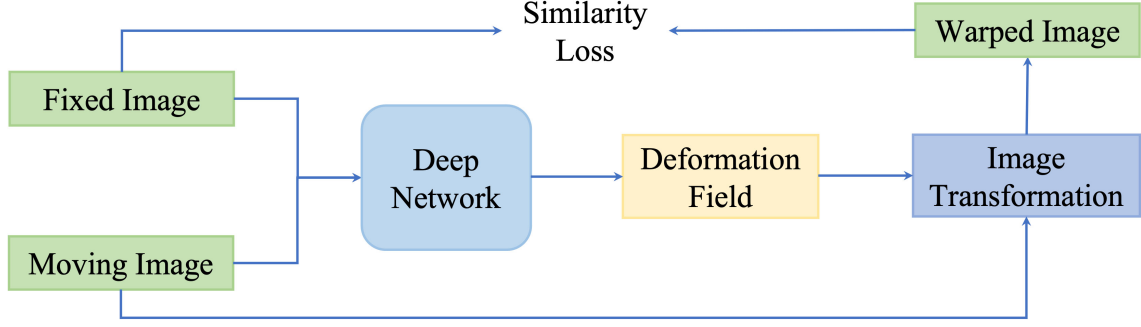


Figure 2.3: Example graph illustrating the training process of an unsupervised network with only a image similarity loss, taken from [4].

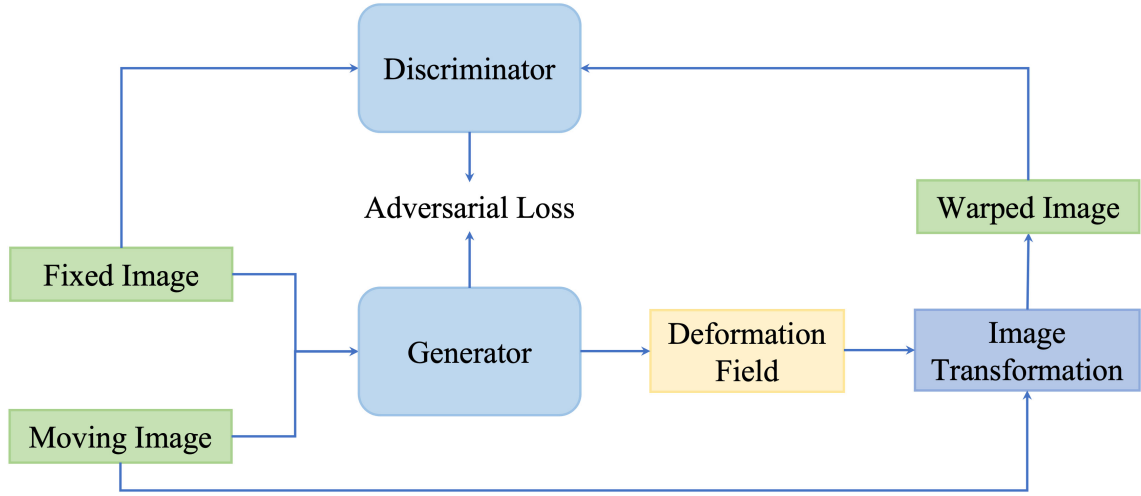


Figure 2.4: Example graph illustrating the training process of a GAN, taken from [4].

(the images are exactly the same).

Another common image metric is the structural similarity index measure (SSIM), which operates between 1 (complete similarity) and 0 (no similarity) and tries to estimate the general similarity instead of a pixel-wise comparison making it more robust against contrast changes compared to e.g. MSE.

$$\text{SSIM} = \frac{(2\mu_F\mu_W + c_1) \cdot (2\sigma_{FW} + c_2)}{(\mu_F^2 + \mu_W^2 + c_1) \cdot (\sigma_F^2 + \sigma_W^2 + c_2)}, \quad (2.7)$$

where μ_F, μ_W are the mean values of the images F and W ; σ_F^2, σ_W^2 the variances of F and W as well as σ_{FW} the covariance for F and W , with c_1, c_2 being constants derived from the dynamic range of the images.

All of these metrics can also be used as loss functions for network training.

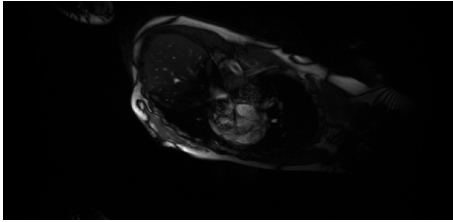
For comparison of segmentations the Dice score is a commonly used metric to estimate the similarity of two segmentations. A score of 1 indicates a complete overlap/match and a score of 0 indicates no overlapping of the segmentations. The Dice score is calculated

be compensated for in reconstruction, however, many methods are restricted to rigid or affine motion due either to computational issues or to the difficulty of modeling complex displacement fields [23]. The latter however seems to become less of a problem as image registration improves with e.g. the help of deep learning.

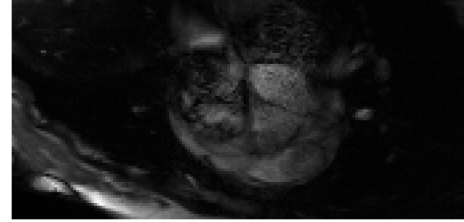
Motion-resolved data acquisition for these applications are usually accelerated by Parallel Imaging or Compressed Sensing techniques yielding sub-Nyquist sampled (subsam-pled) k-space data. In order to reconstruct aliasing-free images, these methods rely on reconstruction schemes that, for example incorporate sparsity or low-rank constraints to solve the ill-posed problem. Fixed sparsity assumptions in Compressed Sensing are often too restrictive and incapable of fully modeling spatio-temporal dynamics. Careful fine-tuning between regularization and data consistency is required and especially in highly subsampled cases residual aliasing may remain in the image (under-regularization) or staircasing and blurring artifacts can occur (over-regularization) which affect the image registration. After reconstruction, non-rigid motion fields can be estimated in image space from reconstructed images by solving a registration problem. A particular interest and challenge lies in the derivation of reliable motion fields which capture the spatio-temporal non-rigid deformations, such as respiratory or cardiac movement. Instead of performing these two steps sequentially, motion-compensated image reconstruction schemes like *GRICS* [23] integrate both motion field estimation and motion correction into the reconstruction process. These methods require reliable motion-resolved images from which the motion fields can be estimated. Motion field estimation can be controlled or supported by external motion surrogate signals, initial motion field estimates, from motion-aliased images or low-frequency image contents. Moreover, spatio-temporal redundancies can be exploited to achieve an aliasing-free image. While these methods have been proven to be more robust against registration errors, they can require a significantly increased computational demand and/or limit imaging acceleration [24].

There are also approaches for learning new subsampling strategies in a data-driven manner (pruning unimportant k-space frequencies) [25] as well as deep learning based radial [26] and non-Cartesian [27] subsampling for MRI acceleration.

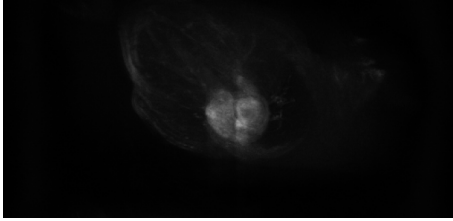
calculated and averaged for each slice. According to the first assumption this process is repeated for all slices and their results summed to create a sort of heatmap as seen in Figure 3.4c with a clearly higher intensity in the cardiac region compared to the background. To further isolate this region a simple thresholding is performed followed by an opening operation, which removes any smaller fragments still left in the image. The cardiac region is now clearly separated from the rest of the image as seen in Figure 3.4d. The only step left is to calculate the approximate center of this region and create a crop around it. To drastically reduce the image size a cropping factor of one third was used, reducing the image size from 246×512 to 82×170 . The result of this drastic crop can be seen in Figure 3.4b when compared to the original image in Figure 3.4a. Note, however, that the cropped image in Figure 3.4 appears larger than it actually is, when compared to the original, to allow for a more detailed look. Thus both images are not presented to scale, but in order to show the most image detail.



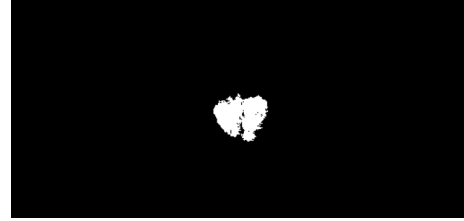
(a) The original full-size image.



(b) The corresponding cropped image.



(c) Sum of the differences between slices and their respective frames with the cardiac region in the middle.



(d) Mask of the cardiac region generated from the sum of differences by thresholding and opening.

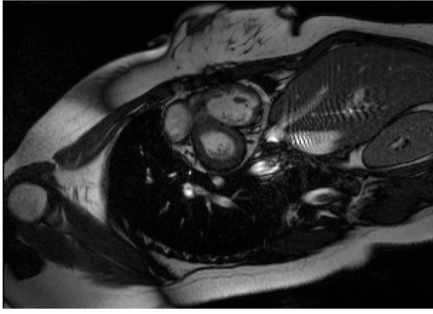
Figure 3.4: Cropping of an example image by locating the movement between frames for each slice. Note that the cropped image appears larger than it actually is compared to the original.

3.3 ACDC Dataset

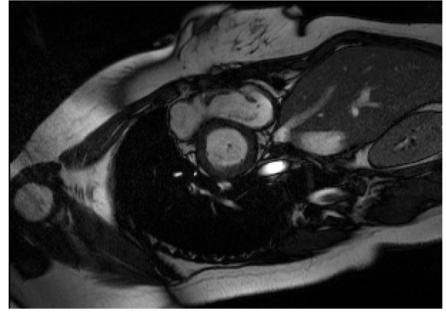
The ACDC dataset [34] from the *Automated Cardiac Diagnosis Challenge (ACDC)* during the MICCAI 2017 conference. It contains cardiac cine-MRI short-axis data from 150 subjects that were divided into 5 subgroups (4 pathological, 1 healthy). For each subject systolic (see Figure 3.5a) and diastolic frames (see Figure 3.5b) are provided with corresponding segmentations (see Figure 3.5c and 3.5d), which enables direct comparison via e.g. the Dice score. The segmentations contain only four values with 0, 1, 2 and 3 representing

pixels located in the background, in the RV cavity, in the myocardium, and in the LV cavity. The frames themselves are 3D volumes with size $216 \times 256 \times 10$, thus ten image slices can be extracted which each have size 216×256 . The same holds true for the segmentation, which means that both the image data and segmentations can be generated in the same manner.

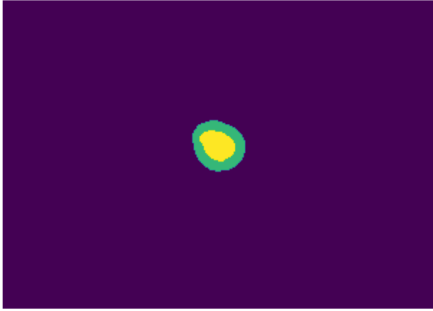
For the training data, the original 4D data was used as we do not require the segmentations for the end-systolic and end-diastolic frames. As the 4D data has size $216 \times 256 \times 10 \times 30$ we can extract 30 frames for each of the ten slices. These can be sorted into 251376 image pairs for training. For the validation and test data the end-systolic and end-diastolic frames with their segmentations were used given us 641 image pairs each as we can only have these two frames to align.



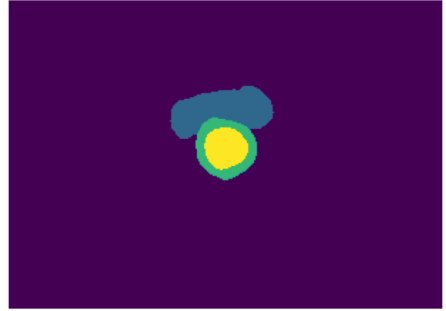
(a) Image of systolic phase.



(b) Image of diastolic phase.



(c) Segmentation of systolic phase.



(d) Segmentation of diastolic phase.

