



# Deep Learning for Classification and Selection of Cine CMR Images to Achieve Fully Automated Quality-Controlled CMR Analysis From Scanner to Report

#### **OPEN ACCESS**

#### Edited by:

Paul Leeson, University of Oxford, United Kingdom

#### Reviewed by:

Richard Nethononda, Chris Hani Baragwanath Hospital, South Africa Emmanuel Androulakis, Royal Brompton and Harefield NHS Foundation Trust, United Kingdom

#### \*Correspondence:

Bram Ruijsink jacobus.ruijsink@kcl.ac.uk Vittoria Vergani vergani.vittoria@gmail.com

<sup>†</sup>These authors have contributed equally to this work and share last authorship

#### Specialty section:

This article was submitted to Cardiovascular Imaging, a section of the journal Frontiers in Cardiovascular Medicine

> Received: 16 July 2021 Accepted: 09 September 2021 Published: 14 October 2021

#### Citation:

Vergani V, Razavi R, Puyol-Antón E and Ruijsink B (2021) Deep Learning for Classification and Selection of Cine CMR Images to Achieve Fully Automated Quality-Controlled CMR Analysis From Scanner to Report. Front. Cardiovasc. Med. 8:742640. doi: 10.3389/fcvm.2021.742640 Vittoria Vergani<sup>1\*</sup>, Reza Razavi<sup>1,2</sup>, Esther Puyol-Antón<sup>1†</sup> and Bram Ruijsink<sup>1,3\*†</sup>

<sup>1</sup> School of Biomedical Engineering and Imaging Sciences, King's College London, London, United Kingdom, <sup>2</sup> Department of Adult and Paediatric Cardiology, Guy's and St. Thomas' NHS Foundation Trust, London, United Kingdom, <sup>3</sup> Department of Cardiology, Division of Heart and Lungs, University Medical Centre Utrecht, Utrecht, Netherlands

# **Introduction:** Deep learning demonstrates great promise for automated analysis of CMR. However, existing limitations, such as insufficient quality control and selection of target acquisitions from the full CMR exam, are holding back the introduction of deep learning tools in the clinical environment. This study aimed to develop a framework for automated detection and quality-controlled selection of standard cine sequences images from clinical CMR exams, prior to analysis of cardiac function.

**Materials and Methods:** Retrospective study of 3,827 subjects that underwent CMR imaging. We used a total of 119,285 CMR acquisitions, acquired with scanners of different magnetic field strengths and from different vendors (1.5T Siemens and 1.5T and 3.0T Phillips). We developed a framework to select one good acquisition for each conventional cine class. The framework consisted of a first pre-processing step to exclude still acquisitions; two sequential convolutional neural networks (CNN), the first (CNN<sub>class</sub>) to classify acquisitions in standard cine views (2/3/4-chamber and short axis), the second (CNN<sub>QC</sub>) to classify acquisitions according to image quality and orientation; a final algorithm to select one good acquisition of each class. For each CNN component, 7 state-of-the-art architectures were trained for 200 epochs, with cross entropy loss and data augmentation. Data were divided into 80% for training, 10% for validation, and 10% for testing.

**Results:**  $CNN_{class}$  selected cine CMR acquisitions with accuracy ranging from 0.989 to 0.998. Accuracy of  $CNN_{QC}$  reached 0.861 for 2-chamber, 0.806 for 3-chamber, and 0.859 for 4-chamber. The complete framework was presented with 379 new full CMR studies, not used for CNN training/validation/testing, and selected one good 2-, 3-, and 4-chamber acquisition from each study with sensitivity to detect erroneous cases of 89.7, 93.2, and 93.9%, respectively.

1

**Conclusions:** We developed an accurate quality-controlled framework for automated selection of cine acquisitions prior to image analysis. This framework is robust and generalizable as it was developed on multivendor data and could be used at the beginning of a pipeline for automated cine CMR analysis to obtain full automatization from scanner to report.

Keywords: cardiac magnetic resonance, deep learning, quality control, cardiac function, view-selection, multivendor

#### INTRODUCTION

Cardiac magnetic resonance (CMR) is the state-of-the-art clinical tool to assess cardiac morphology, function, and tissue characterization (1), and both European and American guidelines advocate its use to diagnose and monitor a large number of cardiovascular diseases (2, 3). The role of CMR continues to grow due to the technical advances that allow increasingly detailed analysis of the cardiovascular system.

However, systematic manual analysis of the different CMR sequences which are acquired during a typical CMR exam is highly time consuming, where the bulk of the time is taken up by repetitive tasks, such as image identification, selection, and segmentation, which are at the basis of CMR post-processing.

Deep learning (DL), a branch of artificial intelligence (AI), is securing an emergent role in the field of CMR, as it provides for automatization of repetitive tasks, significantly reducing the time required for image analysis, while maintaining a high degree of accuracy (4, 5). Physicians' time can thus be optimized and targeted for critical review of clinical and imaging information to reach a correct diagnosis. Moreover, automated analysis allows access to biomarkers of cardiac function that would normally be too labor intensive to obtain, such as peak ejection and filling rates from ventricular volume curves (6, 7) or atrioventricular valve planar motion (8) from long-axis segmentations.

Several groups have shown promising results on the implementation of DL in the analysis of CMR, including segmentation of cine images to derive cardiac function (5), analysis of perfusion defects to detect inducible ischemia (9), and assessment of late gadolinium enhancement and T1 mapping to aid tissue characterization (10, 11).

However, some limitations still need to be addressed before a widespread clinical adoption of DL tools, such as steps to perform automated selection of target images from the full CMR exam, as well as robust systems to flag inadequate quality of image or of analysis, which are the necessary steps that precede analysis of CMR sequences in the clinical setting. We have previously shown that comprehensive quality-control can be introduced into a DL pipeline for accurate, automated analysis of cine CMR images that adheres to clinical safety standards (5). On the other

hand, DL has not yet been systematically implemented for image recognition and selection prior to analysis in CMR.

