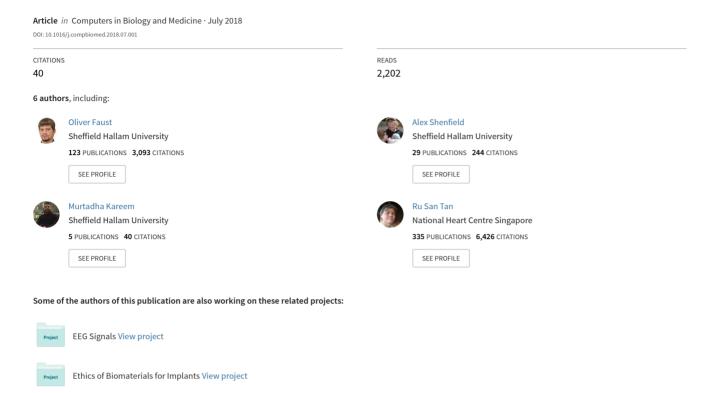
Automated detection of atrial fibrillation using long short-term memory network with RR interval signals



Automated Detection Of Atrial Fibrillation Using Long Short-Term Memory Network With RR Interval Signals

Oliver Faust^{a,1,*}, Alex Shenfield^{a,1}, Murtadha Kareem^a, Tan Ru San^b, Hamido Fujita^c, U Rajendra Acharya^{d,e,f}

^aDepartment of Engineering and Mathematics, Sheffield Hallam University, United Kingdom

^bNational Heart Centre Singapore, Singapore

^cFac. of Software and Information Science, Iwate Prefectural University, Iwate, Japan

^dDepartment of Electronic & Computer Engineering, Ngee Ann Polytechnic, Singapore

^eDepartment of Biomedical Engineering, School of Science and Technology, SUSS, Singapore

^fSchool of Medicine, Faculty of Health and Medical Sciences, Taylor's University, 47500 Subang Jaya, Malaysia

Abstract

Atrial Fibrillation (AF), either permanent or intermittent (paroxysnal AF), increases the risk of cardioembolic stroke. Accurate diagnosis of AF is obligatory for initiation of effective treatment to prevent stroke. Long term cardiac monitoring improves the likelihood of diagnosing paroxysmal AF. We used a deep learning system to detect AF beats in Heart Rate (HR) signals. The data was partitioned with a sliding window of 100 beats. The resulting signal blocks were directly fed into a deep Recurrent Neural Network (RNN) with Long Short-Term Memory (LSTM). The system was validated and tested with data from the MIT-BIH Atrial Fibrillation Database. It achieved 98.51% accuracy with 10-fold cross-validation (20 subjects) and 99.77% with blindfold validation (3 subjects). The proposed system structure is straight forward, because there is no need for information reduction through feature extraction. All the complexity resides in the deep learning system, which gets the entire information from a signal block. This setup leads to the robust performance for unknown data, as measured with the blind fold validation. The proposed Computer-Aided Diagnosis (CAD) system can be used for long-term monitoring of the human heart. To the best of our knowledge, the proposed system is the first to incorporate deep learning for AF beat detection.

Keywords: Deep learning, Recurrent Neural Network, Heart Rate, Atrial Fibrillation

1. Introduction

Atrial Fibrillation (AF) is the most common sustained cardiac rhythm disorder in adults [1]. The disease affects around 0.4% of the adult population and it gets more prevalent with age. Less than 1% of the population under the age of 60 is affected, and over 6% of those over the age of 80 years are affected [2]. It is predicted that the incidence of AF increases, because of the aging population. The disease is associated with inefficiencies in blood flow dynamics which substantially increase the risk of stroke and systemic thromboembolism, resulting in high mortality and morbidity [3, 4]. Some patients with AF are asymptomatic, but others have accompanying symptoms, such as fainting, palpitations, chest pain, fatigue, and heart failure, which seriously diminish the quality of life for the patients [5]. The link between AF and increased stroke risk was established in the Framingham study, which revealed that nonrheumatic AF is an independent stroke risk predictor among 5,070 participants [6]. AF is characterized

^{*}Corresponding author

¹These authors contributed equally to this work.

by highly variable ventricular beat intervals [7]. The distribution of RR intervals during AF differs from the distribution during normal sinus rhythm [8]. In clinical practice, these beat-to-beat variations are detected manually [9]. The sensitivity of this method is above 90% and its specificity is above 70% [10]. Antithrombotic prophylaxis can then be used to reduce the stroke risk from AF [11]. However, a prerequisite for antithrombotic medication is an accurate diagnosis. Unfortunately, antithrombotic medication is associated with bleeding risk and the manual method is neither sensitive nor specific enough to meet the demand for diagnosis, and medication initiation. Furthermore, intra- and inter-observer variability diminish the reliability of manual diagnosis. Automation can potentially improve the sensitivity and specificity [12].