Figure 3.5: Example images for systolic (a) and diastolic frames (b) with corresponding segmentations (c),(d) taken from the ACDC dataset [34].

4

Network Architectures

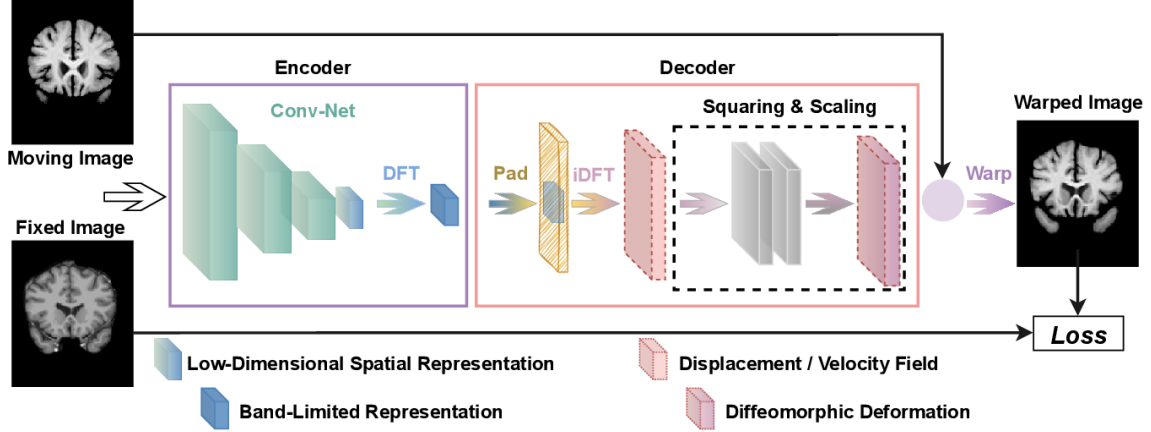
As a starting point *Fourier-Net* [7] and its successor *Fourier-Net+* [8] were used. These networks, which are explained in the following pages, enable fast and accurate registration while needing less resources compared to similar approaches. These attributes are very beneficial for a potential application like motion correction because the current networks, e.g. *LAPNet* [18], are usually supervised and require large computational resources.

4.1 Fourier-Net

Fourier-Net is a new unsupervised approach that aims to learn a low-dimensional representation of the displacement field in a band-limited Fourier domain instead of the full field in the spatial domain. This band-limited representation is then decoded by a model-driven decoder to the dense, full-resolution displacement field in the spatial domain. This allows for fewer parameters and computational operations, resulting in faster inference speeds [7]. The architecture is based on the U-Net [16], like most deep registration approaches, but replaces the expanding path with a parameter-free model-driven decoder as mentioned before. The encoder of *Fourier-Net* consists of a CNN, which takes two images (fixed and moving) as inputs. The output is a displacement field that is then converted from the spatial domain into the Fourier domain via an discrete Fourier transformation (DFT). From there, this band-limiting representation is padded with zeros to the full resolution of the original displacement field. The field is then recovered by using the inverse DFT (iDFT) to convert it back into the spatial domain. This displacement field is then used to warp the moving image into the fixed image. Additionally, squaring and scaling layers [35] can be added before warping the image in order to encourage a diffeomorphism in final deformation.

Encoder

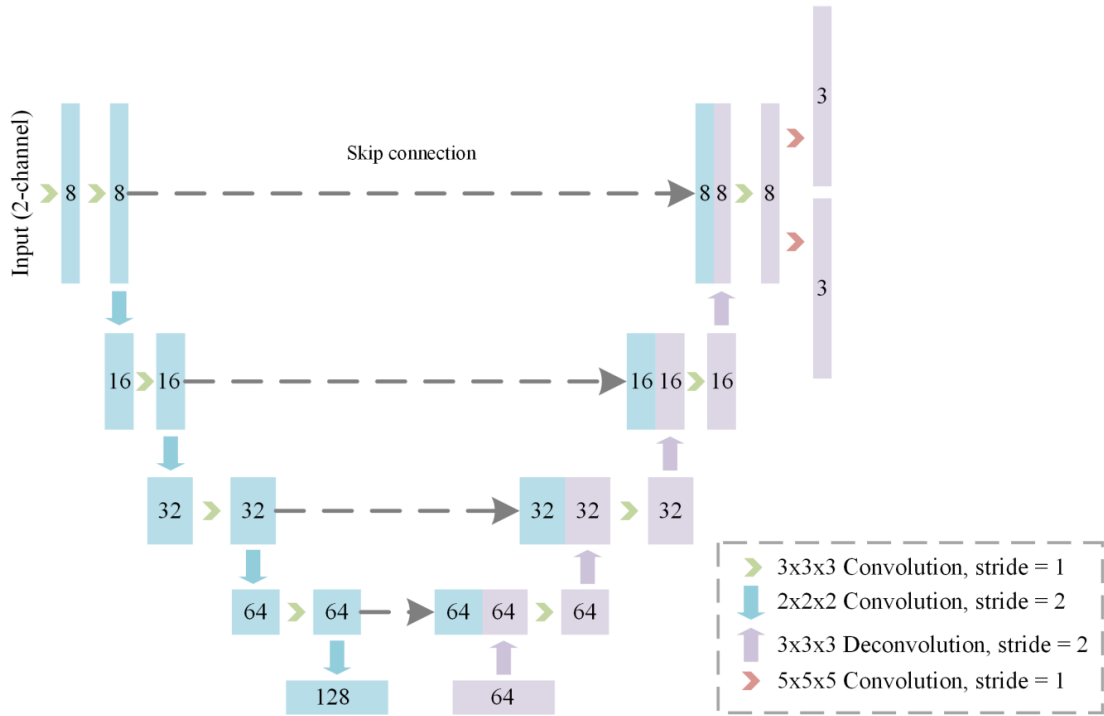
The encoder of *Fourier-Net* consists of a CNN that generates the displacement field between the two inputs followed by a DFT layer that produces a band-limited representation of the full displacement field. The fully convolutional neural network (FCN) from *SYMNet* [21] was modified to function as the CNN of the encoder. Its (original) architec-

Figure 4.1: Architecture of *Fourier-Net* taken from [7].

ture (see Figure 4.2) is again based on the *U-Net* with a contracting and expanding path. The FCN concatenates the inputs images X and Y as a single 2-channel input and estimates two dense, non-linear displacement fields ϕ_{XY} and ϕ_{YX} , however we only need the displacement field for the moving image, denoted as \mathbb{S}_ϕ , since we are not interested in transforming the fixed image. This is actually a low dimensional representation because *Fourier-Net* does not utilize the last two levels of the FCNs expanding path that would be needed to reconstruct the actual full-resolution displacement field ϕ .

For each level in the contracting path of the FCN, two successive convolution layers are applied, which contain one $3 \times 3 \times 3$ convolution layer with a stride of 1, followed by a $3 \times 3 \times 3$ convolution layer with a stride of 2 to further compute the high-level features between the input images as well as downsample the features by half until the lowest level of the network is reached. For each level in the expanding path of the FCN, the feature maps from the contracting path are concatenated through skip connections and apply $3 \times 3 \times 3$ convolution with a stride of 1 and $2 \times 2 \times 2$ deconvolution layer for upsampling the feature maps to twice of its size. At the end of the expanding path, two $5 \times 5 \times 5$ convolution layers with a stride of 1 are appended to the last convolution layer and generate the displacement fields θ_{XY} and θ_{YX} [21]. Each convolution layer in the FCN is followed by a rectified linear unit (ReLU) activation, except for the output convolution layer that does not have an activation function because *Fourier-Net* only uses the first two levels of the expanding path, thus leading the FCN to generate a low dimensional representation \mathbb{S}_ϕ of the full-resolution displacement field ϕ .

As discussed previously, the encoder aims to learn a displacement (or velocity) field in the band-limited Fourier domain. Intuitively, this may require convolutions to be able to handle complex-valued numbers, which can be done by using complex-valued CNNs [36], which are suitable when both input and output are complex values, however these complex-valued operations sacrifice computational efficiency. Other approaches like *DeepFlash* [37] tackle this problem by converting the input images to the Fourier domain and using two individual real-valued CNNs to learn the real and imaginary parts separately. This, however, increases training and inference cost. To bridge the domain gap between real-valued spatial images and complex-valued band-limited displacement fields without increasing



does not need to learn the coefficients of \mathbb{B}_ϕ , but instead only the real-valued coefficients in \mathbb{S}_ϕ , which is the low dimensional spatial representation of the full-resolution spatial displacement field ϕ , which is then reconstructed by the decoder.

Decoder

The decoder contains no learnable parameters, instead the usual expansive path is replaced with a zero-padding layer, an iDFT layer, and an optional squaring and scaling layer.

The output from the encoder is a band-limited representation \mathbb{B}_ϕ in the frequency domain of the low dimensional displacement field \mathbb{S}_ϕ in the spatial domain. To recover the full-resolution displacement field ϕ in the spatial domain, we first pad the patch \mathbb{B}_ϕ , containing mostly low frequency signals, to the original image resolution with zeros. We then feed the zero-padded complex-valued coefficients, denoted as $\mathcal{F}(\phi)$, to an iDFT layer consisting of two steps: shifting the Fourier coefficients from centers to corners and then applying the iDFT to convert them into the spatial domain:

$$\phi_{n,m} = \frac{1}{HW} \sum_{k=0}^{H-1} \sum_{l=0}^{W-1} \mathcal{D}_{k,l} [\mathcal{F}(\phi)]_{k,l} \cdot \exp \left(i \cdot \left(\frac{2\pi n}{H} \cdot k + \frac{2\pi m}{W} \cdot l \right) \right). \quad (4.4)$$

The $H \times W$ sized sampling mask \mathcal{D} is a low-pass filter that has zeros as entries if they are on the positions of high-frequency signals in ϕ and ones if they are on the low-frequency positions. Thus we can reconstruct the full spatial displacement field ϕ from \mathbb{B}_ϕ despite the latter being band-limited. Approaching the problem from the other side we can also think about working backwards from the final displacement. For this, after applying equation (4.1), the low-frequency signals are shifted to a center patch with size $\frac{H}{a} \times \frac{W}{b}$ with $a = 2 \cdot Z_a, b = 2 \cdot Z_b, Z_a, Z_b \in \mathbb{Z}^+$, which is then center-cropped to get \mathbb{B}_ϕ . This crop of $\mathcal{F}(\phi)$ can be reconstructed using the iDFT from equation (4.4) with the cropping functioning as a kind of low-pass filtering:

$$[\mathbb{S}_\phi]_{n,m} = \frac{ab}{HW} \sum_{k=1}^{\frac{H}{a}-1} \sum_{l=1}^{\frac{W}{b}-1} [\mathbb{B}_\phi]_{k,l} \cdot \exp \left(i \cdot \left(\frac{2\pi an}{H} \cdot k + \frac{2\pi bm}{W} \cdot l \right) \right), \quad (4.5)$$

with $n \in [0, \frac{H}{a} - 1]$ and $m \in [0, \frac{W}{b} - 1]$ being the indices of the spatial domain, while $k \in [0, \frac{H}{a} - 1]$ and $l \in [0, \frac{W}{b} - 1]$ are the indices of the frequency domain with i being the imaginary unit. Thus \mathbb{S}_ϕ actually contains all the necessary information from ϕ , as long as they have the same low-frequency coefficients \mathbb{B}_ϕ . This can be formulated as:

$$[\mathbb{S}_\phi]_{n,m} = ab \cdot \phi_{an,bm}, \quad (4.6)$$

because most entries of $\mathcal{F}(\phi)$ are zeros, and the remaining values are exactly the same as in \mathbb{B}_ϕ , which means that \mathbb{S}_ϕ contains all the information ϕ can provide. For this, however, \mathbb{B}_ϕ needs to be padded to the original image size first in order to get the full-resolution displacement and not a low dimensional representation. This ultimately shows that there is a unique mapping between \mathbb{S}_ϕ and ϕ , which means that it is reasonable to use a network to learn \mathbb{S}_ϕ directly from image pairs and then reconstruct the displacement field

in a very efficient manner [7]. The complete reconstruction process is visualized in Figure 4.3.