This study aimed at developing a framework for automated identification and quality-controlled (QC) selection of cine images used for cardiac function analysis from routine clinical CMR exams. This framework was then implemented as the first step of a larger pipeline for QC CMR analysis of cine images previously developed by our group (5).

#### MATERIALS AND METHODS

The framework we present is composed of a set of algorithms combined with two convolutional neural networks (CNN) aimed at identifying conventional cine views (CNN<sub>class</sub>) and at sorting these images according to quality into "correct" and "wrong" (CNN<sub>QC</sub>). This construction allows for the framework, presented with a full CMR exam, to perform a selection of one good quality acquisition for each conventional cine class, which is then used for analysis of cardiac function, and to flag exams when no image of sufficient quality could be identified (see Figure 1).

The developed pipeline was implemented in Python using standard libraries such as Numpy and Pandas as a dedicated deep learning library Pytorch.

#### **Study Population**

This is a retrospective multivendor study conducted on a large CMR dataset. 3,445 CMR exams were included: 1,510 from UK Biobank and 1,935 from Guy's and St. Thomas' NHS Foundation Trust (GSTFT), London, comprising of a total of 119,285 individual CMR acquisitions. Images were acquired on 1.5T Siemens and 1.5T and 3.0T Phillips CMR scanners using a large variety of protocols, with variable voxel- and image-size, acquisition techniques and under-sampling factors.

Our dataset was randomly selected from the pool of available studies in the UK Biobank and GSTFT databases. All exams were acquired between 2004 and 2020. The random selection was used to obtain a heterogeneous population, including both healthy and pathological hearts, with a variety of cardiac pathologies (ischemic heart disease, dilated and hypertrophic cardiomyopathy, valvular heart disease, adult congenital heart disease (ACHD), and others). In the case of grossly disruptive artifacts (e.g., device artifacts covering the majority of the chambers), or grossly distorted anatomy (e.g., patients with single-ventricle morphology or Ebstein's disease), CMR exams were excluded from the dataset.

Abbreviations: 2Ch, 2-Chamber view; 3Ch, 3-Chamber view; 4Ch, 4-Chamber view; ACHD, adult congenital heart disease; AI, artificial intelligence; CMR, cardiac magnetic resonance; CNN, convolutional neural network; DL, deep learning; GE, General Electrics; GSTFT, Guy's and St. Thomas' NHS Foundation Trust; LVOT, left ventricular outflow tract; QC, quality control; SAX, short axis.



the first to classify images in standard cine views (2/3/4-chamber and short axis), the second to classify images according to image quality and orientation; a final algorithm to select one good image of each class. This construction allows for the framework, presented with a full CMR exam, to perform a quality-controlled selection of one good image for each conventional cine class, which is then used for analysis of cardiac function. Ch, chamber; CNN, convolutional neural network; LAX, long axis; SAX, short axis; QC, quality control.

# Identification of Conventional Cine Classes

The first part of our framework aimed to identify the standard cine views used for analysis of cardiac function from the complete exam. These were the 2-Chamber view (2Ch), 3-Chamber view (3Ch), 4-Chamber view (4Ch), and the short-axis stack (SAX).

First, all single-frame acquisitions were excluded from the exams based on the dicom-header information. This allowed to reduce the bulk of acquisitions available in a complete CMR exam to a set of multi-frame acquisitions. The 2Ch, 3Ch, 4Ch, and SAX sequences were subsequently selected from the remaining acquisitions using our view selection algorithm. To identify the class of the cine acquisitions, only a single frame of the acquisition is needed. Therefore, the first frames of each of the remaining data was used for training. All data was manually classified into the conventional cine classes by an expert physician (2,905 2Ch, 1,171 3Ch, 2,963 4Ch, 9,112 SAX images) and a class of "other" (4,043). In case of doubt, a second opinion was sought, and a decision was made by consensus. Prior to CNN training, all images were cropped to a standard size of  $256 \times 256$  pixels and converted to numpy arrays.

The manually classified data were divided as follows: 80% was used for training, 10% was used for validation, and 10% for testing. The training, validation and test data cohorts had a mutually exclusive subject pool, i.e., acquisitions from the same subject could only be used in one of the three cohorts. We trained seven state-of-the-art CNN architectures: AlexNet (12), DenseNet (13), MobileNet (14), ResNet (15), ShuffleNet (16), SqueezeNet (17), and VGG (18). Each network was trained for 200 epochs with cross entropy loss, to classify end-diastolic images into the five classes described. For training data, data augmentation was performed on-the-fly using random translations ( $\pm 30$  pixels), rotations ( $\pm 90^{\circ}$ ), flips (50%) probability) and scalings (up to 20%) to each mini-batch of images before feeding them to the network. The probability of augmentation for each of the parameters was 50%. Augmentation is the only technique we use to prevent over-fitting, as other techniques were not found to improve performance and their omission contributed to a simpler network architecture.

An additional algorithm was used after  $\text{CNN}_{\text{class}}$  to check complete classification of the short axis acquisition. An image classified as SAX was confirmed to belong to a short axis acquisition if the following two criteria, screened by the algorithm, were met: (1) the image belonged to a stack composed of a minimum of 8 slices, (2) at least 2 out the 3 central images of the stack were classified as short axis by  $\text{CNN}_{\text{class}}$ .

#### **Quality Control of Selected Images**

The second part of our framework aimed to scrutinize the quality of the identified long-axis cine images. QC of short axis acquisitions was not performed in this step, as our downstream pipeline for automated CMR analysis already includes short-axis QC (5).