In some patients, the episodes of AF occur paroxysmally, with normal sinus rhythm in between. The timing of occurrence of paroxysmal AF is unpredictable. Both permanent and paroxysmal AF can predispose to increased stroke risk [13]. The challenge for Computer-Aided Diagnosis (CAD) is twofold: How to diagnose AF accurately as well as to detect these AF periods reliably with non-invasive methods that can be used at any time even outside the hospital [14]. Electrical phenomena of the heart muscle is measured by placing passive electrodes on the skin. The electrical activity of the heart activates phasic heart muscle contraction that in turn gives rise to rhythmic cardiac pulse. Electrocardiography is a noninvasive method for measuring that electrical activity [15, 16, 17]. Electrocardiogram (ECG) measurements can be performed at a single seating, or be recorded over an extended period of time, for instance over 24 hours (Holter monitoring). Unfortunately, it has been shown that after catheter ablation procedure for AF, the rate of recurrence of AF episodes is underestimated, and the success rate of catheter ablation is highly overestimated, when determined from 24-h Holter ECG recordings [18]. This issue can be addressed by considerably extending the monitoring period so that the likelihood of detecting AF episodes increases. However, existing techniques for continuous long-term monitoring reduce the patients' quality of life though inconvenient data acquisition methods. The inconvenience can lead to a premature termination of the data acquisition, which adversely affects diagnostic quality [19]. For example, state of the art Holter systems use multiple electrodes, which must be placed by trained technologists. In contrast, beat to beat intervals, which establishes the Heart Rate (HR), once stripped of the morphological component of the ECG signal, can be measured with a single sensor that may be placed by the patient himself. AF detection algorithms are effective for short duration beat interval recordings, offering the prospect of simple and rapid diagnostic tests based on beat intervals alone [20]. CAD systems are mandatory for long-term monitoring, because manual data interpretation of the huge amount data readout is impracticable [21]. For computer based systems, detecting AF periods solely based on HR signals is difficult, because the data is noisy, and the capability of traditional machine learning algorithms is limited [22]. State of the art CAD systems tackle these problems with information reduction through feature extraction, ensuring that the machine learning algorithm is not overwhelmed. The resulting systems under-perform for unknown data and large datasets [23].

We aim to reduce workload of the clinicians and to enable long-term monitoring by providing a robust diagnosis support system for AF. The proposed system structure is minimalist: there is just data partitioning and automated decision support with a deep learning system. That implies the space for design errors is also minimal. Furthermore, the decision-making system gets all the information within the selected data block. There is no information reduction through feature extraction. Consequently, the proposed system is robust and accurate. The AF detection system scored 98.51% accuracy with 10-fold cross-validation. A 99.77% blind fold accuracy indicates a good robustness. These performance figures make us confident that the system will perform well for unknown data in a practical environment, such as AF long-term monitoring. Our work opens up a viable path to extend the monitoring period for diagnosis, treatment monitoring and drug efficacy tests. Furthermore, the results justify our choice to use deep learning for AF detection.

Table 1: Bidirectional Long Short-Term Memory (LSTM) architecture

Layer	Type	Output shape	Number of parameters
1	Input	100, 1	0
2a	LSTM (forward)	100, 400	161600
2b	LSTM (backward)	100, 400	161600
3	Global 1D max pooling	400	0
4	Fully connected Rectified Linear Unit (ReLU)	50	20050
5	Dropout	50	0
6	Fully connected (Sigmoid)	1	51

The remainder of the paper is structured as follows. The next section introduces the materials and methods with a specific focus on the deep learning system. Section 3 states the AF detection results for well-known data. The subsequent discussion section relates our findings to results from other researchers based on the same data. We also state limitations and further work in this section. The conclusion section completes the paper by summarizing the approach and highlighting the main point of the discussion.

2. Materials and methods

This section introduces the data and the processing methods that were used to design the AF detection system. The discussion starts by describing the data and the pre-processing methods. The pre-processed data is directly fed into a deep learning system. As such, that system holds all the computational complexity. Hence, this section focuses on the deep learning algorithm and the design decisions which led to the proposed CAD system.

2.1. Data used

The experiments were conducted based on data from the MIT-BIH Atrial Fibrillation Database [24, 25]. This database includes 23 long-term ECG Holter recordings from different subjects. Each dataset has a duration of 10 hours containing two ECG signals sampled at 250 Hz with AF annotation. These recordings contain also beat annotations and rhythm annotations performed manually by expert clinicians. Furthermore, the R peaks are labelled and the RR interval sequence was extracted based on these labels. The RR interval sequences have been split into overlapping sequences of 100 beats² for each HR trace. A beat sequence is labelled as AF if it contains one or more beats that were classified as showing signs of atrial fibrillation, all other sequences are labelled as normal.

Data from 20 patients has been used for training and 10-fold cross-validation of the model, with the data from the remaining 3 patients completely withheld for use in a blind-fold validation stage after training and validation of the model and tuning of the model hyper-parameters. This step ensures that the proposed method generalises not only to unseen data, but to unseen patients as well.

2.2. Bidirectional Long-Short Term Memory Networks

Deep learning algorithms try to develop the model by using all available information from the input [23]. Extracting this information yields the implicit knowledge which underpins the robust decision making process. Hence, the deep learning approach is more practicable than conventional machine learning, such as Support Vector Machine (SVM) [26].

²99 beats overlap

Recurrent Neural Network (RNN) models have gained increasing popularity in recent years, because they overcome one of the key limitations of using standard machine learning algorithms – the assumption that inputs and outputs to a model are independent of each other [27]. In many problems, such as natural language processing, this assumption is false – for example, to classify sentiment within a sentence it is important to be able to put the individual words of the sentence into context.

RNNs do this by allowing the network to retain and utilise state information (i.e. information about what has happened in previous time steps / inputs). This provides RNNs with a "memory" which captures information about all elements of the input. However, Bengio et al. showed that, whilst standard RNNs can theoretically handle input dependencies over long-intervals, in practice training such networks with gradient descent becomes more inefficient when the temporal span of the input sequence dependencies increases [28]. This results in RNNs becoming difficult to train successfully.

LSTM architecture improves upon standard RNN models by incorporating a gating mechanism which improves the handling of time step information from long-interval input sequences [29]. That mechanism controls the amount of information, from previous time steps, that contributes to the current output. The LSTM gating mechanism implements three layers: 1) input-, 2) forget- and 3) output-layer. The training algorithm determines which information is remembered and indeed which information is forgotten [30]. Much of the current success of RNNs has been achieved using LSTM architecture based models [31, 32].

Schuster and Paliwal proposed the use of bidirectional RNNs for problems where the entire input sequence is available [33]. Bidirectional RNNs utilise past and future data from an input sequence to train both a forward state RNN (operating in the positive time dimension $-t_0, t_1, ..., t_n$) and a backward state RNN (operating in the negative time dimension $-t_n, t_{n-1}, ..., t_0$). This allows the network to make more accurate predictions due to the increased context provided. In recent years, bidirectional LSTM models have shown great promise in fields such as speech recognition, with Graves and Schmidhuber showing that such bidirectional networks can be significantly more effective than unidirectional LSTM architectures [34].