As both padding and iDFT layers are differentiable, *Fourier-Net* can be optimized via back-propagation. For *Diff-Fourier-Net* extra squaring and squaring layers [35] are needed in the decoder turning the displacement field into a stationary velocity field. Typically seven scaling and squaring layers are used to impose such a diffeomorphism [7, 35].

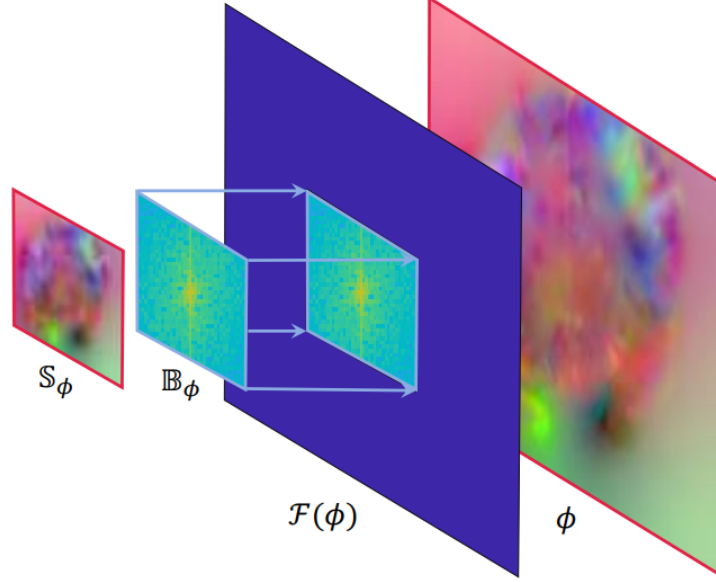


Figure 4.3: Reconstruction of the displacement field via the decoder taken from [8].

Diffeomorphic Transforms

Diffeomorphic deformations are differentiable and invertible, thus preserving topology, which is a desirable property for transformations. $\phi : \mathbb{R}^N \rightarrow \mathbb{R}^N$ represents the deformation that maps the coordinates from one image to coordinates in another image, as long as both images have the dimension N . When using a stationary velocity field representation like e.g. *DARTEL* [38], the deformation field is defined through the following ordinary differential equation (ODE) [35]:

$$\frac{\partial \phi^{(t)}}{\partial t} = v(\phi^{(t)}), \quad (4.7)$$

where $\phi^{(0)} = \text{Id}$ is the identity transformation and t is time. The final registration field $\phi^{(1)}$ can be obtained by integrating the stationary velocity field v over $t = [0, 1]$. This is typically done by integration numerically using scaling and squaring [39]. The integration of a stationary ODE represents a one-parameter subgroup of diffeomorphisms. In group theory, v is a member of the Lie algebra and is exponentiated to produce $\exp(v) = \phi^{(1)}$, which is also a member of the Lie group. From the properties of one-parameter subgroups, for any scalars t and t' , $\exp((t + t') \cdot v) = \exp(t \cdot v) \circ \exp(t' \cdot v)$, where \circ is a composition map associated with the Lie group. Starting from $\phi^{(1/2^T)} = p + v(p)$ where

p is a map of spatial locations, we use the recurrence $\phi^{(1/2)^{t-1}} = \phi^{(1/2^t)} \circ \phi^{(1/2^t)}$ to obtain $\phi^1 = \phi^{(1/2)} \circ \phi^{(1/2)}$. T is chosen so that $v \approx 0$ [35].

As diffeomorphic deformations are defined as smooth and invertible deformations, the output of the iDFT layer in *Fourier-Net* can be regarded as a stationary velocity field v instead of a displacement field ϕ . In Figure 4.1 scaling and squaring layers are visualized. These apply the diffeomorphic transformation in three steps [39]:

1. Scaling: Divide the velocity field v by a factor 2^N , so that $\frac{v}{2^N}$ is close to zero (depending on the desired accuracy).
2. Exponentiation: Compute $\exp(\frac{v}{2^N}) = \phi^{(1)}(\frac{v}{2^N})$ with a first-order explicit numerical scheme.
3. Squaring: N recursive squarings of $\phi^{(1)}(\frac{1}{2^N}) = \exp(\frac{v}{2^N})$ to yield an accurate estimation of $\phi^{(1)}(1) = \phi^{(1)}(\frac{1}{2^N})^{2^N} = \exp(\frac{v}{2^N})^{2^N} = \exp(v)$.

Thus, the diffeomorphic deformation can be efficiently calculated. When used, the specific version is then called *Fourier-Net Diff* to differentiate it from the baseline version.

Spatial Transformer

The warping layer of *Fourier-Net* utilizes the *Spatial Transformer* [40], which allows for spatial image manipulation within the network. This is a differentiable and learnable module for neural networks which applies a spatial transformation to a feature map during a single forward pass. The spatial transformer mechanism is split into three parts as seen in Figure 4.4. First is the localization network, which takes the input and outputs the parameters for the transformation. These are then used to create a sample grid using the grid generator. Lastly, the sampler produces the output feature map based on the input at the grid points.

From the input feature map $U \in \mathbb{R}^{H \times W \times C}$ with width W , height H and channels C the localization network f_{loc} computes the parameters $\theta = f_{\text{loc}}(U)$ of the transformation \mathcal{T}_θ which is later applied to the feature map. Thus the size of θ varies depending on the transformation. The localization network function can both be implemented as a fully-connected network or as a CNN, but should include a final regression layer to produce the transformation parameters.

In order to warp the input feature map, each output pixel is computed by applying a sampling kernel centered at a particular location in the input feature map. The output pixels are defined to lie on a regular grid $G = G_i$ of pixels, forming an output feature map $V \in \mathbb{R}^{H' \times W' \times C}$, where H' and W' are the height and width of the grid with C again being the number of channels, which is the same for input and output.

In order to perform a spatial transformation of the input feature map U , the sampler must take the set of sampling points $\mathcal{T}_\theta(G)$, along and produce the sampled output feature map V . Each coordinate (x_i^s, y_i^s) in $\mathcal{T}_\theta(G)$ defines the spatial location in the input where a sampling kernel is applied to get the value at a particular pixel in the output:

$$V_i^c = \sum_n^H \sum_m^W U_{nm}^c k(x_i^s - m; \Phi_x) k(y_i^s - n; \Phi_y), \quad (4.8)$$

with Φ_x and Φ_y being the parameters for a generic sampling kernel k that defines the image interpolation, U_{nm}^c is the value of the input feature maps at location (n, m) in the channel $c \in [1, \dots, C]$ and V_i^c is the value for every pixel $i \in [1, \dots, H'W']$ for the output feature map. Any sampling kernel can be used, as long as (sub-) gradients can be defined with respect to (x_i^s, y_i^s) to allow the loss gradients to flow back not only to the input feature map, but also to the sampling grid coordinates and therefore back to the transformation parameters θ and localization network, thus enabling back-propagation [40].

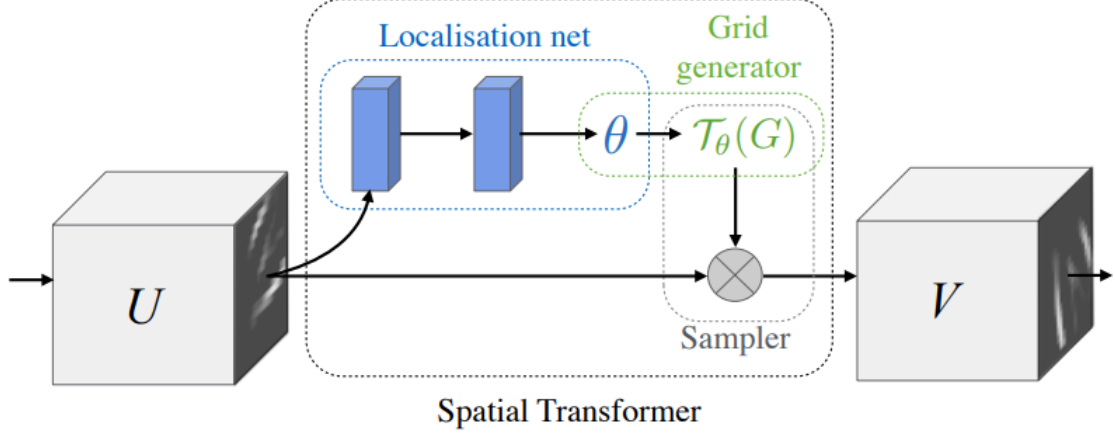


Figure 4.4: Architecture of the *Spatial Transformer* taken from [40].

Loss Function

The loss function consists of two parts to enable unsupervised learning, which are balanced using the scalar parameter λ . The first, \mathcal{L}_1 , measures the similarity between the fixed image and the moving image after warping, while the second, \mathcal{L}_2 , ensures a smooth displacement field. Thus, the unsupervised loss \mathcal{L} can be calculated as follows:

$$\begin{aligned} \mathcal{L}(\Theta) &= \min \left(\mathcal{L}_1(\phi(\Theta)) + \lambda \cdot \mathcal{L}_2(\phi(\Theta)) \right) \\ &= \min \left(\mathcal{L}_1(v(\Theta)) + \lambda \cdot \mathcal{L}_2(v(\Theta)) \right), \end{aligned} \quad (4.9)$$

for both displacement fields ϕ and velocity fields v . The first part of the loss function consists of:

$$\mathcal{L}_1(\phi(\Theta)) = \frac{1}{N} \sum_{i=1}^N \mathcal{L}_{Sim}(M_i \circ (\phi_i(\Theta) + \text{Id}) - F_i), \quad (4.10)$$

where \circ denotes the warping operation, N the number of training pairs with moving images M_i and fixed images F_i , Θ the network parameters, ϕ_i the displacement field, Id the identity grid. \mathcal{L}_{Sim} determines the similarity between warped moving images and fixed images via MSE or NCC, and the second term of the unsupervised loss, \mathcal{L}_2 , defines the

smoothness regularization function that controls smoothness of the displacement fields:

$$\mathcal{L}_2(\phi(\Theta)) = \frac{1}{N} \sum_{i=1}^N \|\nabla \phi_i(\Theta)\|_2^2, \quad (4.11)$$

with ∇ denoting the first order gradient and $\|\cdot\|_2^2$ denoting the squared L_2 -Norm. When using the squaring and scaling layers, thus making the deformation of the moving image diffeomorphic, the loss needs to be modified by replacing the displacement field θ with the velocity field v . Thus, both parts of the of the loss function need to be changed:

$$\mathcal{L}_1(v(\Theta)) = \frac{1}{N} \sum_{i=1}^N \mathcal{L}_{Sim}(M_i \circ \text{Exp}(v_i(\Theta)) - F_i), \quad (4.12)$$

$$\mathcal{L}_2(v(\Theta)) = \frac{1}{N} \sum_{i=1}^N \|\nabla v_i(\Theta)\|_2^2 \quad (4.13)$$

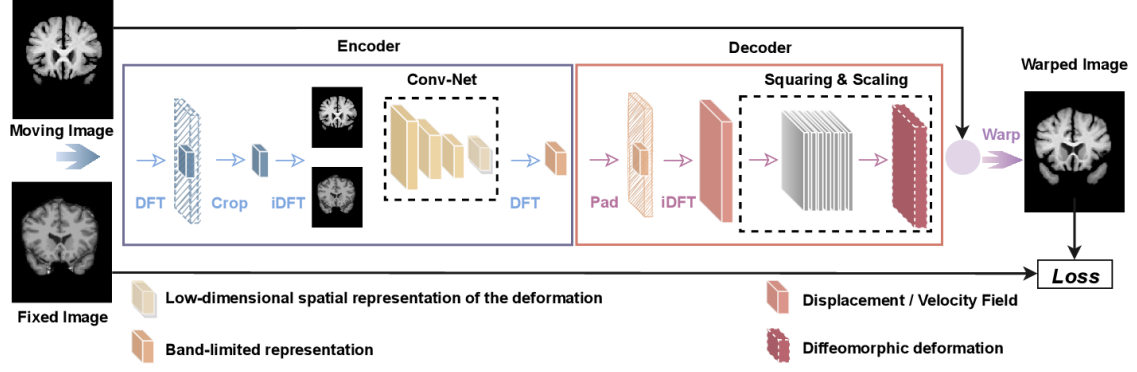
4.2 Fourier Net+

Fourier-Net+, as the name suggests, is an extension of *Fourier-Net* which takes the band-limited spatial representation of the images as input, instead of their original full-resolution counterparts. This leads to further reduction in the number of convolutional layers in the contracting path of the network, resulting in a decrease of parameters, memory usage, and computational operations. This makes *Fourier-Net+* even more efficient than its predecessor [8].