To train the networks for QC ( $CNN_{QC}$ ), a set of 2Ch (1,937), 3Ch (1,591), 4Ch (2,003) images from our database were reviewed by an expert physician and classified as "correct" or "wrong" based on whether image quality was satisfactory for subsequent analysis. Images that included

mis-triggering, breathing, implant or fold-over artifacts were classified as "wrong" if the detection of myocardial borders was hindered. Moreover, all off-axis orientations (i.e., presence of left ventricular outflow tract (LVOT) in 4Ch, fore-shortening of the apex, absence of any of the valves in 3Ch) were also classified as "wrong." In case of doubt, a second opinion was sought, and a decision was made by consensus. The resulting database consisted of 1,444 "correct" and 493 "wrong" 2Ch, 1,098 "correct" and 493 "wrong" 3Ch images and 1,393 "correct" and 610 "wrong" 4Ch images.

The manually classified data were used to train QC networks for each class (2Ch-CNN<sub>QC</sub>, 3Ch-CNN<sub>QC</sub>, 4Ch-CNN<sub>QC</sub>). The data were divided as follows: 80% training, 10% for validation, 10% for testing. We trained the same seven CNN architectures as described in the previous section. We used the same training process as described for training of CNN<sub>class</sub>, with the difference that CNN<sub>OC</sub> was trained as binary classifiers, i.e., a two-class classification problem as opposed to five, and therefore used binary cross entropy with a logit loss function. Additionally, we implemented an adaptive learning rate scheduler, which decreases the learning rate by a constant factor of 0.1 after 5 epochs stopping on plateau on the validation/test set (commonly known as ReduceLRonPlateau). This step was added as it improves CNN training when presented with unbalanced datasets. The CNN<sub>OC</sub>'s output was a binary classification ("correct" vs. "wrong"), as well as the probabilities (i.e., certainty) associated with the classification for each case.

# Complete Framework: From Full Study to Selection

To complete the framework, the  $\text{CNN}_{\text{class}}$  and  $\text{CNN}_{\text{QC}}$  were combined with a final selection algorithm. This algorithm selected one good quality acquisition of each standard cine view for image analysis, when multiple acquisitions of a single class were present in the exam. For long axis data, it did so by identifying the case with the highest probability of being scored "correct" by the  $\text{CNN}_{\text{QC}}$ . For short axis, the stack with the highest probability of belonging to SAX (obtained from the output of the  $\text{CNN}_{\text{class}}$ ) was selected. If any of the classes was absent in an exam, or the framework did not identify an image of sufficient quality, the case was flagged for clinician review.

As the individual components act in series in the complete framework, their sequential action will yield an overall performance that is different from the sum of the individual ones. To find the best combination of components, each possible combination of the trained CNN architectures trained in the previous steps was tested using an additional testset of 379 scans randomly selected from the database, not previously used for CNN training. For each exam, a manual operator selected the best cine long and short axis acquisitions. To determine the intra- and inter-observer variability present in the manual analysis, 100 randomly selected scans were re-analyzed by the same operator and by a second operator.

#### TABLE 1 | Population characteristics.

		CNN training	Complete framework
Number		3,445	379
Age (years)		$57 \pm 16$	$49 \pm 19$
Sex (males)		1,911 (55)	228 (60)
Height (cm)		$176\pm32$	$171 \pm 18$
Weight (kg)		$79\pm18$	$80\pm19$
BMI (kg/m <sup>2</sup> )		$27 \pm 5$	$27 \pm 7$
Ethnicity	Caucasian	2,401 (69.7)	231 (60.9)
	Afro-Caribbean	172 (5.0)	54 (14.2)
	Asian	85 (2.5)	10 (2.6)
	Other	21 (0.6)	7 (1.8)
	Not stated	766 (22.2)	77 (20.3)
Cardiac pathology	Healthy	1,886 (54.7)	68 (17.9)
	IHD	315 (9.1)	43 (11.3)
	DCM	167 (4.8)	27 (7.1)
	HCM	77 (2.2)	16 (4.2)
	ACHD	185 (5.4)	59 (15.6)
	Valvular	133 (3.9)	37 (9.8)
	Vascular	104 (3.0)	32 (8.4)
	Arrhythmic	159 (4.6)	26 (6.9)
	Other	419 (12.2)	71 (18.7)

Age, sex, height, weight, and cardiac pathology of subjects used for training of CNNs, framework and full pipeline validation. All continuous values are reported as mean  $\pm$  standard deviation, while categorical variables are reported as number (percentage). ACHD, adult congenital heart disease (excluding valvular and vascular abnormalities); CNN, convolutional neural network; HCM, hypertrophic cardiomyopathy; IHD, ischaemic heart disease; SD, standard deviation.

#### **Statistics**

#### **Class Identification CNN**

Precision, recall, and F1-score of each class ("4Ch," "3Ch," "2Ch," "SAX," "other") and overall accuracy were computed at test-time to evaluate the performance of each trained CNN<sub>class</sub>.

#### **Quality Control CNNs**

Precision, recall, and F1-score of each class ("correct," "wrong") and overall accuracy were assessed to evaluate the performance at test-time of each trained  $2Ch/3Ch/4Ch-CNN_{QC}$ .

#### **Complete Framework**

Sensitivity (defined as: the percentage of incorrect cases identified as incorrect), specificity (defined as: the percentage of correct cases identified as correct), and balanced accuracy were computed for each framework. Cohen kappa coefficient was used to assess intra- and inter-observer variability.

#### **Full Pipeline: From Scanner to Report**

Finally, we added the complete framework as the first step of our previously validated pipeline for quality-controlled AIbased analysis of cardiac function from CMR (5). Broadly, this pipeline consists of quality-controlled image segmentation and analysis of cine images to obtain LV and RV volumes and mass, LV ejection and filling dynamics, and longitudinal, radial and circumferential strain.