2.3. Proposed system architecture

The details of the proposed bidirectional LSTM model are shown in both Table 1 and Figure 1. The number of LSTM cells in each of the forward and backward layers was set to twice the input sequence length (empirically this has been shown to perform well on a range of natural language and time series classification tasks studied by the authors), with two fully connected layers used as the top model to produce the final classification. Global max pooling in one dimension was used between the bidirectional LSTM layers and the fully connected layers to compress the features of the output sequences produced by the bidirectional LSTM layers.

Effectively the bidirectional LSTM layers act to learn and extract the features from the input HR data sequences, before passing these features to the fully connected top model for classification as to whether signs of AF are present or not. The model proposed in this paper was implemented using Keras and Tensorflow [35, 36, 37].

2.4. Model training

Xavier initialisation [38] was used to initialise all the weights of the model, and gradient descent backpropagation, using the Adam optimiser [39], was used to update the weights. The initial learning rate of the Adam optimiser was set to 1e-3 and the binary cross-entropy function was used to evaluate the loss of the network. A minibatch size of 1024 input sequences was used during this training process – providing a good trade-off between available Graphics Processing Unit (GPU) memory and speed of training.

Recurrent dropout [40] was applied (with a probability of 0.1) during training to both the inputs and hidden states of the LSTM cells and standard dropout [41] was applied between the

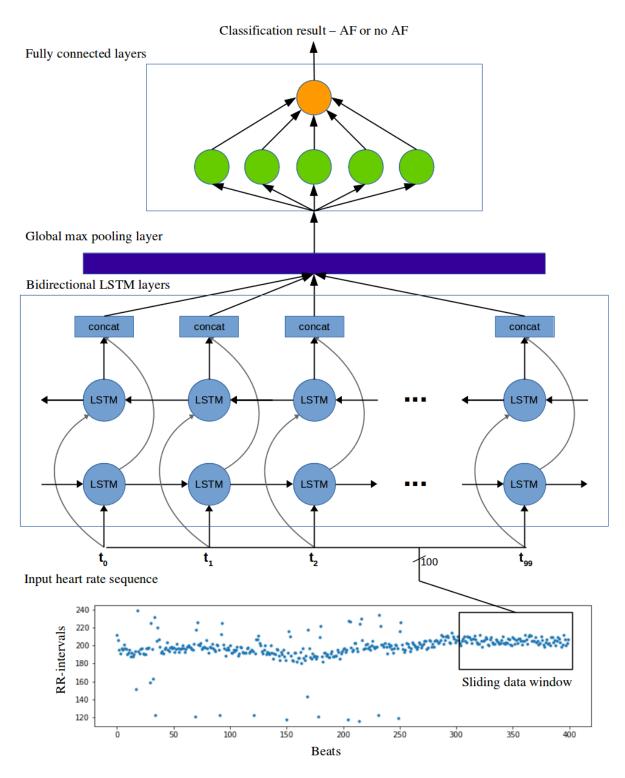


Figure 1: The bidirectional LSTM architecture used for AF classification $\,$

fully connected layers (again, with a probability of 0.1) to reduce overfitting of the model and improve model generalisation. Furthermore, the training performance of the proposed model was evaluated using the binary cross-entropy function as this provides a better understanding of the model performance across a range of operating conditions (as opposed to the classification accuracy which only describes the performance of a model at a single point). The binary cross-entropy function compares two probability distributions (that of the true distribution and the predicted distribution) to provide more information about the nature of the search landscape.

A stratified 10-fold cross-validation strategy was used to evaluate the model performance and to tune both the model architecture and the hyperparameters (although minimal tuning was used to obtain the results shown in the next section). Stratified cross-validation was necessary to ensure that each fold was representative of the balance of the full data set. Results from this stratified 10-fold cross-validation are shown in Section 3.1.

3. Results

This section introduces the 10-fold cross-validation results for training and testing as well as the blind-fold validation results. The bi-directional LSTM model, proposed in this paper, was trained using an nVidia Quadro M5000 GPU with 8GB of GDDR5 graphics RAM. The average time needed to train a single epoch of this model was approximately 215 s. Initial experiments showed that the model converges after between 60-80 epochs of training, so 80 epochs were used in this study to ensure full convergence of the model and limit opportunities for overfitting.

3.1. 10-fold cross-validation results

Figures 2 and 3 show the training and validation set performance against the number of epochs³. These figures show the mean of the performance for each of the 10-folds as a solid line, and the standard deviation of the performance as the shaded region. As can be seen from these figures, whilst the performance on the training set is slightly better than that of the validation set, the model has converged to a stable value with none of the typical signs of overfitting such as the training performance continuing to improve, whilst validation performance stagnates (and even worsens).

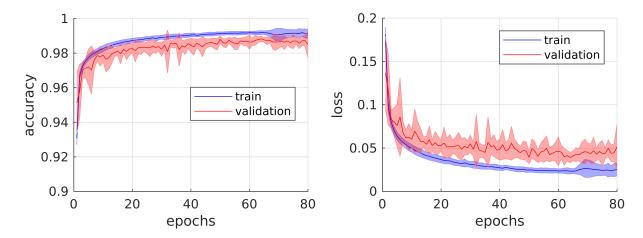


Figure 2: Training and validation accuracy over epochs. The solid line is the mean of the 10 folds and the shaded area indicates the variance.

Figure 3: Training and testing loss function over the epochs. The solid line is the mean and the shaded area indicates the variance.

³One epoch is a single pass through all the training data, followed by evaluation of the model on the validation data.