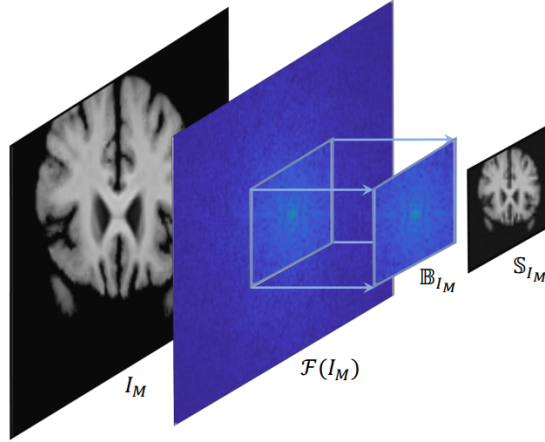
As seen in Figure 4.5, the network architecture is almost the same as for *Fourier-Net* (see Figure 4.1 for comparison). However, while the decoder, and thus the loss function, remain the same, the encoder is slightly altered to make the network even more efficient. For this, similarly to the decoder, a DFT is used, however this time the idea is applied to the input images. These are first transformed into the Fourier domain, then low-pass filtered by center-cropping and finally reconstructed from their band-limited representation back into the spatial domain via an iDFT. The two images, now compressed, are the input for the encoder of *Fourier-Net*, meaning the CNN and following DFT. However, due to the band-limiting before the CNN, the latter can be made much more light-weight, thus reducing computational cost. This is visualized in Figure 4.7. Thus, *Fourier-Net+* too is overall lighter than the baseline *Fourier-Net* in terms of the number of parameters and computations. However, such a light network may face limitations in accurately capturing complex deformations. To counter this potential weakness, the authors propose a cascaded version of *Fourier-Net+*, which uses multiple versions of *Fourier-Net+* cascaded one after the other to achieve a better overall displacement field [8]. A schematic for this can be seen in Figure 4.8.

Changes to the Encoder

In order to further reduce the amount of computational operations, *Fourier-Net+* discards the early layers of the encoder. Instead, a DFT $\mathcal{F}(I_M)$ followed by a center-crop to produce

Figure 4.5: Architecture of *Fourier-Net+* taken from [8].

the band-limited representation \mathbb{B}_{I_M} in the frequency domain and iDFT are used to get the spatial patch \mathbb{S}_{I_M} , while the rest of the encoder from *Fourier-Net* stays the same. The input I_M is thus compressed to a lower resolution (i.e. band-limited) using the frequency space, which reduces the computational cost. This process is visualized in Figure 4.6. The encoder of *Fourier-Net+* has several convolutional layers less in the contracting path compared to *Fourier-Net*, which leads to a further accelerated registration process while reducing the memory footprint. These advantages are visualized in Figure 4.7 where the amount of different layers between a conventional *U-Net*, *Fourier-Net* (with the smaller decoder) and *Fourier-Net+* (with a smaller encoder and decoder) are shown.

Figure 4.6: Compression in the frequency domain of the encoder used in *Fourier-Net+* taken from [8].

Effects of Cascading

As seen in the previous section, *Fourier-Net+* is lighter than *Fourier-Net* due to the band-limited representation of both images and deformations lowering the number of parameters and computations. This, however, can lead to limitations when trying to accurately capture complex deformations. To this end, a cascaded version of *Fourier-Net+*

4 Network Architectures

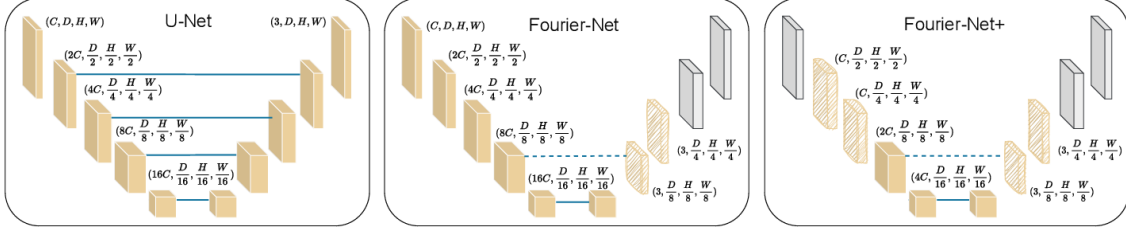


Figure 4.7: Architecture of the CNN for a typical U-Net, *Fourier-Net* and *Fourier-Net+* taken from [8].

called $K \times \text{Fourier-Net+}$ (see Figure 4.8), with K denoting the amount of cascades, can be used where the warped image of one cascade is used as the moving image of the next. It is important to note that the weights are not shared between cascades. The squaring and scaling layers, in case a diffeomorphic deformation is wanted, are applied after the last cascade. The same is true for the calculation of the loss function.

In order to more accurately describe this version of *Fourier-Net+* one can look at the in- and outputs of the different cascades. The first cascade of $K \times \text{Fourier-Net+}$ has the moving image I_M and fixed image I_F as inputs. While the latter always stays the same for all cascades the warped image $I_M^{w(1)}$ from the first cascade is used as input for the second cascade instead of the original moving image. Thus, in general $I_M^{w(k-1)}$ and I_F are the inputs for a cascade $k \in [1, K]$ with output $\delta\phi^{(k)}$. Furthermore, $I_M^{w(k)}$ can be defined as:

$$I_M^{w(k)} = (((I_M \circ \delta\phi^{(1)}) \circ \delta\phi^{(2)}) \circ \dots) \circ \delta\phi^{(k-1)} \circ \delta\phi^{(k)} = I_M \circ \phi^{(k)}, \quad (4.14)$$

with $\phi^{(k)}$ being the displacement field computed by composing the outputs of the first cascade up to the k -th cascade:

$$\phi^{(k)} = \delta\phi^{(1)} \circ \delta\phi^{(2)} \circ \dots \circ \delta\phi^{(k-1)} \circ \delta\phi^{(k)}. \quad (4.15)$$

Thus, the output displacement field of the K -th cascade $\phi^{(K)}$ is the final displacement which is then used to warp I_M , the original moving image, in order to compute the loss [8].

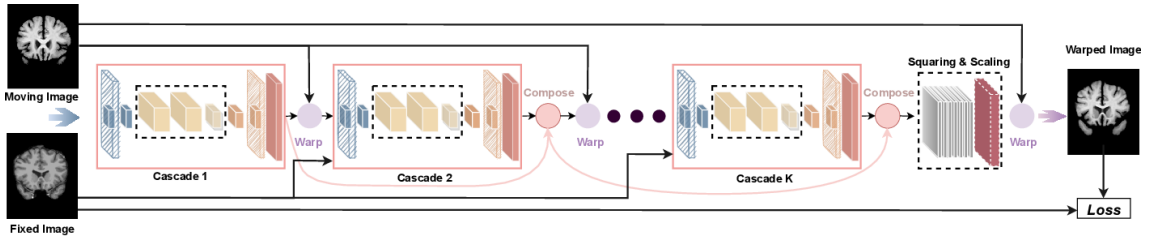


Figure 4.8: Cascaded version of *Fourier-Net+* taken from [8].

5

Experiments

In the following chapter the different experiments that were conducted are described and explained. First, experiments were conducted on the *CMRxRecon* dataset to find the optimal parameters for our model. As this dataset lacks ground truth segmentations, the *ACDC* dataset was used to extend these tests.

5.1 Parameter Studies on the *CMRxRecon* Dataset

As a first step, frame-to-frame registration on the *CMRxRecon* dataset was tested. The goal of these initial experiments was to examine whether the network could learn to align the cardiac image pairs despite the movement between the frames. To start, different parameters were tested in order to find the optimal model parameters for the *CMRxRecon* dataset.

As discussed in section 3.2, the CMR images were cropped to extract the moving cardiac region. This was done for both the fully sampled and subsampled (Acc4, Acc8, Acc10) data. Image pairs for each slice from each patient of the training set were generated to train the network. In order to evaluate the training progress the mean MSE and mean SSIM metrics calculated the validation data at each checkpoint (usually every 1000 iterations). The best model parameters, as determined by the metrics, were saved in a separate folder. To further evaluate the training success the unseen test data containing new patients was used. To benchmark the networks training success the MSE and SSIM were again used, but the mean inference time and the percentage of non-positive Jacobian determinant of deformation ($\% |J_\phi| \leq 0$) were also calculated. The procedure, however, was slightly altered for the subsampled data as only the displacement field of the model was used to align the corresponding fully sampled image pairs. The evaluation metrics were then calculated on these images for better comparability.

Fourier-Net versus Fourier-Net+

First, the baseline network *Fourier-Net* was compared to its newer version *4xFourier-Net+* once with both using the diffeomorphic version and once without the transform. Thus the effect of the diffeomorphic deformation can be compared to the normal transforma-

tion on *Fourier-Net* and *4xFourier-Net+*. All of the experiments were conducted on the fully sampled data as the subsampling should not effect the tested properties. The other parameters were also kept constant with a learning rate of 0.0001, MSE-loss and $\lambda = 0.01$. FT crop in *4xFourier-Net+* was set to 24×24 .

Starting Channel Size

Next, the impact of the starting channels was examined. For this, *4xFourier-Net+* was trained with MSE-loss, a learning rate of 0.0001, $\lambda = 0.01$, FT crop of 24×24 and no diffeomorphic transform. The experiments were again only conducted on the fully sampled data. Three different channel sizes were used for training *Fourier-Net+*: 8, 16 and 32. Due to the early stopping the networks were trained for 17, 17 and 10 epochs respectively.

Quantitative Comparison with NiftyReg

To establish a baseline, the similarity metrics were calculated between the unaligned test data. *4xFourier-Net+* was then trained with the MSE-loss, 8 channels, $\lambda = 0.01$ and a learning rate of 0.0001. *NiftyReg*, a traditional registration algorithm that iteratively optimizes the displacement [22], was used for comparison. *NiftyReg* provides the warped image for the fully sampled data directly, but for the subsampled data the control point grid produced by *NiftyReg* was used to resample the corresponding fully sampled data for comparability. Thus only the fully sampled test data is needed for an unregistered baseline as the subsampled data is only used to produce the displacement which is then used to warp the fully sampled data.

5.2 Parameter Tests on the ACDC Dataset

As the evaluation on the *CMRxRecon* dataset is limited to image similarity measures, further testing was done on the *ACDC* dataset which contains cardiac data with segmentations. These can be used to better evaluate the registration performance by using e.g. the Dice score.

Fourier-Net versus Fourier-Net+

First, *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net* are compared for both normal and diffeomorphic versions. Additionally, network versions creating dense instead of band-limited displacements were used as a baseline for best achievable performance. The registration performance was evaluated on the test set using the percentage of Dice scores calculated with and without the background label, percentage of SSIM, MSE multiplied by 10^{-3} (denoted by m for milli) and the percentage of negative Jacobian determinant of deformation. The mean inference time on GPU together with memory consumption for each model is also added to the comparison with the latter containing number of trainable model parameters, mult-add operations (in millions or billion as denoted by M or G) and the total memory in Megabyte (MB). The latter is only given for the normal versions as the

diffeomorphic version require the same amount of memory and the baseline dense versions. The first network was trained for a maximum of 100 epochs with early stopping activating after 3 epochs without improvement of the validation Dice score. The other variants were then trained with the same number of epochs, which worked out to be only 6 epochs due to the large training set in the experiments. All networks were trained on the fully sampled ACDC data with MSE as the unsupervised similarity loss, a channel size of 8, learning rate of 0.0001 and $\lambda = 0.01$. *Fourier-Net+* and *4xFourier-Net* used a FT crop size of 48×48 for these experiments. All of these parameters were the same for the diffeomorphic and dense versions of the networks. The results and subsequent discussion can be found in section 6.2.