		AlexNet		J	DenseNet		Σ	MobileNet		-	ResNet		S	ShuffleNet	it	Ň	SqueezeNet	et		VGG	
. —	Precisior	Recall I	<sup>r</sup> 1-score	Precision	Recall F	F1-score	Precision	Recall F	1-score F	Precision	Recall F	-1-score	Precision	Recall	F1-score	Precisior	ר Recall	Precision Recall F1-score	Precision	Recall F	1-score
2-chamber	0.998	0.993	0.996	0.998	1.000	0.999	0.997	1.000	0.998	0.998	1.000	0.999	0.993	1.000	0.997	0.986	0.991	0.989	0.993	1.000	0.997
3-chamber	0.991	0.953	0.972	1.000	1.000	1.000	1.000	0.996	0.998	0.996	0.996	0.996	1.000	0.983	0.991	0.987	0.979	0.983	0.987	0.987	0.987
4-chamber	1.000	0.990	0.995	1.000	0.998	0.999	0.990	0.995	0.992	0.998	0.995	0.997	0.997	1.000	0.998	0.987	0.992	0.989	0.993	0.997	0.995
Short axis	0.995	0.999	0.997	0.996	0.999	0.998	0.997	1.000	0.998	0.999	0.999	0.999	0.996	1.000	0.998	0.992	0.997	0.994	0.996	1.000	0.998
Other	0.973	0.985	0.979	0.998	0.990	0.994	0.996	0.984	066.0	0.994	0.995	0.994	0.996	0.984	066.0	0.984	0.969	0.976	0.995	0.978	0.986
Accuracy		0.992			0.998			0.996			0.998			0.996			0.989			0.994	

Frontiers in Cardiovascular Medicine | www.frontiersin.org

		AlexNet		-	DenseNet	it.	2	MobileNet			ResNet		S	Shufflenet		Sq	SqueezeNet	st		VGG	
. —	Precisior	ו Recall	F1-score	Precision Recall F1-score precision recall F1-score	recall	F1-score	precisior	recall	-1-score	Precision	recall F	1-score	precision recall F1-score	recall F	-1-score	precision	recall	-1-score	precision	recall F	-1-score
2Ch-CNN <sub>QC</sub>	2 2																				
Correct	0.856	0.844	0.850	0.931	0.841	0.884	0.889	0.917	0.903	0.877	0.910	0.893	0.881	0.869	0.875	0.755	0.993	0.858	0.873	0.955	0.912
Wrong	0.536	0.559	0.547	0.620	0.806	0.701	0.714	0.645	0.678	0.683	0.602	0.640	0.608	0.634	0.621	0.000	0.000	0.000	0.803	0.570	0.667
Accuracy		0.775			0.832			0.851			0.835			0.812			0.751			0.861	
3Ch-CNNac	30																				
Correct	0.788	0.795	0.792	0.815	0.923	0.866	0.850	0.873	0.861	0.830	0.868	0.849	0.880	0.800	0.838	0.690	1.000	0.816	0.824	0.873	0.848
Wrong	0.536	0.525	0.531	0.757	0.535	0.627	0.699	0.657	0.677	0.674	0.606	0.638	0.630	0.758	0.688	0.000	0.000	0.000	0.674	0.586	0.627
Accuracy		0.712			0.803			0.806			0.787			0.786			0.690			0.784	
4Ch-CNN <sub>QC</sub>	50																				
Correct	0.877	0.821	0.848	0.900	0.900	0.900	0.891	0.878	0.884	0.878	0.900	0.888	0.906	0.832	0.867	0.705	1.000	0.827	0.891	0.878	0.884
Wrong	0.630	0.726	0.675	0.761	0.761	0.761	0.719	0.744	0.731	0.745	0.701	0.722	0.664	0.795	0.724	0.000	0.000	0.000	0.719	0.744	0.731
Accuracy		0.793			0.859			0.838			0.841			0.821			0.705			0.838	

We present the feasibility and importance of a fully automated multi-step QC pipeline by (1) running 700 new CMR exams cases (not earlier seen during training) through the pipeline, and (2) presenting a video (**Supplementary Video 1**) of the full analysis for a good-quality case. For the 700 cases we ran through the pipeline we report the average time for selection and complete cine analysis from a full CMR study, and again report sensitivity, specificity and balanced accuracy of error detection.

# RESULTS

#### **Study Population**

Of the 3,827 CMR exams used for this study 3,448 were used for the training and validation of  $\text{CNN}_{\text{class}}$  and  $\text{CN}_{\text{QC}}$  [1,026 acquisition of 16,151 (6.4%) were excluded because of grossly disruptive artifacts or grossly distorted anatomy]. These included patients undergoing clinical scans at GSTFT as well as subjects voluntarily enrolling onto the UK BioBank project, yielding a heterogeneous population in terms of sex (55% male) and clinical presentation (43% healthy, the remaining displaying a wide variety of cardiovascular pathologies, as shown in **Table 1**).

The remaining 379 CMR scans were used to test the complete framework. These were all selected from the GSTFT database to obtain a population representative of routine clinical practice. Demographic characteristics are comparable to the training population, but clinical presentation was more variable, with only 18% of patients having no cardiovascular pathology.

Population characteristics are summarized in Table 1.

#### **Class Identification CNN**

Precision, recall, F1-score, accuracy for all  $CNN_{class}$  are presented in **Table 2**. All trained architectures showed excellence performance, with accuracy ranging from 0.989 to 0.998. DenseNet and ResNet reached highest accuracy, i.e., 0.988. DenseNet showed best precision, recall and F1-score for conventional cine classes: 0.998, 1.00, 0.999 for 2Ch, 1.00, 1.00, 1.00 for 3Ch, 1.00, 0.998, 1.00 for 4Ch, and 0.996, 0.999, 0.998 for SAX.