Table 2: Overall cross validation performance of the proposed bi-directional LSTM classifier

TP	TN	FP	FN	Accuracy	Sensitivity	Specificity	Area Under Curve (AUC)
430,615	523,241	7,040	7,407	98.51%	98.32%	98.67%	0.9986

Table 3: Overall blind fold validation performance of the proposed bi-directional LSTM classifier

TP	TN	FP	FN	Accuracy	Sensitivity	Specificity	AUC
91,888	65.699	255	116	99.77%	99.87%	99.61%	1^4

The confusion plot for the stratified 10-fold cross-validation process is shown in Figure 4 and the Receiver Operating Characteristic (ROC) curve is shown in Figure 5. In these plots, the results from applying the model to the validation set in all 10 folds of the cross-validation process were aggregated and then used to create the confusion plot and the ROC curve. Table 2 shows the overall performance of the cross-validation process.

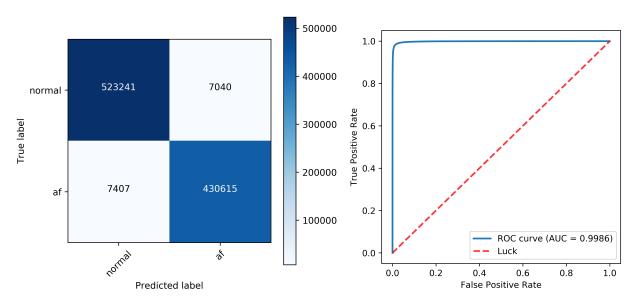


Figure 4: Confusion plot from the stratified 10-fold cross-validation process

Figure 5: ROC curve from the stratified 10-fold cross-validation process

It can been seen from the results in Figures 4 and 5 and Table 2, that the proposed bidirectional LSTM classifier achieves an overall classification accuracy of 98.51% with stratified 10-fold cross-validation - classifying 98.67% of normal HR sequences correctly and 98.32% of HR sequences showing signs of atrial fibrillation correctly.

3.2. Blind-fold validation results

Following this stratified 10-fold cross-validation process, further evaluation of the proposed model was undertaken using blindfold validation of AF and normal HR sequences from 3 completely held out patients, as described in Section 2.1. Results from this completely held out test set are shown in Figures 6 and 7, and summarised in Table 3.

It can been seen from the results in Figures 6 and 7 and Table 3, that the proposed bidirectional LSTM classifier achieves an overall classification accuracy of 99.77% on the test set of completely unseen patients - classifying 99.61% of normal HR sequences correctly and 99.87% of HR sequences showing signs of atrial fibrillation correctly.

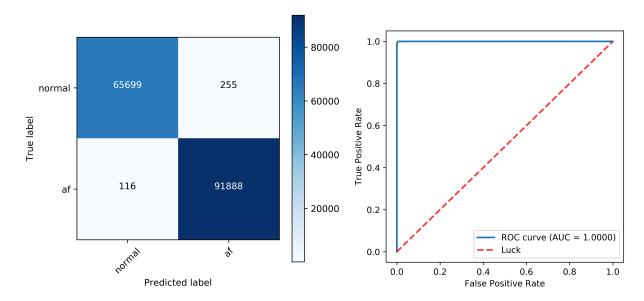


Figure 6: Confusion plot from the 3 completely withheld Figure 7: ROC curve from the 3 completely witpatients

hheld patients

4. Discussion

Data is always a concern for learning systems. Indeed, the proposed automated detection system for AF is no exception. We have used the physionet MIT-BIH AF database, because this enables competition and cooperation. Competition comes from the fact that the database is well known and classification results are available from other research projects, see Table 4. Cooperation is possible, because the data is freely available and our processing methods were stated in Section 2. However, the data is balanced for sinus rhythm and AF. That implies that it is heavily weighted with AF patients compared to a real patient population. That is a problem, especially for AF screening. For screening applications, we expect that our performance figures are biased toward high sensitivity and false positives for lower prevalence of the abnormality.

Table 4 shows a selection of scientific studies which are based on heart rate from the physionet MIT-BIH AF database [21]. All these studies have achieved good validation results with a conventional processing structure of feature extraction followed by machine learning. The processing structure we propose does not need feature extraction, all the information available in the training data set is passed on to the deep learning system. Hence, it can extract the implicit knowledge which enables it to make good decisions even for unknown data.

Looking beyond the narrow area of AF detection with Deep Learning (DL) applied to HR, we found only one similar work. Chen et al. who used deep learning to detect Congestive Heart Failure (CHF) beats with an accuracy of 72.41% [42]. Their system incorporated the Sparse-Auto-Encoder (SAE) algorithm which increases the robustness of the model. Apart from HR, there are other physiological signals that can be used to diagnose Cardiovascular Diseases (CVDs). Deep learning was used successfully to analyze a wide range of these physiological signals [23]. Sannino De Pietro achieved an accuracy of 99.09% for AF detection with tensorfolow deep learning on the ECG signals in the Physionet AF database [43]. Zihlmann et al. have used a convolutional RNN for ECG classification [44]. Their system achieved and accuracy of 80.5%. Deep Convolutional Neural Network (CNN) was used extensively for ECG analysis [45, 46, 47, 48, 49]. Deep Artificial Neural Network (ANN) have be used for multiple heath care applications [50, 51, 52].

Table 4: Selected work on the automated detection of AF beats using RR interval signals from the MIT-BIH Atrial Fibrillation database.