Starting Channel Size

Next, the impact of the starting channels on the ACDC data was examined. For this, *Fourier-Net+* and *4xFourier-Net+* was trained with MSE-loss, a learning rate of 0.0001, $\lambda = 0.01$, FT crop of 24×24 and no diffeomorphic transform. The experiments were again only conducted on the fully sampled data. Four different channel sizes were used for training: 8, 16, 32 and 64. The first network variant was trained with early stopping, which ended at 6 epochs. For better comparability, the other network variants were also trained with the same number of epochs. The registration performance was evaluated on the test set using the percentage of the Dice score calculated once on all labels, once without the background using only the cardiac labels, as well as the percentage of SSIM, MSE multiplied by 10^{-3} and the percentage of negative Jacobian determinant of deformation. Additionally the number of network parameters, number of mult-add operations (in millions or billion as denoted by M or G) and memory consumption in Megabyte (MB) are listed for all network variants to better compare the relationship between registration performance and memory consumption to potentially find an optimal configuration for both performance and efficiency. The mean inference time was measured on GPU in seconds. The results and subsequent discussion can be found in section 6.2.

Fourier-Transform Crop Size

The FT crop size used for compressing the images is a parameter specific to *Fourier-Net+* and *4xFourier-Net+*. Four different sizes of the FT crop were analyzed: 80×168 , 40×84 , 48×48 and 24×24 . A larger crop would obviously compress the images less, thus leading to less efficiency, however, a very small crop, while very efficient, might lose some of the image details leading to a decrease in performance. The experiments were again only conducted on the fully sampled data as one would expect similar results on subsampled data. The network variants were trained with MSE as the unsupervised similarity loss, a channel size of 8, learning rate of 0.0001 and $\lambda = 0.01$. Again, the networks were trained for only 5 epochs, as this is where the first model ended training due to early stopping. The registration performance was evaluated on the test set using the Dice score calculated once on all labels, once without the background, as well as the SSIM, MSE and the percentage of negative Jacobian determinant of deformation. Additionally, the number of network parameters, number of mult-add operations in millions (M) and memory

consumption in Megabyte (MB) are listed for all network variants to better compare the relationship between registration performance and memory consumption to potentially find a optimal configuration for both performance and efficiency. The mean inference time was measured on GPU in seconds. The results and subsequent discussion can be found in section 6.2.

Comparison with VoxelMorph

After finding parameters which optimize registration performance while minimizing memory consumption, a comparison to another commonly used registration network should be made. For this, the famous *VoxelMorph* [6] was used. It was trained with the default number of layers, MSE as the similarity loss, a learning rate of 0.0001, $\lambda = 0.01$ and provides a dense displacement field. For comparison *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* were trained for 6 epochs with MSE as the unsupervised similarity loss, a channel size of 16, learning rate of 0.0001 and $\lambda = 0.01$ as well as a FT crop size of 48×48 , which provide a good balance of performance and memory imprint for the networks. To evaluate registration performance, the Dice score was again computed for all labels and without the background as well as the similarity metrics SSIM, MSE and the percentage of negative Jacobian determinant of deformation. All networks were trained on the fully sampled ACDC data. The inference times of all networks were computed on the GPU and the number of network parameters, number of mult-add operations in billions (G) and memory consumption in Megabyte (MB) are also given to further compare the efficiency of the networks.

Dense Displacement on Accelerated Data

The difference between a dense displacement field and a band-limited one was already explored in section 5.2. This, however, only includes fully sampled data, not accelerated data. We will explore potential performance changes on the latter in this section with the hypothesis that the network variants with the dense displacement will perform worse than the variants with the band-limited displacement field. This expectation stems from the fact, that frequencies outside the center region of the k-space are dropped for the accelerated data thus reducing the advantage of the dense displacement in comparison to the band-limited (i.e. center-cropped) displacement.

The tests again included *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+*, which were all trained for 6 epochs with MSE as the unsupervised similarity loss, a channel size of 16, learning rate of 0.0001 and $\lambda = 0.01$. The registration performance was evaluated on the test set using the Dice score calculated once on all labels, once without the background, as well as the SSIM, MSE and the percentage of negative Jacobian determinant of deformation. Additionally, the number of network parameters, number of mult-add operations in millions (M) and memory consumption in Megabyte (MB) are given.

Comparison on Subsampled Data

After comparing *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* with both dense and band-limited displacement fields as well as a comparison with *VoxelMorph*, which also has a

dense displacement field, on fully sampled data to access the difference in registration performance and memory consumption an extension to subsampled data needs to be made. For this *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* were compared to a baseline consisting of the unaligned test image pairs as well as *NiftyReg*, *VoxelMorph* and *LAPNet*. Thus, observations of the performance on accelerated data can be made as well as comparisons to the aligned baseline (which all methods should hopefully beat), a traditional registration algorithm, a dense registration network working in the image domain and another dense network which works in the k-space domain.

Fourier-Net, *Fourier-Net+* and *4xFourier-Net+* were trained for 6 epochs with MSE as the unsupervised similarity loss, a channel size of 16, learning rate of 0.0001 and $\lambda = 0.01$ as well as a FT crop size of 48×48 for all subsampling data. *VoxelMorph* was trained with default layers and MSE as the similarity loss, a learning rate of 0.0001 and $\lambda = 0.01$. Three different accelerations were tested: *Acc4* (R=4), *Acc8* (R=8) and *Acc10* (R=10). For reference, the data for the fully sampled data (R=0) is also provided. The registration performance of all methods was evaluated on the test set using the Dice score calculated once on all labels, once without the background, as well as the SSIM and MSE as image metrics. All times are computed on CPU for better comparability as *NiftyReg* currently only works on CPU as the GPU capable versions were deprecated and the other networks would gain an unfair advantage when ran on GPU. The baseline obviously has no time associated as it only denotes the metrics on the test data before registration.

5.3 Data Consistency Experiments

As seen in the previous sections, the performance for all methods decreases drastically for highly subsampled/accelerated data due to reconstruction artifacts in the image domain. However, since we are interested in maintaining performance on subsampled data, it might prove useful to use k-space information for the loss calculation and thus for the network training as it should not contain artifacts compared to the image domain. Thus, it would act as a kind of data consistency constraint during training. In order to achieve this, the similarity loss was calculated not on the warped and fixed images, but instead on their corresponding k-spaces. This would hopefully still provide the network with useful information during training that is free of any aliasing artifacts, which might decrease performance on subsampled data.

After trying the MSE as the similarity loss, it quickly became apparent that this yielded unrealistic and wildly deformed displacements which could no longer be reigned in by the smoothness constraint of the loss function. Thus, normalized cross-correlation (NCC) was used for the calculation of the similarity loss with $\lambda = 1$ to place a greater emphasize on the smoothness of the displacement.

5.4 Integration into a Motion-Corrected Reconstruction Pipeline

Describe qualitative test to see whether the network can be used to improve a reconstruction pipeline for cardiac data.

General Reconstruction Pipeline

Overview of the reconstruction pipeline using *Fourier-Net+*.

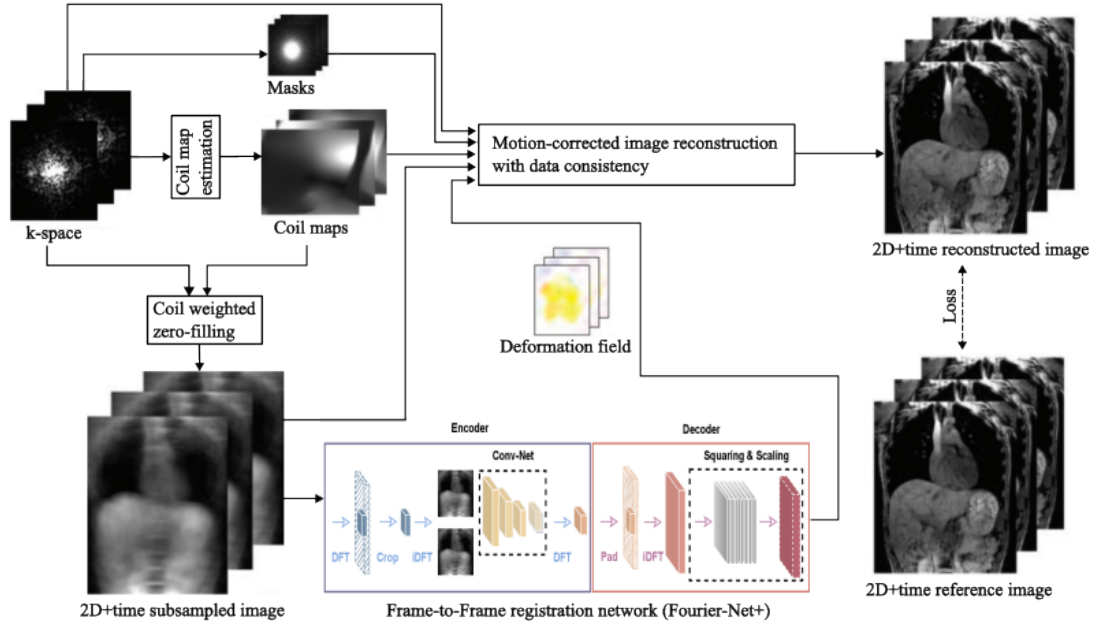


Figure 5.1: Overview of the general reconstruction pipeline using *Fourier-Net+* as a registration network.

6

Results and Discussion

In this chapter, the results for the experiment described in the previous chapter will be examined. Possible implications will be discussed to determine the effectiveness of this new method.

6.1 Parameter Studies on the CMRxRecon Dataset

Different parameters were tested in order to find the optimal model parameters for usage on the *CMRxRecon* dataset. After some tests on the fully sampled data, a comparison with the traditional iterative algorithm *NiftyReg* was conducted on both the fully sampled and subsampled data.

Fourier-Net versus Fourier-Net+

First, the baseline network *Fourier-Net* was compared to its more efficient successor *4xFourier-Net+* with both using the diffeomorphic version as we can then compare the effect of the diffeomorphic deformation compared to the normal transformation on *4xFourier-Net+*. However, this effect should be similar for *Fourier-Net*. All of the experiments were conducted on the fully sampled data as the subsampling should not effect the tested properties. The baseline MSE (note that all MSE values are multiplied by 10^{-3} as denoted by the m) for unregistered test images is 0.535 and the SSIM percentage (% SSIM) is 91.71. The results for *Fourier-Net* and *4xFourier-Net+* can be seen in Table 6.1.

The *Fourier-Net* model without the diffeomorphic transform was trained for 17 epochs, while the diffeomorphic version trained for 100 epochs. Despite this, the latter is only marginally better in terms of MSE and SSIM. The percentage of negative Jacobian determinants was actually better for the normal transformation compared to the diffeomorphic one. The normal *Fourier-Net+* model trained for 37 epochs, while the diffeomorphic version trained for 100 epochs. *Fourier-Net+* performs slightly worse than *Fourier-Net*, however it is more efficient.

Table 6.1: Results for *Fourier-Net* and *Fourier-Net+* on the fully sampled CMRxRecon test data with a diffeomorphic transform.