# **Quality Control CNN**

Precision, recall, F1-score, and accuracy for all  $\text{CNN}_{QC}$  are shown in **Table 3**. Accuracy was variable for different architectures and ranged from 0.751 to 0.861 for 2Ch, from 0.690 to 0.806 for 3Ch, and from 0.705 to 0.859 for 4Ch. Precision, recall and F1score were consistently lower for the "wrong" class compared to the "correct" class for all trained architectures and across the 3 different chamber views.

# **Complete Framework**

Sensitivity, specificity, and balanced accuracy of each constructed framework to identify and select one good quality 2Ch, one good quality 3Ch, and one good quality 4Ch image for each exam are shown in **Table 4**. For the sake of brevity, we present the results of one  $\text{CNN}_{\text{class}}$ , i.e., DenseNet, given the very high and similar performance of all different architectures, combined with all possible  $\text{CNN}_{\text{QC}}$ 's.

#### TABLE 4 | Framework validation.

			Fra	amework for im	age identification	on and selection			
				Der	nseNet CNN <sub>class</sub>	+			
		2Ch-CNN <sub>QC</sub>			3Ch-CNN <sub>QC</sub>			4Ch-CNN <sub>QC</sub>	
Network	SEN	SPE	BACC	SEN	SPE	BACC	SPE	SEN	BACC
AlexNet	79.4	89.8	84.6	84.4	85.8	85.1	70.3	90.9	80.6
DenseNet	75.0	91.1	83.0	93.2	85.3	89.2	88.2	90.9	89.5
MobileNet	78.1	92.4	85.2	88.9	85.9	87.4	72.2	91.9	82.1
ResNet	77.8	91.4	84.6	88.9	85.1	87.0	83.3	90.9	87.1
ShuffleNet	89.7	91.5	90.6	88.9	85.9	87.4	93.9	89.2	91.6
SqueezeNet	55.3	88.2	71.8	51.8	91.0	71.4	23.3	95.5	59.4
VGG	84.4	91.8	88.1	84.8	85.1	84.9	79.4	91.4	85.4

SEN, Sensitivity; SPE, Specificity; and BACC, balanced accuracy of each constructed framework for image identification and selection. Best performing framework per class is highlighted in blue.

The best performing framework was: DenseNet  $CNN_{class}$  + ShuffleNet 2Ch- $CNN_{QC}$  (sensitivity = 89.7%, specificity = 91.5%, balanced accuracy = 90.6%), DenseNet 3Ch- $CNN_{QC}$  (sensitivity = 93.2%, specificity = 85.3%, balanced accuracy = 89.2%), ShuffleNet 4Ch- $CNN_{QC}$  (sensitivity = 93.9%, specificity = 89.2%, balanced accuracy = 91.6%).

Cohen's k for intra- and inter-observer agreement for the same manual operator and between the two different operators were 0.79 and 0.60, respectively.

#### **Full Pipeline: From Scanner to Report**

The sensitivity, specificity and balanced accuracy of the integrated view selection and quality-controlled cardiac analysis pipelines was 96.3, 85.0, and 90.6%, respectively. Performance was also assessed for healthy and pathological cases separately. Results are summarized in **Table 5**.

The average time for selection and complete cine analysis from a full CMR study was between 4 and 7 min for a clinical CMR exam.

**Supplementary Video 1** portrays how implementing the new framework prior to segmentation results in good quality analysis.

#### DISCUSSION

In this study we present a DL-based framework to identify all conventional cine views from a full CMR exam, and subsequently select one image per class of good quality for further automated image analysis. To the best of our knowledge, this is the first automated framework developed for this purpose.

The framework was trained on a large database, a prerequisite to develop DL tools of good quality. It was also trained on multivendor and clinically heterogeneous data, which makes it generalizable to be implemented as the first step for other existing tools for image analysis. Moreover, the framework was developed through training and testing of 7 state-of the art CNN architectures for each step. In DL, several network variants are available, each exhibiting different strengths and weaknesses. Studies often focus on a single highly individualized network, TABLE 5 | Full pipeline performance.

		Full pipeline	
	BACC	SEN	SPE
Healthy	92.3	96.5	88.1
Pathological	87.3	95.7	78.9
Global	90.6	96.3	85.0

SEN, Sensitivity; SPE, Specificity; and BACC, balanced accuracy of full pipeline, global, and for left (LV) and right ventricle (RV), and healthy and pathological hearts.

tailored for a task through multiple trial-and-error experiments. This makes reproduction of the methods and appreciation of its performance in the context of other datasets challenging. In our work, we present the data of all trained CNN architectures, thus displaying our selection process in a reproducible, fair, and meaningful way.

Finally, we integrated the new framework as the first step of a larger pipeline we had previously developed (8), and we demonstrated that it could produce highly accurate, rapid, and fully-automated cine analysis from a complete collection of images routinely acquired during a clinical study.

#### **Class Identification CNN**

Class identification is the first necessary step for image analysis, making algorithmic classification of standard views a fundamental step for true automatization of analysis (19). DL has been used to meet this need for automated analysis of echo images (20, 21), but not in the field of CMR.

Identification of conventional cine classes can be seen as a trivial task. Nonetheless it is time consuming, especially in long CMR studies, where a multitude of sequences are acquired. Moreover, view recognition cannot rely on the name of the sequences, as these are not replicated across groups and misnaming is common, especially when images are repeated due to insufficient quality or slight errors in view-planning, or added during acquisition. These characteristics make the problem of class identification well-suited for automation. In computer vision tasks in particular, DL has shown excellent performance (22). This is reflected by the high performance of all trained  $\text{CNN}_{\text{class}}$  architectures, which had an accuracy approaching 100%. Precision, recall and F1-scores were mostly between 0.99 and 1 for all classes.