Author	Data pre-processing	Feature extraction met-	Analysis method	Results
		hod		
Artis et al., 1991	100 beat window 99 be-	Threshold based inter-	ANN	92.86% sensitivity and $92.34%$
[53]	ats overlap	vals		positive predictive accuracy
Tateno and Glass,	50 consecutive AF be-	ΔRR is defined as being	Kolmogorov-Smirnov	Sensitivity 94.4%, Specificity
2001 [54]	ats, no overlap	the difference between	test and CV test	97.2% both with Kolmogorov-
		two successive RR inter-		Smimov statistics.
		vals.		
Logan and Hea-	600 consecutive beats,	ΔRR and RR interval	Kolmogorov-Smirnov	sensitivity = 96% and specifi-
ley, 2005 [55]	no overlap	histogram	test	city = 89%.
Ghodrati et al.,	30 beat window no over-	Normalized absolute de-	Threshold	sensitivity of 89%
2008 [56]	lap reported	viation and normalized		
		absolute difference		
Ghodrati and Ma-	30 beat window no over-	Probability density	Threshold evaluated	sensitivity of 92%
rinello, 2008 [57]	lap reported	function	with ROC	
Babaeizadeh et	Beat to beat analysis	RR interval	Hidden Markov model	92% sensitivity and 97% posi-
al., 2009 [58]				tive predictive value
Yaghouby et al.,	64 beat window, overlap	Statistical and geo-	Genetic algorithm	99.11% Accuracy
2010 [59]	not reported	metrical features plus		
		feature dimension		
		reduction		
Lian et al., 2011	128 beat window	ΔRR	Threshold on statistical	95.9% sensitivity and $95.4%$
[60]			values	positive predictive value
Huang et al., 2011	bins of 50 beats	ΔRR and RR interval	Kolmogorov-Smirnov	specificity = 96.1% .
[61]		histogram	test	
Yaghouby et al.,	30 beat window no over-	Six morphological fea-	Gaussian mixture mo-	sensitivity of 98.09%, specifi-
2012 [5]	lap reported	tures	del	city=91.66%
Rincón et al.,	600 beat window no	Hybrid RR interval and	Fuzzy classifier	sensitivity of 96%, specifi-
2012 [62]	overlap reported	ECG features		city=93%

Zhou et al., 2014	Median filter	Shannon entropy	Threshold evaluated	sensitivity 96.72%, specificity
[63]			with ROC	95.07%, accuracy 96.05%
Petrėnas et al.,	8 beat sliding window	Median filter and thres-	Threshold	sensitivity of 97.1%, specifi-
2015 [64]		holding for data label-		city of 98.3%
		ling		
Henzel et al., 2017	beat by beat evalua-	4 statistical features	Generalized Linear Mo-	accuracy 93%, sensitivity of
[65]	tion. Windows of va-	and the beat itself	del evaluated with ROC	90%, specificity $95%$
	rying length are used			
	to extract the statistical			
	features.			
Proposed sy-	100 beat window 99	None	Recurrent neural	Cross validation: accu-
Proposed system	100 beat window 99 beats overlap	None	Recurrent neural network	Cross validation: accuracy 98.51%, Sensiti-
		None		
		None		racy 98.51%, Sensiti-
		None		racy 98.51%, Sensitivity 98.32%, Specificity
		None		racy 98.51%, Sensitivity 98.32%, Specificity 98.67%, Positive pre-
		None		racy 98.51%, Sensitivity 98.32%, Specificity 98.67%, Positive predictive accuracy 98.39%.
		None		racy 98.51%, Sensitivity 98.32%, Specificity 98.67%, Positive predictive accuracy 98.39%. Blind fold validation:
		None		racy 98.51%, Sensitivity 98.32%, Specificity 98.67%, Positive predictive accuracy 98.39%. Blind fold validation: accuracy 99.77%, Sensi-

4.1. Limitations

A limitation of our study comes from the fact that we have used data blocks with 100 RR intervals. These short segments might not capture all the nonlinear properties of the signal adequately and therefore the deep learning system does not get all the information. However, the choice of a 100 beat window was made with regard to a practical real-time diagnosis system. This 100 beat window would allow for minimal latency in any automated alert of AF detection via heart rate monitoring. Another shortcoming is the limited number of subjects. More data from a larger subject range would lead to the extraction of more implicit knowledge. In turn that might result in a model that is even more robust. A problem we encountered was the fact that it takes design time to train the model. During the design phase, training speed is major limitation, because it prevented us to explore all possible network parameters. Even massively parallel⁵ GPU processors could not speed up the processing sufficiently.

Another limitation of our study comes from instrumentation. We have used RR intervals that were extracted from ECG signals. However, to realize the advantages of HR the signal has to be measured directly, i.e. not being extracted from ECG. Appropriate instrumentation has to ensure that these measurements reflect the RR intervals in the same way as if they were extracted from ECG. In future, we plan to investigate the impact of the heart rate sensor.

4.2. Future work

The future work is centered on the long-term monitoring of the human heart [66]. Internet of Things (IoT) technology is needed to facilitate data transfer and disseminate control messages [67]. Apart from the technology aspects, there is also much scope to acquire more depth and breadth through academic research. More depth is achieved by detecting different AF types and predict the onset of AF. Having cost effective long-term monitoring systems allows us to extend the breath of research by investigating a range of different heart diseases, such as CHF [68]. Apart from addressing these medical engineering aspects, further work is planned to conduct a full ablation study to investigate the importance of individual architectural and algorithmic components of the proposed model (such as the number of LSTM cells, optimisers, learning rates, and fully connected classification layer sizes).

5. Conclusion

Our approach to AF detection in HR signals is straight forward: we partition the data with a sliding window and feed the resulting blocks into a deep learning system. There is no need for information reduction through feature extraction. During the training phase, the learning algorithm can extract all the available information to create the implicit knowledge which underpins the subsequent decision-making processes. As a consequence, the proposed decision support system delivers accurate and robust results. To be specific, the system achieved an accuracy of 98.51% with 10-fold cross-validation. The 99.77% blind fold accuracy indicates a good robustness. These results are similar to performance measures that were reported for ECG based CAD systems. However, HR measurements are more convenient for the patient and they can be carried out for longer.