Model	MSE (m)	% SSIM	% $ J_\phi \leq 0$	Time [s]
Fourier-Net	0.319 ± 0.352	93.29 ± 3.85	0.0022 ± 0.0183	0.0025
4xFourier-Net+	0.352 ± 0.449	93.08 ± 4.09	0.0003 ± 0.0057	0.0123
Diff-Fourier-Net	0.318 ± 0.388	93.39 ± 3.82	0.0273 ± 0.0897	0.0028
Diff-4xFourier-Net+	0.352 ± 0.448	93.08 ± 4.11	0.0029 ± 0.0214	0.0103

Starting Channel Size

Next, the impact of the starting channels was examined. For this, *4xFourier-Net+* was trained with three different channel sizes: 8, 16, 32 and 64. Due to the early stopping the *4xFourier-Net+* variants were trained for x, 17 and 10 epochs respectively. The results can be seen in Table 6.2 with the inference time for each version being calculated on the GPU. Despite training for the least amount of epochs channel size 32 performs the best for *4xFourier-Net+*.

Table 6.2: Results for three different starting channel sizes of *Fourier-Net+* on the fully sampled CMRxRecon test data.

Starting Channels	MSE (m)	% SSIM	% $ J_\phi \leq 0$	Time [s]
8	0.352 ± 0.449	93.08 ± 4.09	0.0003 ± 0.0057	0.0123
16	0.347 ± 0.439	93.12 ± 4.04	0.0003 ± 0.0059	0.0113
32	0.344 ± 0.429	93.13 ± 4.01	0.0008 ± 0.0098	0.0106

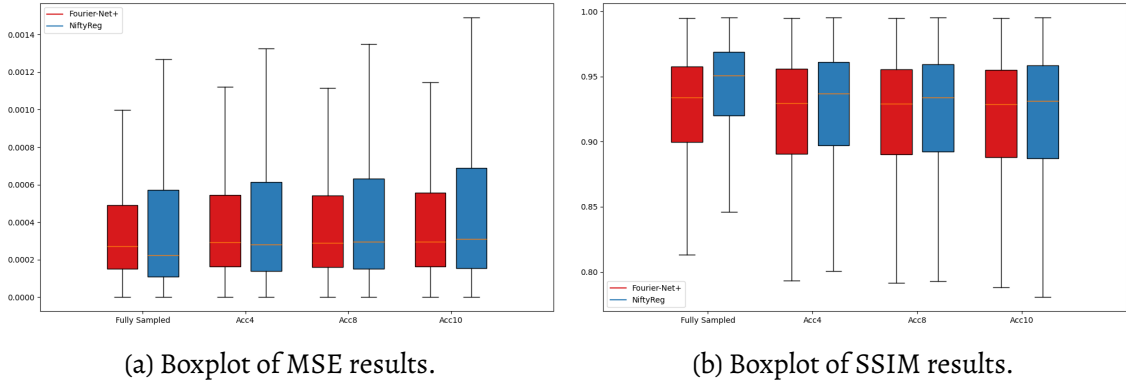
Quantitative Comparison with NiftyReg

For this first quantitative test similarity metrics such as MSE and SSIM were used to judge image similarity as an indicator for proper image alignment. The results can be seen in Table 6.3 with the SSIM given in percent % and the MSE multiplied by 10^{-3} for better readability (this is denoted by m for milli although the MSE has no unit strictly speaking). They are further visualized in Figure 6.4a for the MSE and Figure 6.4b for the SSIM results.

Fourier-Net+ needed about 0.0104 seconds (s) per frame pair for registration with $|J|_{<0} \% < 0.001$ for all levels of subsampling, while *NiftyReg* took up to 15 s for aligning an image pair. The results for *Fourier-Net+* look good on their own as the metrics are stable even for the subsampled data compared to the fully sampled data. However, its baseline performance is slightly worse than *NiftyReg* except for the Acc10 (R=10) data, where both perform worse than the unaligned baseline of the test data, however *4xFourier-Net+* is slightly better than *NiftyReg*. Overall, the network is much faster than the iterative registration algorithm as expected, but the latter performs better in terms of similarity metrics.

Table 6.3: Benchmark results for *Fourier-Net+* and *NiftyReg* on the *CMRxRecon* test data.

Subsampling	Method	MSE (m)	% SSIM
R=0	4xFourier-Net+	0.435 ± 0.554	92.35 ± 4.59
	NiftyReg	0.514 ± 0.771	93.97 ± 3.91
R=4	4xFourier-Net+	0.482 ± 0.626	91.81 ± 5.01
	NiftyReg	0.551 ± 0.798	92.38 ± 4.94
R=8	4xFourier-Net+	0.479 ± 0.620	91.74 ± 5.12
	NiftyReg	0.565 ± 0.803	92.06 ± 5.09
R=10	4xFourier-Net+	0.491 ± 0.640	91.59 ± 5.23
	NiftyReg	0.926 ± 2.867	90.66 ± 10.32

Figure 6.4: Boxplots for MSE and SSIM results of the test set for fully sampled and subsampled (Acc4, Acc8, Acc10) data for both *Fourier-Net+* and *NiftyReg*.

6.2 Further Tests on the ACDC Dataset

Due to the limitations of the *CMRxRecon* dataset in terms of segmentations, the registration performance is further evaluated on the *ACDC* dataset. Some of the parameter studies done on *CMRxRecon* were repeated, others were added to further test the networks ability.

Fourier-Net versus Fourier-Net+

Again, *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net* are compared in terms of registration performance and inference time (on GPU) together with their diffeomorphic versions, however, versions with dense instead of band-limited displacements are also added as a baseline for peak performance as these utilize all of the k-space information. The memory consumption is also added to the comparison with the number of parameters, mult-add operations and the total memory. The latter is only given for the dense and normal versions as the diffeomorphic version required roughly the same amount of memory as the normal ones, while the dense versions have a slightly larger encoder. Note that all net-

work variants were trained for 6 epochs. The results can be seen in Table 6.5.

For the dense displacement variants the diffeomorphic transform increases the performance in terms of Dice score (with and without the background label) slightly for *Fourier-Net* and *4xFourier-Net*, but decreases for *Fourier-Net+*. The SSIM decreases for all models slightly, while the MSE stays about the same. Unsurprisingly, the percentage of non-positive Jacobian determinants goes down for the diffeomorphic variants. The times between the baseline and diffeomorphic versions do not vary a lot with the *Fourier-Net* being faster without the diffeomorphis, while *Fourier-Net+* and *4xFourier-Net* are faster.

The results for the band-limited versions are very similar with *Fourier-Net* and *Fourier-Net+* performing slightly worse in terms of Dice (with background) with the diffeomorphis, while *4xFourier-Net* performs better by almost 2%. When excluding the background label *Fourier-Net* again performs slightly worse, but *Fourier-Net+* and *4xFourier-Net* both improve, the latter with almost 3% more than the baseline version. These changes are not consistent with the dense displacement versions, however, the band-limited *4xFourier-Net* version might be an outlier, as all of the other models are not affected as much by the diffeomorphic transform. The SSIM values slightly decrease for *Fourier-Net* and *Fourier-Net+* with the diffeomorphism, but again improve slightly for *4xFourier-Net*. The MSE is slightly better for *Fourier-Net* and *4xFourier-Net*, but a bit worse for *Fourier-Net+*. The percentage of non-positive Jacobian determinants again decreases for the diffeomorphic variants, which is consistent with the previous observations on the dense displacement versions. The times again vary slightly with *Fourier-Net* and *Fourier-Net+* being a bit slower, while *4xFourier-Net* is quite a bit faster.

Now to the difference between the dense and band-limited displacement. Overall, the dense displacement versions perform better in terms of Dice score for all models both with and without the background label. This is to be expected as the dense displacement versions can use all of the available k-space data for the registration task, while the band-limited versions have only a limited amount of k-space data, directly impacting and limiting their performance. The dense displacement is also better in the SSIM metric, however the difference is not quite as large as with the Dice scores. As the differences between the frames is not quite as large, the weaker registration performance does not impact this metric as much as the Dice score, which is focused on the moving cardiac region. The same is true for the MSE, but the dense displacement is again far superior. Only in terms of the percentage of non-positive Jacobian determinants does the band-limited displacement actually perform better as the band-limiting probably allows for only a smaller deformation. However, the difference is still very small, especially for the diffeomorphic versions, and it could be argued that a better registration performance at the cost of some image folding is an acceptable trade-off. In terms of inference time the dense displacement variants are surprisingly slightly faster despite having a larger encoder, however all of the model are in the low milliseconds (< 40 ms). Last but not least, the memory consumption needs to be addressed. The dense displacement variants have a larger encoder as discussed before and thus have a lot more parameters, especially for the more optimized *Fourier-Net+* and *4xFourier-Net* (about 5 times more). The number of Mult-Adds and the amount of total memory are more than doubled (almost tripled for *4xFourier-Net*) for the dense displacement versions compared to those with a band-limited displacement across all models.

Table 6.5: Results for *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* with both dense and band-limited displacement fields as well as diffeomorphic transforms on the fully sampled ACDC test data.

Metrics	Dense Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	78.22 ± 13.84	78.43 ± 13.96	79.34 ± 14.03
% Dice*	71.03 ± 18.87	70.50 ± 19.17	72.02 ± 18.94
% SSIM	91.92 ± 3.32	89.09 ± 4.10	89.65 ± 3.85
MSE (m)	0.09 ± 0.06	0.17 ± 0.13	0.15 ± 0.12
% $ J_\phi \leq 0$	0.53 ± 0.52	0.32 ± 0.55	0.04 ± 0.09
Time [s]	0.0070	0.0107	0.0219
	Diff-Fourier-Net	Diff-Fourier-Net+	Diff-4xFourier-Net+
% Dice	78.56 ± 14.13	77.88 ± 13.81	79.49 ± 14.26
% Dice*	71.31 ± 19.13	70.19 ± 18.82	72.12 ± 19.33
% SSIM	91.79 ± 3.38	89.08 ± 4.10	89.62 ± 3.98
MSE (m)	0.09 ± 0.06	0.16 ± 0.13	0.15 ± 0.12
% $ J_\phi \leq 0$	0.03 ± 0.06	0.00 ± 0.00	0.00 ± 0.00
Time [s]	0.0082	0.0075	0.0203
Parameters	645,216	380,470	1,521,880
Mult-Adds (G)	1.12	0.04	0.18
Memory [MB]	97.63	5.81	21.90
Metrics	Band-limited Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	77.95 ± 14.71	75.35 ± 14.28	76.59 ± 14.84
% Dice*	69.96 ± 21.55	64.74 ± 21.12	66.82 ± 21.55
% SSIM	91.35 ± 3.51	88.42 ± 3.94	88.59 ± 4.23
MSE (m)	1.00 ± 0.71	2.06 ± 1.54	1.92 ± 1.46
% $ J_\phi \leq 0$	0.36 ± 0.38	0.13 ± 0.35	0.03 ± 0.12
Time [s]	0.0145	0.0145	0.0335
	Diff-Fourier-Net	Diff-Fourier-Net+	Diff-4xFourier-Net+
% Dice	77.93 ± 14.93	75.17 ± 13.43	78.20 ± 14.01
% Dice*	69.91 ± 21.99	65.67 ± 18.61	69.61 ± 19.11
% SSIM	91.24 ± 3.56	87.81 ± 3.94	88.81 ± 4.13
MSE (m)	0.98 ± 0.67	2.29 ± 1.61	1.85 ± 1.41
% $ J_\phi \leq 0$	0.01 ± 0.03	0.00 ± 0.00	0.00 ± 0.00
Time [s]	0.0150	0.0207	0.0188
Parameters	434,519	75,429	301,716
Mult-Adds (M)	595.04	10.98	43.91
Memory [MB]	44.69	2.25	7.66

Starting Channel Size

Next, the impact of the starting channels was examined. For this, *Fourier-Net+* and *4xFourier-Net+* were trained with four different channel sizes: 8, 16, 32 and 64. The memory consumption as well as inference time on the GPU were calculated to extend the evaluation given by the other metrics. All network variants were trained for 6 epochs. The results can be seen in Table 6.6.

For both *Fourier-Net+* and *4xFourier-Net+* there is a clear increase in Dice (with and without the background label) with larger channel sizes as more features increase the registration performance. The SSIM and MSE metrics only increase marginally with larger channel size. The percentage of non-positive Jacobian determinants actually decreases as the registration performance increases, likely due to better/more features enabling a smoother and better displacement field to be generated by the models.