# **Quality Control CNN**

The second DL component of our framework was trained to add a quality-control step to our framework by identifying images of insufficient quality or inadequate planning to inform the automated image analysis process.

Quality control is crucial to transfer DL research tools to the clinical reality in a safe manner, and its importance is increasingly recognized (5, 10, 23, 24). The performance of our CNN<sub>QC</sub> was lower compared to the CNN<sub>class</sub> with highest recorded accuracy of 0.86 for 2Ch and 4Ch, and 0.80 for 3Ch. This is explained by several reasons. First, the input data were highly unbalanced, which is a natural consequence of the fact that radiographers aim to acquire good quality images, resulting in the poor quality class being significantly underrepresented. This is reflected by the significantly lower precision, recall and f1-scores for the identification of "wrong" images compared to that of "correct" ones. To reduce the bias of unbalanced data, we used cross entropy loss, adaptive learning rate scheduler, and balanced accuracy, but such bias can never be fully controlled. Second, images to be considered of insufficient quality have a wide range of problems, from motion artifacts to off-axis planning of different types, making their grouping into one class difficult for the CNN. In particular, when evaluating 3Ch views, the quality of both the cardiac chambers and the aorta were considered, which might explain the lower performance compared to 2Ch and 4Ch views. On the other hand, separation into different classes would have resulted in further imbalance of the data, with insufficient numbers in each hypothetical poor-quality class. Therefore, we decided to group them together. Last, there is a degree of subjectivity in this task, as the same problem can be present to a varying degree of severity; for example, off-axis planning of the 4Ch view can result in the presence of a clear LVOT, or just a small disruption of the basal septum. The subjectivity of QC is reflected in the intra- and inter-observer variability during manual assessment. Consequently, it is both unlikely and unnecessary for any algorithm to reach 100% accuracy in this task. The cases with low degree of severity were the most likely to be misclassified, as well as the most frequent source of interand intra-observer disagreement, as displayed in Figure 2.

#### **Complete Framework**

For the complete framework, we selected the  $\rm CNN_{class}$  and  $\rm CNN_{QC}$  that performed best in combination with a selection algorithm, rather than the networks that performed best in the validation of the individual steps. This was done for two reasons. First, a sequential process can leverage individual strengths and weaknesses to obtain the best combined result. Second, the addition of the selection algorithm after the two CNNs aimed at achieving a more complex and possibly more clinically relevant task: selection of images for further analysis, which can be highly

time-consuming, especially in long exams containing several acquisitions. It was therefore our intent to test the integrated framework and select the one with the best overall performance.

The integration of the three steps of the framework yielded an accurate and rapid system to select images of interest for analysis. The best combination was DenseNet  $\text{CNN}_{\text{class}}$ , ShuffleNet 2Ch-CNN<sub>QC</sub>, DenseNet 2Ch-CNN<sub>QC</sub>, ShuffleNet 4Ch-CNN<sub>QC</sub>, which had a 90% sensitivity for 2Ch, 93% for 3Ch, and 94% for 4Ch acquisitions. This is achieved at the cost of a small proportion of good quality images being mistakenly labeled as erroneous, thus requiring clinician review. However, we believe this is a reasonable compromise to ensure clinical safety within an automated process. Moreover, the process of review is fast in case of data falsely labeled as erroneous, which only requires a visual check from the clinician to accept the analysis results.

#### Full Pipeline: From Scanner to Report

Using the image-processing steps developed in this paper, we were able to present the first pipeline for analysis of cardiac function from cine CMR that automates the complete process from scanner to report, offering an automated system that reproduces manual analysis in current clinical practice. This pipeline is characterized by a high degree of QC (one step in the new framework, two steps in the previously published one). Sequential QC steps focusing on different quality problems ensures a "Swiss cheese" framework, where if a poor-quality image slips through a first barrier, it will likely be flagged up in a later stage. **Supplementary Video 1** displays how the new quality control step aids in selecting images where segmentation can be performed at high standards by subsequent pipeline steps.

Moreover, the addition of the new framework offers automated selection of all standard cine views, which can be further exploited for analysis of parameters beyond conventional ones, such as longitudinal strain and atrioventricular valve systolic excursion, expanding the role of CMR for the assessment of systolic and diastolic function.

The full pipeline is highly accurate, with a focus on high sensitivity, showing an improvement compared to our previously published work (5). The pipeline is also significantly timeefficient, producing outcome measures in about 4 min for standard scans, and in up to 7 min for longer research scans. This is faster than the time reported for the initial pipeline (5), due to changes in the previously developed code.

In the future, we aim to extend our framework to identify and analyze other CMR sequences, including late gadolinium enhancement, flow and T2 mapping. Moreover, with an expanding data set we will be able to train quality control CNNs to recognize specific types of quality and planning errors. In order to decrease subjectivity of this task, a collaborative initiative to build a consensus across a vast number of operators, similar to a recent one developed in the field of echo and AI (25), would be of great value. Lastly, our method now requires postprocessing and is separated from the CMR scanner. In the future, effort should be made for direct implementation on the scanner console. In particular, the implementation of CNN<sub>QC</sub> at the time of image acquisition would aid radiographers by



CCN<sub>QC</sub>; (second line) cases classified as "wrong" both by manual assessment (both operators) and CCN<sub>QC</sub>; (third line) cases classified as "wrong" by manual assessment (with disagreement between operators for 2-chamber) and as "correct" by CCN<sub>QC</sub>. 0, correct; 1, wrong; Ch, chamber; GT, ground truth; R1, first operator; R2, second operator.

promptly detecting images of unsatisfactory quality. This would improve image quality upstream and yield a greater accuracy of downstream image analysis (26).