Having an accurate and robust AF detection system using RR intervals is a prerequisite for long-term monitoring. The goal for such long-term monitoring is to produce more data which holds more information about the patient health. Deep learning can help us to make sense of that data and thereby reduce the workload of clinicians. Furthermore, such computer aided diagnosis reduces the risk of inter- and intra-observer variability. Computer based systems do not suffer from fatigue and the results are reproducible. Furthermore, the learning algorithm

⁵The term 'massively parallel' refers to the use of a large number of processors

model can be updated, such that the decision support system improves over time. These systems have the potential to benefit patients by delivering an accurate diagnosis as well as unintrusive and uninterrupted treatment monitoring. We hope that this work is the first in a series of scientific studies on deep learning for AF detection.

Acknowledgements

Funding: We gratefully acknowledge the support of NVIDIA Corporation with the donation of the Quadro M5000 GPU used in this research. No external funding sources supported this study.

Competing interests: The authors declare that they have no conflict of interest.

Statements of ethical approval: All the data for this study comes from the well known MIT-BIH Atrial Fibrillation Database. The authors did not undertake measurements that involved human or animal participants.

Acronyms

AF Atrial Fibrilla	ation
---------------------------	-------

ANN Artificial Neural Network

AUC Area Under Curve

CAD Computer-Aided DiagnosisCHF Congestive Heart Failure

CNN Convolutional Neural Network

CVD Cardiovascular Disease

DL Deep LearningECG Electrocardiogram

GPU Graphics Processing Unit

HR Heart Rate

IoT Internet of Things

ReLU Rectified Linear Unit
RNN Recurrent Neural Network

ROC Receiver Operating Characteristic

SAE Sparse-Auto-EncoderSVM Support Vector Machine

References

- [1] S. T. Mathew, J. Patel, S. Joseph, Atrial fibrillation: mechanistic insights and treatment options, European journal of internal medicine 20 (2009) 672–681.
- [2] S. Petrutiu, J. Ng, G. M. Nijm, H. Al-Angari, S. Swiryn, A. V. Sahakian, Atrial fibrillation and waveform characterization, IEEE engineering in medicine and biology magazine 25 (2006) 24–30.
- [3] E. J. Benjamin, P. A. Wolf, R. B. D'Agostino, H. Silbershatz, W. B. Kannel, D. Levy, Impact of atrial fibrillation on the risk of death: the framingham heart study, Circulation 98 (1998) 946–952.
- [4] G. Y. Lip, R. Nieuwlaat, R. Pisters, D. A. Lane, H. J. Crijns, Refining clinical risk stratification for predicting stroke and thromboembolism in atrial fibrillation using a novel risk factor-based approach: the euro heart survey on atrial fibrillation, Chest 137 (2010) 263–272.

- [5] F. Yaghouby, A. Ayatollahi, R. Bahramali, M. Yaghouby, Robust genetic programming-based detection of atrial fibrillation using rr intervals, Expert Systems 29 (2012) 183–199.
- [6] P. A. Wolf, R. D. Abbott, W. B. Kannel, Atrial fibrillation as an independent risk factor for stroke: the framingham study., Stroke 22 (1991) 983–988.
- [7] P.-S. Yang, S. Ryu, D. Kim, E. Jang, H. T. Yu, T.-H. Kim, J. Hwang, B. Joung, G. Y. Lip, Variations of prevalence and incidence of atrial fibrillation and oral anticoagulation rate according to different analysis approaches, Scientific Reports 8 (2018) 6856.
- [8] T. Hennig, P. Maass, J. Hayano, S. Heinrichs, Exponential distribution of long heart beat intervals during atrial fibrillation and their relevance for white noise behaviour in power spectrum, Journal of biological physics 32 (2006) 383–392.
- [9] S. Somerville, J. Somerville, P. Croft, M. Lewis, Atrial fibrillation: a comparison of methods to identify cases in general practice., Br J Gen Pract 50 (2000) 727–729.
- [10] N. Lowres, L. Neubeck, J. Redfern, S. B. Freedman, Screening to identify unknown atrial fibrillation, Thrombosis and haemostasis 110 (2013) 213–222.
- [11] R. G. Hart, L. A. Pearce, M. I. Aguilar, Meta-analysis: antithrombotic therapy to prevent stroke in patients who have nonvalvular atrial fibrillation, Annals of internal medicine 146 (2007) 857–867.
- [12] O. Faust, U. R. Acharya, T. Tamura, Formal design methods for reliable computer-aided diagnosis: a review, IEEE reviews in biomedical engineering 5 (2012) 15–28.
- [13] O. Faust, R. Acharya, S. Krishnan, L. C. Min, Analysis of cardiac signals using spatial filling index and time-frequency domain, BioMedical Engineering OnLine 3 (2004) 30.
- [14] R. J. Martis, U. R. Acharya, H. Adeli, H. Prasad, J. H. Tan, K. C. Chua, C. L. Too, S. W. J. Yeo, L. Tong, Computer aided diagnosis of atrial arrhythmia using dimensionality reduction methods on transform domain representation, Biomedical signal processing and control 13 (2014) 295–305.
- [15] R. J. Martis, U. R. Acharya, L. C. Min, Ecg beat classification using pca, lda, ica and discrete wavelet transform, Biomedical Signal Processing and Control 8 (2013) 437–448.
- [16] M. Kumar, R. B. Pachori, U. R. Acharya, Automated diagnosis of atrial fibrillation ecg signals using entropy features extracted from flexible analytic wavelet transform, Biocybernetics and Biomedical Engineering (2018) 1–10.
- [17] J. Ródenas, M. García, R. Alcaraz, J. J. Rieta, Wavelet entropy automatically detects episodes of atrial fibrillation from single-lead electrocardiograms, Entropy 17 (2015) 6179– 6199.
- [18] T. Hanke, E. I. Charitos, U. Stierle, A. Karluss, E. Kraatz, B. Graf, A. Hagemann, M. Misfeld, H. H. Sievers, Twenty-four-hour holter monitor follow-up does not provide accurate heart rhythm status after surgical atrial fibrillation ablation therapy: up to 12 months experience with a novel permanently implantable heart rhythm monitor device, Circulation 120 (2009) S177–S184.
- [19] A. F. Members, V. Fuster, L. E. Rydén, D. S. Cannom, H. J. Crijns, A. B. Curtis, K. A. Ellenbogen, J. L. Halperin, J.-Y. Le Heuzey, G. N. Kay, et al., Acc/aha/esc 2006 guidelines for the management of patients with atrial fibrillation—executive summary: A report of the

