Learning disentangled representation from 12-lead electrograms: application in localizing the origin of Ventricular Tachycardia

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Abstract

The increasing availability of electrocardiogram (ECG) data has motivated the use of data-driven models for automating various clinical tasks based on ECG data. The development of subject-specific models are limited by the cost and difficulty of obtaining sufficient training data for each individual. The alternative of population model, however, faces challenges caused by the significant inter-subject variations within the ECG data. We address this challenge by investigating for the first time the problem of learning representations for clinically-informative variables while disentangling other factors of variations within the ECG data. In this work, we present a conditional variational autoencoder (VAE) to extract the subject-specific adjustment to the ECG data, conditioned on task-specific representations learned from a deterministic encoder. To encourage the representation for intersubject variations to be independent from the task-specific representation, maximum mean discrepancy is used to match all the moments between the distributions learned by the VAE conditioning on the code from the deterministic encoder. The learning of the task-specific representation is regularized by a weak supervision in the form of contrastive regularization. We apply the proposed method to a novel yet important clinical task of classifying the origin of ventricular tachycardia (VT) into pre-defined segments, demonstrating the efficacy of the proposed method against the standard VAE.

Introduction

The increasing availability of electrocardiogram (ECG) data has motivated the use of data-driven models for automating various clinical tasks based on ECG data (Rajpurkar et al. 2017). However, ECG signals exhibit significant physiological variations across individuals (Park, Kim, and Marchlinski 2012), (Rajpurkar et al. 2017), (Wiens and Guttag 2010), a manifestation of anatomical differences in the heart and torso among individuals along with electrode positioning on the body surface during ECG acquisition (Plonsey and Fleming 1969). There are also various pathological confounders that are tied to the specific clinical task under consideration, such as sub-groups under a population group, or different characteristics of structural abnormality of the heart. These inter-subject variations are not always observed and they

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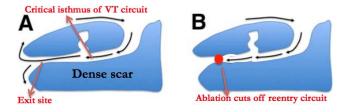


Figure 1: Schematics of a VT reentry circuit. A: an electrical "short circuit" travels through narrow strands of surviving tissue inside the scar and exits from the scar to depolarize the rest of the ventricle. B: Cutting off the isthmus or exits that form the "short circuit" in catheter ablation.

present a great challenge for building accurate populationbased models for patient-specific uses. At the same time, the alternative of patient-specific models are difficult to implement in clinical practice due to the challenge and cost in obtaining sufficient data from each patient.

One particular example is the clinical task of localizing the origin of ventricular tachycardia (VT) from an ECG signal. VT is a significant cause of sudden cardiac deaths in the United States (Stevenson 2009). Such VT usually involves an electrical "short circuit" formed by narrow strands of surviving tissue inside the myocardial scar as illustrated in Fig. 1A. Clinically, this is treated by cutting off the circuit at its exit (VT exit) from the scar using catheter ablation (Stevenson 2009) as shown in Fig. 1B. Because a VT exit acts like the origin of ventricular activation to be seen on ECG, these exit sites are currently located by repeatedly stimulating (i.e. pacing) different locations of the heart until localizing the site where pacing reproduce the QRS morphology of the VT on all 12 leads of the ECG (Stevenson 2009).

Rather than this "trial-and-error" status quo, naturally, an automatic real-time identification of the VT exit from 12-lead ECG data would improve the efficiency of ablation procedure. Limited attempts by the use of methods such as template matching (Sapp et al. 2012) and support vector machines (Yokokawa et al. 2012) have been made to use the data from a large cohort of patient to learn the relationship between the origin of ventricular activation and the ECG data. However, limited accuracy has been reported so far. One of the major challenges, as mentioned earlier, is the

significant inter-subject variations across individuals (Park, Kim, and Marchlinski 2012).