#### **Study Limitations**

Our dataset did not include CMR studies acquired with General Electrics (GE), thus limiting our framework's generalizability. However, using Philips and Siemens granted a high degree of variability, which would facilitate further training with GE data.

DL algorithms inherently are black boxes. Therefore, interpretation of decisions remains challenging. In this paper we used a stepwise approach of classification and QC algorithms instead of a fused algorithm to allow at least some interpretation of the DL-based decisions.

Moreover, although we included a large number of patients with ACHD to train and test the model, exclusion of grossly

distorted anatomy limits the use of this framework in patients with severe ACHD.

The framework presented in this paper performs limited quality control for short axis. We decided not to train a further CNN for this view as already present in the previously validated pipeline, and it would have therefore been redundant.

# CONCLUSIONS

We developed and validated a framework to select cine acquisitions and perform QC of the selected images prior to automated cine CMR image analysis. We show that our network is able to select cine CMR from a full clinical CMR exam accurately and screen for image quality with a high rate of detecting erroneous acquisitions. We implemented our developed framework as the first step of a wider qualitycontrolled pipeline to obtain automated, quality-controlled analysis of cardiac function from short and long axis cine images from complete CMR clinical studies.

### DATA AVAILABILITY STATEMENT

The datasets presented in this study can be found in online repositories. The names of the repository/repositories and accession number(s) can be found below: The UK Biobank data set is publicly available for approved research projects from https://www.ukbiobank.ac.uk/. The GSTFT data set cannot be made publicly available due to restricted access under hospital ethics and because informed consent from participants did not cover public deposition of data.

# **ETHICS STATEMENT**

The studies involving human participants were reviewed and approved by the North West Research Ethics Committee (REC 11/NW/0382; for UK Biobank data) and the South London Research Ethics Committee (REC 15/NS/0030; for GSTFT data) and was conducted according to the Declaration of Helsinki and Good Clinical Practice guidelines. The patients/participants provided their written informed consent to participate in this study.

# **AUTHOR CONTRIBUTIONS**

VV, EP-A, and BR: conception, study design, and data analysis. EP-A: development of algorithms and analysis software. VV and EP-A: data pre-processing. VV, BR, and RR: clinical advice. VV and BR: manual image annotation. EP-A, BR, and RR: interpretation of data and results. VV, EP-A, BR, and RR: drafting and revising. All authors have read and approved the final manuscript.

#### REFERENCES

- 1. American College of Cardiology Foundation Task Force on Expert Consensus Documents, Hundley WG, Bluemke DA, Finn JP, Flamm SD, Fogel MA, et al. ACCF/ACR/AHA/NASCI/SCMR 2010 expert consensus document on cardiovascular magnetic resonance: a report of the American College of Cardiology Foundation Task Force on Expert Consensus Documents. *J Am Coll Cardiol.* (2010) 55:2614–62. doi: 10.1016/j.jacc.2009.11.011
- Von Knobelsdorff-Brenkenhoff F, Schulz-Menger J. Role of cardiovascular magnetic resonance in the guidelines of the European Society of Cardiology. J Cardiovasc Magn Reson. (2016) 18:6. doi: 10.1186/s12968-016-0225-6
- Leiner T, Bogaert J, Friedrich MG, Mohiaddin R, Muthurangu V, Myerson S, et al. SCMR Position Paper (2020) on clinical indications for cardiovascular magnetic resonance. J Cardiovasc Magn Reson. (2020) 22:76. doi: 10.1186/s12968-020-00682-4
- Bernard O, Lalande A, Zotti C, Cervenansky F, Yang X, Heng P-A, et al. Deep learning techniques for automatic MRI cardiac multi-structures segmentation and diagnosis: is the problem solved? *IEEE Trans Med Imaging*. (2018) 37:2514–25. doi: 10.1109/TMI.2018.2837502
- Ruijsink B, Puyol-Antón E, Oksuz I, Sinclair M, Bai W, Schnabel JA, et al. Fully automated, quality-controlled cardiac analysis from CMR: validation and large-scale application to characterize cardiac function. *JACC Cardiovasc Imaging*. (2020) 13:684–95. doi: 10.1016/j.jcmg.2019.05.030

#### FUNDING

This work was supported by the Wellcome/EPSRC Center for Medical Engineering at Kings College London (WT203148/Z/16/Z). EP-A was supported by the EPSRC (EP/R005516/1 and EP/P001009/1) and by core funding from the Wellcome/EPSRC Center for Medical Engineering (WT203148/Z/16/Z) and BR was supported by the NIHR Cardiovascular MedTech Co-operative award to the Guy's and St. Thomas' NHS Foundation Trust. This research was funded in whole, or in part, by the Wellcome Trust WT203148/Z/16/Z. For the purpose of open access, the author has applied a CC BY public copyright license to any author accepted manuscript version arising from this submission. The authors acknowledge financial support from the Department of Health through the National Institute for Health Research (NIHR) comprehensive Biomedical Research Center award to Guy's and St. Thomas' NHS Foundation Trust in partnership with King's College London.

# ACKNOWLEDGMENTS

This research has been conducted using the UK Biobank Resource under Application Number 17806. The authors wish to thank all participants and staff from UK Biobank and GSTFT.

# SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fcvm. 2021.742640/full#supplementary-material

Supplementary Video 1 | Full pipeline: from scanner to report. It displays the fully automated multi-step QC pipeline from scanner to image analysis and how implementing the new framework prior to segmentation results in good quality analysis.