- american college of cardiology/american heart association task force on practice guidelines and the european society of cardiology committee for practice guidelines (writing committee to revise the 2001 guidelines for the management of patients with atrial fibrillation) developed in collaboration with the european heart rhythm association and the heart rhythm society, European Heart Journal 27 (2006) 1979–2030.
- [20] P. Langley, M. Dewhurst, L. Di Marco, P. Adams, F. Dewhurst, J. Mwita, R. Walker, A. Murray, Accuracy of algorithms for detection of atrial fibrillation from short duration beat interval recordings, Medical Engineering and Physics 34 (2012) 1441–1447.
- [21] N. Larburu, T. Lopetegi, I. Romero, Comparative study of algorithms for atrial fibrillation detection, in: Computing in Cardiology, 2011, IEEE, pp. 265–268.
- [22] K. Tateno, L. Glass, A method for detection of atrial fibrillation using rr intervals, in: Computers in Cardiology 2000, IEEE, pp. 391–394.
- [23] O. Faust, Y. Hagiwara, T. J. Hong, O. S. Lih, U. R. Acharya, Deep learning for healthcare applications based on physiological signals: a review, Computer Methods and Programs in Biomedicine 161 (2018) 1–13.
- [24] G. Moody, A new method for detecting atrial fibrillation using rr intervals, Computers in Cardiology (1983) 227–230.
- [25] A. L. Goldberger, L. A. Amaral, L. Glass, J. M. Hausdorff, P. C. Ivanov, R. G. Mark, J. E. Mietus, G. B. Moody, C.-K. Peng, H. E. Stanley, Physiobank, physiotoolkit, and physionet, Circulation 101 (2000) e215–e220.
- [26] O. Faust, Documenting and predicting topic changes in computers in biology and medicine: A bibliometric keyword analysis from 1990 to 2017, Informatics in Medicine Unlocked 11 (2018) 15–27.
- [27] J. Schmidhuber, Deep learning in neural networks: An overview, Neural networks 61 (2015) 85–117.
- [28] Y. Bengio, P. Simard, P. Frasconi, Learning long-term dependencies with gradient descent is difficult, IEEE transactions on neural networks 5 (1994) 157–166.
- [29] S. Hochreiter, J. Schmidhuber, Long short-term memory, Neural computation 9 (1997) 1735–1780.
- [30] J. H. Tan, Y. Hagiwara, W. Pang, I. Lim, S. L. Oh, M. Adam, R. San Tan, M. Chen, U. R. Acharya, Application of stacked convolutional and long short-term memory network for accurate identification of cad ecg signals, Computers in Biology and Medicine (2018).
- [31] R. Pascanu, C. Gulcehre, K. Cho, Y. Bengio, How to construct deep recurrent neural networks, arXiv preprint arXiv:1312.6026 (2013).
- [32] B. A. Pearlmutter, Learning state space trajectories in recurrent neural networks, Neural Computation 1 (1989) 263–269.
- [33] M. Schuster, K. K. Paliwal, Bidirectional recurrent neural networks, IEEE Transactions on Signal Processing 45 (1997) 2673–2681.
- [34] A. Graves, J. Schmidhuber, Framewise phoneme classification with bidirectional lstm and other neural network architectures, Neural Networks 18 (2005) 602–610.

- [35] F. Chollet, et al., Keras: Deep learning library for theano and tensorflow, URL: https://keras. io/k 7 (2015) 8.
- [36] A. Gulli, S. Pal, Deep Learning with Keras, Packt Publishing Ltd, 2017.
- [37] M. Abadi, P. Barham, J. Chen, Z. Chen, A. Davis, J. Dean, M. Devin, S. Ghemawat, G. Irving, M. Isard, et al., Tensorflow: A system for large-scale machine learning., in: OSDI, volume 16, pp. 265–283.
- [38] X. Glorot, Y. Bengio, Understanding the difficulty of training deep feedforward neural networks, in: Proceedings of the thirteenth international conference on artificial intelligence and statistics, pp. 249–256.
- [39] A. Radford, L. Metz, S. Chintala, Unsupervised representation learning with deep convolutional generative adversarial networks, arXiv preprint arXiv:1511.06434 (2015).
- [40] Y. Gal, Z. Ghahramani, Bayesian convolutional neural networks with bernoulli approximate variational inference, arXiv preprint arXiv:1506.02158 (2015).
- [41] N. Srivastava, G. Hinton, A. Krizhevsky, I. Sutskever, R. Salakhutdinov, Dropout: A simple way to prevent neural networks from overfitting, The Journal of Machine Learning Research 15 (2014) 1929–1958.
- [42] W. Chen, G. Liu, S. Su, Q. Jiang, H. Nguyen, A chf detection method based on deep learning with rr intervals, in: Engineering in Medicine and Biology Society (EMBC), 2017 39th Annual International Conference of the IEEE, IEEE, pp. 3369–3372.
- [43] G. Sannino, G. De Pietro, A deep learning approach for ecg-based heartbeat classification for arrhythmia detection, Future Generation Computer Systems (2018) Article in press.
- [44] M. Zihlmann, D. Perekrestenko, M. Tschannen, Convolutional recurrent neural networks for electrocardiogram classification, arXiv preprint arXiv:1710.06122 (2017).
- [45] U. R. Acharya, S. L. Oh, Y. Hagiwara, J. H. Tan, M. Adam, A. Gertych, R. San Tan, A deep convolutional neural network model to classify heartbeats, Computers in biology and medicine 89 (2017) 389–396.
- [46] U. R. Acharya, H. Fujita, S. L. Oh, Y. Hagiwara, J. H. Tan, M. Adam, Application of deep convolutional neural network for automated detection of myocardial infarction using ecg signals, Information Sciences 415 (2017) 190–198.
- [47] U. R. Acharya, H. Fujita, O. S. Lih, Y. Hagiwara, J. H. Tan, M. Adam, Automated detection of arrhythmias using different intervals of tachycardia ecg segments with convolutional neural network, Information sciences 405 (2017) 81–90.
- [48] U. R. Acharya, H. Fujita, O. S. Lih, M. Adam, J. H. Tan, C. K. Chua, Automated detection of coronary artery disease using different durations of ecg segments with convolutional neural network, Knowledge-Based Systems 132 (2017) 62–71.
- [49] U. R. Acharya, H. Fujita, S. L. Oh, Y. Hagiwara, J. H. Tan, M. Adam, R. S. Tan, Deep convolutional neural network for the automated diagnosis of congestive heart failure using ecg signals, Applied Intelligence (2018) 1–12.
- [50] A. Shenfield, S. Rostami, Multi-objective evolution of artificial neural networks in multiclass medical diagnosis problems with class imbalance, in: Computational Intelligence in Bioinformatics and Computational Biology (CIBCB), 2017 IEEE Conference on, IEEE, pp. 1–8.