The inference time is not heavily impacted by channel size as the times remain very similar for all sizes. The memory however increases exponentially when doubling the channel sizes as all layers are effected, not just the starting one as the name suggests. Thus the number of parameters drastically increases with starting size leading to an increase in Mult-Add and thus overall memory. As *4xFourier-Net+* is just a cascaded version of *Fourier-Net+* a channel size of 16 for the latter is about the same as channel size 8 for the cascaded version and so on. Thus it seems that there is no point at which both the performance and memory consumption are maximized at the same time. This can also be seen in the direct comparison between *Fourier-Net+* and *4xFourier-Net+*, where the latter has a better performance, but also needs more memory. While an increase of channel size also increases performance in terms of Dice (with the background label) by 0.36%, 0.43% and 0.41% for the latter, which is quite consistent, it can be observed that an increase of channel size for *Fourier-Net+* has a bigger impact at the beginning (1.38%, 0.78% and 0.15%) and fades towards larger channels sizes. This effect can also be seen when excluding the background label (0.69%, 0.78% and 0.42% for *4xFourier-Net+* compared to 2.16%, 1.09% and 0.10% for *Fourier-Net+*), but is less pronounced for the SSIM and MSE as these do not capture the difference in the cardiac regions as well. Note that all versions of *Fourier-Net+* and *4xFourier-Net+* are smaller in terms of total memory than *Fourier-Net* with channel size 8 (44.69 MB), except for *4xFourier-Net+* with channel size 64.

Fourier-Transform Crop Size

As a parameter specific to *Fourier-Net+* and *4xFourier-Net+*, the impact of the FT crop size used for compressing the images was analyzed. Four different sizes of the FT crop were used for training: 80×168 , 40×84 , 48×48 and 24×24 . The experiments were again only conducted on the fully sampled data and each network trained for 5 epochs. The results can be seen in Table 6.7.

The registration performance, as measured by the Dice score both with and without the background label, decreases drastically with a smaller crop size. This is as expected as the smaller FT drop compresses the images more thus making an accurate registration harder. The SSIM and MSE also get worse (SSIM decreases, MSE increases), however the amount of performance lost is hard to gauge, especially for the MSE metric. Interestingly,

Table 6.6: Results for different starting channel sizes of *Fourier-Net+* and *4xFourier-Net+* on the fully sampled ACDC test data.

Metrics	Starting Channels - Fourier-Net+			
	8	16	32	64
% Dice	75.50 ± 13.79	76.88 ± 13.86	77.66 ± 13.60	77.81 ± 13.76
% Dice*	65.70 ± 18.96	67.86 ± 19.06	68.95 ± 18.65	69.05 ± 18.80
% SSIM	88.05 ± 3.89	88.67 ± 3.88	88.83 ± 3.83	89.02 ± 3.67
MSE (m)	0.22 ± 0.16	0.20 ± 0.15	0.19 ± 0.15	0.19 ± 0.15
% $ J_\phi \leq 0$	0.07 ± 0.25	0.04 ± 0.14	0.01 ± 0.04	0.00 ± 0.03
Time [s]	0.0072	0.0072	0.0080	0.0079
Parameters	75,429	300,477	1,199,469	4,793,037
Mult-Adds (M)	10.98	42.89	169.54	674.10
Memory [MB]	2.25	4.64	11.22	31.57

Metrics	Starting Channels - 4xFourier-Net+			
	8	16	32	64
% Dice	77.54 ± 13.73	77.90 ± 13.92	78.33 ± 14.14	78.74 ± 13.91
% Dice*	68.52 ± 18.62	69.21 ± 19.02	69.99 ± 19.21	70.41 ± 19.02
% SSIM	88.70 ± 4.17	88.89 ± 4.05	89.08 ± 4.01	89.29 ± 3.74
MSE (m)	0.19 ± 0.14	0.19 ± 0.14	0.18 ± 0.14	0.18 ± 0.14
% $ J_\phi \leq 0$	0.02 ± 0.07	0.01 ± 0.04	0.01 ± 0.05	0.00 ± 0.00
Time [s]	0.0301	0.0244	0.0297	0.0277
Parameters	301,716	1,201,908	4,797,876	19,172,148
Mult-Adds (G)	0.04	0.17	0.68	2.70
Memory [MB]	7.66	17.23	43.56	124.94

the percentage of non-positive Jacobian determinants actually decreases for smaller crop sizes, perhaps because the displacements on the compressed images are not as extreme and thus more smooth. Note however, that the crop size of 48×48 is a bit of an outlier in this regard breaking the trend with a higher percentage as 40×84 . The time also seems to decrease slightly with a smaller crop, although this is quite noisy as the time from 80×168 to 40×84 is halved for *Fourier-Net+*, but the time needed for 48×48 and 24×24 increases again very slightly. For *4xFourier-Net+* the time decrease for all smaller crop sizes, except for 24×24 .

The number of parameters does not change for different crop sizes as the networks themselves do not change, however the number of Mult-Adds and the total memory still change with the image size. Both decrease with a larger crop size as the image gets smaller. This effect is again not linear as a reduction in image size yields a larger reduction in memory going from 80×168 to 40×84 than from 48×48 to 24×24 . Again, there is no sweet-spot to be found that maximizes both memory efficiency and registration performance similar to the experiments in the previous chapter.

Table 6.7: Results for four different FT crop sizes for *Fourier-Net+* and *4xFourier-Net+* examined on the fully sampled ACDC test data.

Metrics	FT crop size - <i>Fourier-Net+</i>			
	80×168	40×84	48×48	24×24
% Dice	78.24 ± 14.43	76.61 ± 13.90	75.27 ± 13.49	73.77 ± 14.42
% Dice*	70.66 ± 19.70	67.60 ± 19.24	65.48 ± 18.76	63.37 ± 20.07
% SSIM	89.81 ± 4.09	88.58 ± 3.87	87.98 ± 3.91	87.00 ± 3.99
MSE (m)	0.15 ± 0.12	0.20 ± 0.15	0.22 ± 0.16	0.27 ± 0.19
% $ J_\phi \leq 0$	0.22 ± 0.46	0.05 ± 0.15	0.09 ± 0.25	0.00 ± 0.02
Time [s]	0.0158	0.0077	0.0079	0.0081
Parameters	75,429	75,429	75,429	75,429
Mult-Adds (M)	64.03	16.37	10.98	2.74
Memory [MB]	9.51	2.97	2.25	1.12

Metrics	FT crop size - <i>4xFourier-Net+</i>			
	80×168	40×84	48×48	24×24
% Dice	78.12 ± 15.01	77.49 ± 14.67	74.95 ± 14.15	72.58 ± 14.67
% Dice*	70.08 ± 21.92	68.03 ± 21.57	64.54 ± 20.75	61.19 ± 21.63
% SSIM	89.91 ± 4.01	89.03 ± 3.98	87.98 ± 3.94	87.02 ± 3.95
MSE (m)	1.45 ± 1.11	1.81 ± 1.37	2.22 ± 1.61	2.64 ± 1.84
% $ J_\phi \leq 0$	0.12 ± 0.23	0.03 ± 0.15	0.06 ± 0.23	0.02 ± 0.15
Time [s]	0.0390	0.0301	0.0277	0.0289
Parameters	301,716	301,716	301,716	301,716
Mult-Adds (M)	256.14	65.50	43.91	10.98
Memory [MB]	36.70	10.54	7.66	3.15

Comparison with VoxelMorph

In this test *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* are compared to the famous *VoxelMorph* [6]. This unsupervised network brought deep learning based registration approaches into the mainstream and computes a dense displacement field in contrast to the band-limited displacement of the other networks. From the previous experiments a FT crop size of 48×48 and channel size of 16 was chosen to strike a good balance between performance and memory efficiency. The experiment was only done on the fully sampled test data and all networks were trained for 6 epochs. The results can be seen in Table 6.8. *VoxelMorph* performs worst in terms of Dice score with the background label, however it does outperform *Fourier-Net+* in Dice without the background label. The best registration performance in terms of Dice (both with and without the background label) has *Fourier-Net* followed by *4xFourier-Net+* as expected. *VoxelMorph*, however, does perform best in terms of SSIM and MSE followed by *Fourier-Net*, so it seems that the dense network aligns the overall image well, but struggles to compensate the strong changes in the cardiac region between frames. It is conspicuous, however, that the percentage of non-positive Jacobian determinants for *VoxelMorph* is quite high with almost 90% while the other models are all under one percent. This is perhaps caused by the dense displacement which causes a better overall alignment (as indicated by the good SSIM and MSE values)

at the cost of more folding occurring. In terms of time *4xFourier-Net+* is slowest followed by *VoxelMorph*, *Fourier-Net* and *Fourier-Net+*, however all pretty fast ($< 30\text{ms}$). The model with the least amount of parameters and Mult-Adds is *VoxelMorph*, however perhaps due to the dense displacement it needs quite a lot of memory with almost 40 MB. *Fourier-Net* is the largest model in terms of model parameters, Mult-Adds and total memory (90 MB). *Fourier-Net+* and *4xFourier-Net+* both have a higher number of parameters and Mult-Adds, but a lower total amount of memory needed. Overall, *4xFourier-Net+* is the only model that is truly more efficient and has better performance in terms of Dice than *VoxelMorph*, as *Fourier-Net+* is more efficient, but not necessarily better in terms of Dice score, while *Fourier-Net* is far superior in registration performance (as measured by the Dice score), but also less efficient in terms of memory consumption.

Table 6.8: Comparison of *Fourier-Net*, *Fourier-Net+*, *4xFourier-Net+* and *VoxelMorph* with similarity metrics and memory consumption on the fully sampled ACDC test data.

	Fourier-Net	Fourier-Net+	4xFourier-Net+	VoxelMorph
% Dice	78.31 ± 13.96	76.88 ± 13.86	77.90 ± 13.92	75.84 ± 13.46
% Dice*	71.17 ± 18.99	67.86 ± 19.06	69.21 ± 19.02	68.25 ± 18.36
% SSIM	91.53 ± 3.49	88.67 ± 3.88	88.89 ± 4.05	93.53 ± 3.30
MSE (m)	0.09 ± 0.07	0.20 ± 0.15	0.19 ± 0.14	0.06 ± 0.04
$\% J_\phi \leq 0$	0.44 ± 0.45	0.04 ± 0.14	0.01 ± 0.04	89.96 ± 1.32
Time [s]	0.0099	0.0071	0.0244	0.0145
Parameters	1,735,447	300,477	1,201,908	84,322
Mult-Adds (G)	2.35	0.04289	0.17157	0.00157
Memory [MB]	90.08	4.64	17.23	39.04

Dense Displacement on Accelerated Data

The difference between a dense displacement field and a band-limited one was already explored in section 6.2. However, the results in Table 6.5 only include fully sampled data, not accelerated data. These new tests again included *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+*, but are extended to subsampled data. Results for Acc4 data can be seen in Table 6.9, for Acc8 data in Table 6.10 and for Acc10 in Table 6.11.

For Acc4, the dense version of *Fourier-Net* performed worse than the band-limited version in terms of Dice, as we expected, however this is not the case for *Fourier-Net+* and *4xFourier-Net+* where the performance decreases for the band-limited versions as indicated by the dice score. The same is true for the SSIM and MSE values. The percentage of non-positive Jacobian determinants is higher for the band-limited *Fourier-Net*, while band-limited *Fourier-Net+* and *4xFourier-Net+* show no folding. While this at first seems like a positive metric it can also indicate a lack of complex displacements which could explain the lower performance. All band-limited versions are faster due to the smaller network size as the encoder lack some layers compared to the dense versions.

For Acc8 all band-limited networks perform worse in terms of Dice, SSIM and MSE. The percentage of non-positive Jacobian determinants is again higher for the band-limited

Fourier-Net compared to the dense version, while the band-limited *Fourier-Net+* and *4xFourier-Net+* show no folding. The band-limited versions surprisingly are slower compared to the dense versions except for *Fourier-Net+*.