- Komi S, Inoue Y, Hata H, Nakajima A, Miyatake H. Cardiovascular magnetic resonance evaluation of left ventricular peak filling rate using steady-state free precession and phase contrast sequences. *Springerplus*. (2016) 5:1163. doi: 10.1186/s40064-016-2878-x
- Mendoza D, Codella N, Wang Y, Prince MR, Sethi S, Manoushiagian SJ, et al. Impact of diastolic dysfunction severity on global left ventricular volumetric filling - assessment by automated segmentation of routine cine cardiovascular magnetic resonance. J Cardiovasc Magn Reson. (2010) 12:46. doi: 10.1186/1532-429X-12-S1-P86
- Seemann F, Pahlm U, Steding-Ehrenborg K, Ostenfeld E, Erlinge D. Dubois-Rande JC. Time-resolved tracking of the atrioventricular plane displacement in Cardiovascular Magnetic Resonance (CMR) images. *BMC Med Imaging*. (2017) 17:19. doi: 10.1186/s12880-017-0189-5
- Scannell CM, Veta M, Villa ADM, Sammut EC, Lee J, Breeuwer M, et al. Deeplearning-based preprocessing for quantitative myocardial perfusion MRI. J Magn Reson Imaging. (2020) 51:1689–96. doi: 10.1002/jmri.26983
- Fahmy AS, Rausch J, Neisius U, Chan RH, Maron MS, Appelbaum E, et al. Automated cardiac MR scar quantification in hypertrophic cardiomyopathy using deep convolutional neural networks. *JACC Cardiovasc Imaging*. (2018) 11:1917–8. doi: 10.1016/j.jcmg.2018.04.030
- Puyol-Antón E, Ruijsink B, Baumgartner CF, Masci PG, Sinclair M, Konukoglu E, et al. Automated quantification of myocardial tissue characteristics from native T1 mapping using neural networks with

uncertainty-based quality-control. J Cardiovasc Magn Reson. (2020) 22:60. doi: 10.1186/s12968-020-00650-y

- Krizhevsky A, Sutskever I, Hinton GE. Imagenet classification with deep convolutional neural networks. *Adv Neural Inf Process Syst.* (2012) 25:1097–105.
- Huang G, Liu Z, Weinberger KQ. Densely connected convolutional networks. In: *IEEE Conference on Computer Vision and Pattern Recognition (CVPR)*. Honolulu, HI (2017). p. 4700–8.
- Howard AG, Zhu M, Chen B, Kalenichenko D, Wang W, Weyand T, et al. MobileNets: efficient convolutional neural networks for mobile vision applications. *ArXiv*. (2017) 1704:04861. Available online at: https://arxiv.org/ abs/1704.04861
- He K, Zhang X, Ren S, Sun J. Deep residual learning for image recognition. *ArXiv*. (2015) 1512:03385. doi: 10.1109/CVPR.2016.90
- Zhang X, Zhou X, Lin M, Sun J. ShuffleNet: an extremely efficient convolutional neural network for mobile devices. In: 2018 IEEE/CVF Conference on Computer Vision and Pattern Recognition. Salt Lake City, UT (2018). p. 6848–56. doi: 10.1109/CVPR.2018.00716
- Iandola FN, Han S, Moskewicz MW, Ashraf K, Dally WJ, Keutzer K. SqueezeNet: AlexNet-level accuracy with 50x fewer parameters and <0.5MB model size. *ArXiv*. (2016) 1602:07360. Available online at: https://arxiv.org/ abs/1602.07360
- Simonyan K, Zisserman A. Very deep convolutional networks for large-scale image recognition. ArXiv. (2014) 409:1556. Available online at: https://arxiv. org/abs/1409.1556
- Siegersma KR, Leiner T, Chew DP, Appelman Y, Hofstra L. Artificial intelligence in cardiovascular imaging: state of the art and implications for the imaging cardiologist. *Neth Heart J.* (2019) 27:403–13. doi: 10.1007/s12471-019-01311-1
- Khamis H, Zurakhov G, Azar V, Raz A, Friedman Z, Adam D. Automatic apical view classification of echocardiograms using a discriminative learning dictionary. *Med Image Anal.* (2017) 36:15–21. doi: 10.1016/j.media.2016.10.007
- Madani A, Arnaout R, Mofrad M, Arnaout R. Fast and accurate view classification of echocardiograms using deep learning. *NPJ Digit Med.* (2018) 1:1–8. doi: 10.1038/s41746-017-0013-1
- Voulodimos A, Doulamis N, Doulamis A, Protopapadakis E. Deep learning for computer vision: a brief review. *Comput Intell Neurosci.* (2018) 2018:7068349. doi: 10.1155/2018/7068349

- Robinson R, Valindria VV, Bai W, Oktay O, Kainz B, Suzuki H, et al. Automated quality control in image segmentation: application to the UK biobank cardiovascular magnetic resonance imaging study. J Cardiovasc Magn Reson. (2019) 21:18. doi: 10.1186/s12968-019-0523-x
- 24. Tarroni G, Oktay O, Bai W, Schuh A, Suzuki H, Passerat-Palmbach J, et al. Learning-based quality control for cardiac MR images. *IEEE Trans Med Imaging*. (2019) 38:1127–38. doi: 10.1109/TMI.2018. 2878509
- Howard JP, Stowell CC, Cole GD, Ananthan K, Demetrescu CD, Pearce K, et al. Automated left ventricular dimension assessment using artificial intelligence developed and validated by a UK-wide collaborative. *Circ Cardiovasc Imaging*. (2021) 14:951. doi: 10.1161/CIRCIMAGING.120. 011951
- Chen C, Qin C, Qiu H, Tarroni G, Duan J, Bai W, et al. Deep learning for cardiac image segmentation: a review. *Front Cardiovasc Med.* (2020) 7:25. doi: 10.3389/fcvm.2020.00025

**Author Disclaimer:** The views expressed are those of the authors and not necessarily those of the NHS, the NIHR, EPSRC, or the Department of Health.

**Conflict of Interest:** The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

**Publisher's Note:** All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

Copyright © 2021 Vergani, Razavi, Puyol-Antón and Ruijsink. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.