- [51] A. Shenfield, M. Rodrigues, H. Nooreldeen, J. Moreno-Cuesta, A novel hybrid differential evolution strategy applied to classifier design for mortality prediction in adult critical care admissions, in: Computational Intelligence in Bioinformatics and Computational Biology (CIBCB), 2017 IEEE Conference on, IEEE, pp. 1–7.
- [52] A. Shenfield, S. Rostami, A multi objective approach to evolving artificial neural networks for coronary heart disease classification, in: Computational Intelligence in Bioinformatics and Computational Biology (CIBCB), 2015 IEEE Conference on, IEEE, pp. 1–8.
- [53] S. G. Artis, R. Mark, G. Moody, Detection of atrial fibrillation using artificial neural networks, in: Computers in Cardiology 1991, Proceedings., IEEE, pp. 173–176.
- [54] K. Tateno, L. Glass, Automatic detection of atrial fibrillation using the coefficient of variation and density histograms of rr and δ rr intervals, Medical and Biological Engineering and Computing 39 (2001) 664–671.
- [55] B. Logan, J. Healey, Robust detection of atrial fibrillation for a long term telemonitoring system, in: Computers in Cardiology, 2005, IEEE, pp. 619–622.
- [56] A. Ghodrati, B. Murray, S. Marinello, Rr interval analysis for detection of atrial fibrillation in ecg monitors., in: Conference proceedings:... Annual International Conference of the IEEE Engineering in Medicine and Biology Society. IEEE Engineering in Medicine and Biology Society. Annual Conference, volume 2008, p. 601.
- [57] A. Ghodrati, S. Marinello, Statistical analysis of rr interval irregularities for detection of atrial fibrillation, in: Computers in Cardiology, 2008, IEEE, pp. 1057–1060.
- [58] S. Babaeizadeh, R. E. Gregg, E. D. Helfenbein, J. M. Lindauer, S. H. Zhou, Improvements in atrial fibrillation detection for real-time monitoring, Journal of electrocardiology 42 (2009) 522–526.
- [59] F. Yaghouby, A. Ayatollahi, R. Bahramali, M. Yaghouby, A. H. Alavi, Towards automatic detection of atrial fibrillation: A hybrid computational approach, Computers in Biology and Medicine 40 (2010) 919–930.
- [60] J. Lian, L. Wang, D. Muessig, A simple method to detect atrial fibrillation using rr intervals, American journal of cardiology 107 (2011) 1494–1497.
- [61] C. Huang, S. Ye, H. Chen, D. Li, F. He, Y. Tu, A novel method for detection of the transition between atrial fibrillation and sinus rhythm, IEEE Transactions on Biomedical Engineering 58 (2011) 1113–1119.
- [62] F. Rincón, P. R. Grassi, N. Khaled, D. Atienza, D. Sciuto, Automated real-time atrial fibrillation detection on a wearable wireless sensor platform, in: Engineering in Medicine and Biology Society (EMBC), 2012 Annual International Conference of the IEEE, IEEE, pp. 2472–2475.
- [63] X. Zhou, H. Ding, B. Ung, E. Pickwell-MacPherson, Y. Zhang, Automatic online detection of atrial fibrillation based on symbolic dynamics and shannon entropy, Biomedical engineering online 13 (2014) 18.
- [64] A. Petrėnas, V. Marozas, L. Sörnmo, Low-complexity detection of atrial fibrillation in continuous long-term monitoring, Computers in biology and medicine 65 (2015) 184–191.

- [65] N. Henzel, J. Wróbel, K. Horoba, Atrial fibrillation episodes detection based on classification of heart rate derived features, in: Mixed Design of Integrated Circuits and Systems, 2017 MIXDES-24th International Conference, IEEE, pp. 571–576.
- [66] U. R. Acharya, O. Faust, D. N. Ghista, S. V. Sree, A. P. C. Alvin, S. Chattopadhyay, T.-C. Lim, E. Y.-K. Ng, W. Yu, A systems approach to cardiac health diagnosis, Journal of Medical Imaging and Health Informatics 3 (2013) 261–267.
- [67] M. S. Hossain, G. Muhammad, Cloud-assisted industrial internet of things (iiot)—enabled framework for health monitoring, Computer Networks 101 (2016) 192–202.
- [68] U. R. Acharya, O. Faust, V. Sree, G. Swapna, R. J. Martis, N. A. Kadri, J. S. Suri, Linear and nonlinear analysis of normal and cad-affected heart rate signals, Computer methods and programs in biomedicine 113 (2014) 55–68.