For Acc10, similar to Acc4, band-limited *Fourier-Net* performs better in terms of Dice, SSIM and MSE, while band-limited *Fourier-Net+* and *4xFourier-Net+* are again worse compared to the dense versions. The percentage of non-positive Jacobian determinants for the band-limited *Fourier-Net* is lower than for the dense version this time, while band-limited *Fourier-Net+* and *4xFourier-Net+* again show no folding. As for the Acc4 data, the band-limited networks are faster in terms of inference time.

The results for the accelerated data are not quite consistent as the band-limited *Fourier-Net* performs better for Acc4 and Acc10, but not for Acc8. Band-limited *Fourier-Net+* and *4xFourier-Net+* perform consistently worse than their dense versions across all acceleration levels. The band-limiting of the displacement seems to help with avoiding folding and in most cases decrease inference time. Our hypothesis that the advantage of the dense displacement compared to the band-limited displacement disappears for accelerated data does not seem to hold except for *Fourier-Net* on Acc4 and Acc10 data.

Table 6.9: Results for *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* with both dense and band-limited displacement fields on the Acc4 ACDC test data.

Metrics	Dense Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	67.84 \pm 14.07	73.84 \pm 15.18	72.83 \pm 15.64
% Dice*	56.14 \pm 18.45	64.31 \pm 21.00	63.19 \pm 21.59
% SSIM	71.91 \pm 4.85	77.58 \pm 10.64	77.32 \pm 10.53
MSE (m)	0.28 \pm 0.12	0.20 \pm 0.14	0.21 \pm 0.15
% $ J_\phi \leq 0$	0.37 \pm 0.37	0.15 \pm 0.26	0.15 \pm 0.27
Time [s]	0.0082	0.0092	0.0337
Metrics	Band-limited Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	72.30 \pm 14.04	70.18 \pm 17.43	70.36 \pm 16.63
% Dice*	61.87 \pm 19.32	59.36 \pm 23.90	59.47 \pm 22.80
% SSIM	78.77 \pm 7.22	75.78 \pm 10.63	75.91 \pm 10.72
MSE (m)	0.16 \pm 0.10	0.28 \pm 0.19	0.27 \pm 0.19
% $ J_\phi \leq 0$	0.46 \pm 0.54	0.00 \pm 0.00	0.00 \pm 0.00
Time [s]	0.0068	0.0079	0.0183

Table 6.10: Results for *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* with both dense and band-limited displacement fields on the Acc8 ACDC test data.

Metrics	Dense Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	69.12 ± 14.05	74.20 ± 14.50	74.36 ± 14.97
% Dice*	57.99 ± 18.66	64.50 ± 20.57	65.01 ± 20.82
% SSIM	77.27 ± 3.74	86.08 ± 4.19	85.53 ± 3.99
MSE (m)	0.21 ± 0.10	0.15 ± 0.12	0.15 ± 0.12
% $ J_\phi \leq 0$	0.47 ± 0.30	0.15 ± 0.27	0.15 ± 0.28
Time [s]	0.0117	0.0169	0.0412

Metrics	Band-limited Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	66.07 ± 13.72	71.13 ± 15.37	73.13 ± 15.26
% Dice*	53.76 ± 17.03	60.10 ± 21.34	62.61 ± 21.39
% SSIM	73.34 ± 3.83	84.02 ± 4.36	84.61 ± 4.44
MSE (m)	0.26 ± 0.09	0.21 ± 0.16	0.19 ± 0.15
% $ J_\phi \leq 0$	0.78 ± 0.37	0.00 ± 0.00	0.01 ± 0.04
Time [s]	0.0138	0.0133	0.0418

Table 6.11: Results for *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* with both dense and band-limited displacement fields on the Acc10 ACDC test data.

Metrics	Dense Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	69.78 ± 14.35	73.62 ± 15.13	73.88 ± 15.05
% Dice*	58.27 ± 19.54	63.39 ± 21.54	64.18 ± 21.00
% SSIM	83.27 ± 2.86	89.69 ± 3.34	89.83 ± 3.27
MSE (m)	0.16 ± 0.10	0.14 ± 0.12	0.13 ± 0.12
% $ J_\phi \leq 0$	0.37 ± 0.31	0.21 ± 0.39	0.12 ± 0.19
Time [s]	0.0112	0.0116	0.0433

Metrics	Band-limited Displacement		
	Fourier-Net	Fourier-Net+	4xFourier-Net+
% Dice	72.28 ± 14.65	71.83 ± 15.91	72.55 ± 15.66
% Dice*	62.58 ± 19.99	61.10 ± 22.07	62.00 ± 21.97
% SSIM	90.68 ± 2.93	87.77 ± 3.50	89.13 ± 3.47
MSE (m)	0.10 ± 0.08	0.19 ± 0.15	0.17 ± 0.15
% $ J_\phi \leq 0$	0.23 ± 0.26	0.00 ± 0.01	0.00 ± 0.01
Time [s]	0.0091	0.0088	0.0310

Comparison on Subsampled Data

In section 5.2, *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* were already compared to *VoxelMorph*, however, these comparisons were only done on fully sampled data, not accelerated ones. To further add to the comparison, *NiftyReg* was again used as a traditional registration algorithm. The unaligned test image pairs were used as a baseline for the lower bound of registration performance. Values which are worse than the baseline are marked in red, while the best results for each acceleration level is highlighted with blue. The results can be seen in Table 6.12. All times are computed on CPU due to *NiftyReg* only working on the CPU as the GPU capable versions were deprecated.

Table 6.12: Test results for *NiftyReg*, *VoxelMorph*, *Fourier-Net*, *Fourier-Net+* and *4xFourier-Net+* on the ACDC test data with an unaligned baseline for comparison. The best results for each metric and subsampling are highlighted in blue, while values worse than the unaligned baseline are marked with red.

	Method	% DICE	% DICE*	% SSIM	MSE (m)	Time [s]
$R=0$	Baseline	70.85 \pm 18.27	60.35 \pm 25.24	86.39 \pm 4.08	0.33 \pm 0.23	-
	NiftyReg	70.74 \pm 13.77	57.56 \pm 18.86	91.26 \pm 3.18	1.94 \pm 1.61	20
	VoxelMorph	75.84 \pm 13.46	68.25 \pm 18.36	93.53 \pm 3.30	0.06 \pm 0.04	0.1845
	Fourier-Net	78.31 \pm 13.96	71.17 \pm 18.99	91.53 \pm 3.49	0.09 \pm 0.07	0.1918
	Fourier-Net+	76.88 \pm 13.86	67.86 \pm 19.06	88.67 \pm 3.88	0.20 \pm 0.15	0.0893
	4xFourier-Net+	77.90 \pm 13.92	69.21 \pm 19.02	88.89 \pm 4.05	0.19 \pm 0.14	0.3262
$R=4$	Baseline	70.85 \pm 18.27	60.35 \pm 25.24	76.80 \pm 11.02	0.28 \pm 0.19	-
	NiftyReg	69.89 \pm 13.73	56.47 \pm 17.86	86.03 \pm 6.41	0.15 \pm 0.14	80.08
	VoxelMorph	73.45 \pm 13.40	63.75 \pm 18.51	78.56 \pm 7.32	0.15 \pm 0.08	0.1858
	Fourier-Net	72.30 \pm 14.04	61.87 \pm 19.32	78.77 \pm 7.22	0.16 \pm 0.10	0.1890
	Fourier-Net+	70.18 \pm 17.43	59.36 \pm 23.90	75.78 \pm 10.63	0.28 \pm 0.19	0.0935
	4xFourier-Net+	70.36 \pm 16.63	59.47 \pm 22.80	75.91 \pm 10.72	0.27 \pm 0.19	0.3462
$R=8$	Baseline	70.85 \pm 18.27	60.35 \pm 25.24	85.35 \pm 4.43	0.22 \pm 0.17	-
	NiftyReg	70.04 \pm 13.42	56.37 \pm 17.63	91.07 \pm 2.80	0.12 \pm 0.13	88.36
	VoxelMorph	72.94 \pm 14.20	64.27 \pm 18.97	88.32 \pm 3.01	0.09 \pm 0.06	0.1725
	Fourier-Net	66.07 \pm 13.72	53.76 \pm 17.03	73.34 \pm 3.83	0.26 \pm 0.09	0.2109
	Fourier-Net+	71.13 \pm 15.37	60.10 \pm 21.34	84.02 \pm 4.36	0.21 \pm 0.16	0.1487
	4xFourier-Net+	73.13 \pm 15.26	62.61 \pm 21.39	84.61 \pm 4.44	0.19 \pm 0.15	0.5373
$R=10$	Baseline	70.85 \pm 18.27	60.35 \pm 25.24	89.17 \pm 3.58	0.20 \pm 0.17	-
	NiftyReg	70.40 \pm 13.34	56.61 \pm 17.71	93.47 \pm 2.37	0.11 \pm 0.13	47.44
	VoxelMorph	72.46 \pm 13.20	62.82 \pm 18.08	88.27 \pm 2.36	0.10 \pm 0.06	0.1173
	Fourier-Net	72.28 \pm 14.65	62.58 \pm 19.99	90.68 \pm 2.93	0.10 \pm 0.08	0.1088
	Fourier-Net+	71.83 \pm 15.91	61.10 \pm 22.07	87.77 \pm 3.50	0.19 \pm 0.15	0.0438
	4xFourier-Net+	72.55 \pm 15.66	62.00 \pm 21.97	89.13 \pm 3.47	0.17 \pm 0.15	0.1539

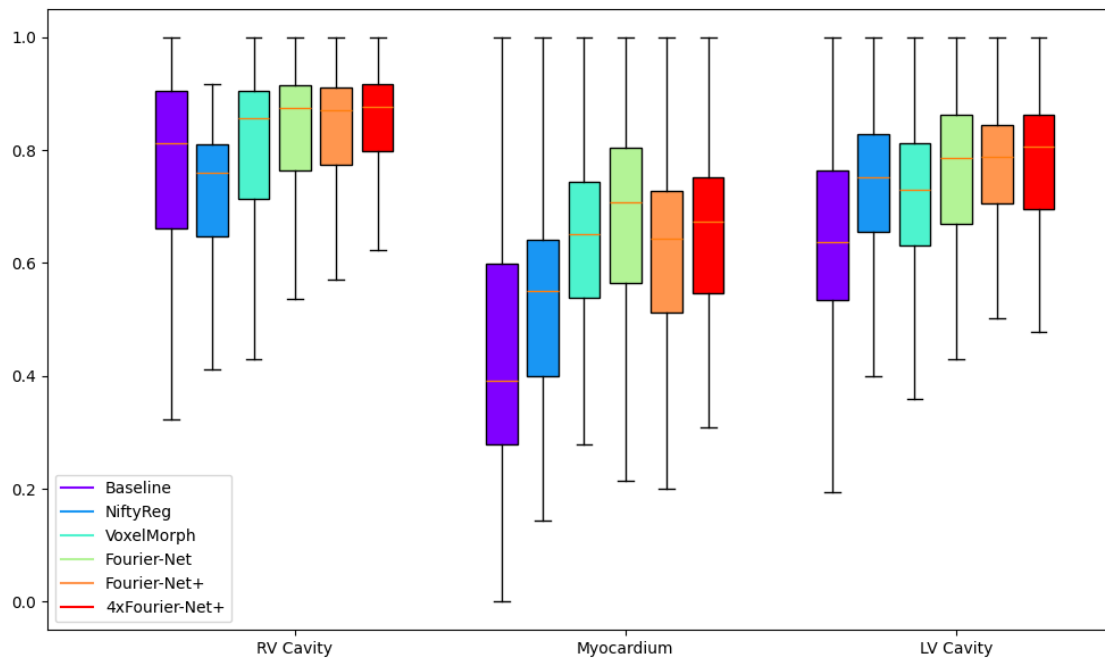


Figure 6.13: Boxplot of the Dice scores for all models sorted by label (excluding the background) on the fully sampled *ACDC* test data.

7

Conclusion

Summery of all stuff...

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