In this paper, we investigate for the first time the use of deep generative models (Kingma and Welling 2013) (Rezende, Mohamed, and Wierstra 2014) to explicitly separate subject-specific variations in the ECG data from the hidden representation informative for the clinical task at hand. Learning to disentangle the underlying causal factors of the data has received increasing attention in the recent advancement of deep learning (Bengio, Courville, and Vincent 2013). A variety of models, many built off variational autoencoder (VAEs) and generative adversarial network (GANs), have been shown to be effective in learning deep representations informative for the task at hand while removing the non-informative factors of variations in domains such as computer vision (Mathieu et al. 2016), computer graphics (Kulkarni et al. 2015) and natural language processing (Hu et al. 2017). Outside these domains and into domain such as medicine and biology, however, the notion of learning disentangled representations to improve the task at hand has been little explored.

In this paper, we recognize the challenge of inter-subject variations inside clinical data with the potential solution of learning disentangled representations. In specific, we deal with a scenario where training labels for the task-specific representations are available and none of the factors of variations is observed. To achieve disentanglement in this setting, we leverage both variational autoencoders (VAEs) (Kingma and Welling 2013) (Rezende, Mohamed, and Wierstra 2014) and maximum mean discrepancy (MMD) (Gretton et al. 2007) along with contrastive regularization. Formally, for the specific task of predicting VT origin from ECG data, we aim to learn a latent representation z to capture the individual level physiological and pathological variations, and a representation for the task-specific factor (VT factor) v to capture the relationship between origins of ventricular activation and ECG data as defined by the underlying biophysics. The latter is learned by a deterministic encoder, while the former is learned by a variational encoder conditioned on the latter. These two representations are encouraged to be statistically independent by matching all order of moments of the marginal posterior distribution $q(\mathbf{z}|\mathbf{v}_i)$ with $q(\mathbf{z}|\mathbf{v}_i)$ where $i \neq j$ and $k = \{1,..,i,..,j,..,c\}$ represents c categories of the task-specific representation v. To further increase the discriminative power of the task-specific representation, contrastive regularization is imposed on the deterministic encoder for a weak supervision given the label information in the training data. Our main contributions include:

- We investigate the first use of deep generative models to disentangle inter-subject variations within ECG data from the representation informative for clinical task at hand.
- We propose a variational encoder conditioned on the codes of a deterministic encoder, to learn to separate intersubject variations from task-specific representations while learning the latter with a weak supervision via contrastive loss.
- We propose a mechanism to encourage the statistical in-

- dependence between the task-specific representation and inter-subject variations by matching all order of moments between the distributions learned from conditional VAE for the latter conditioning on the former.
- We demonstrate the performance of our approach on a challenging clinical task of localizing the origin of VT from ECG data, and compare its performance with the standard VAE.

The article is organized as follows. We introduce several related works in the next section. We provide a brief background of some preliminaries about variational autoencoder and maximum mean discrepancy. In model section, we present our proposed methodology along with training procedure. In experiment section, we present qualitative and quantitative analysis of presented model. Finally, we conclude the paper with limitations and possible future avenue.

Related Works

It has been widely accepted that different representations can entangle and hide explanatory factors of variation to a different extent within the data (Bengio, Courville, and Vincent 2013). The challenge of separating factors of variation from the data has been a topic of emerging interest in computer vision and related fields. As an early approach, a bilinear model (Tenenbaum and Freeman 2000) was proposed to capture sufficiently expressive representations of factors of variations in the data. They used the proposed model to separate the handwriting style from the content when recognizing handwritten digits. The work in (Chen et al. 2017) presents the discriminative architecture using denoising autoencoder to address inter-subject variations in ECG data. In the context of generative models, the work in (Desjardins, Courville, and Bengio 2012) and (Reed et al. 2014) used restricted Boltzman machine to model multiplicative interactions between latent factors of variation, separating identity from emotion (Desjardins, Courville, and Bengio 2012) or expression from pose (Reed et al. 2014) in facial images.

Recent works in learning disentangled representation started to exploit the power of deep generative models like VAE and generative adversarial networks (GAN). The work in (Mathieu et al. 2016) performs unsupervised disentanglement leveraging GAN. Similarly, the work in (Chen et al. 2016) used information-theuritic approach to learn disentangled representations in GAN framework. However, training such adversarial network requires careful optimization of a difficult minimax problem (Li, Swersky, and Zemel 2015).

The work in (Higgins et al. 2016) used a VAE framework where they demonstrated that by enforcing independence of features in the latent state it is possible to learn representations generalizing well to the new task. In (Kingma et al. 2014), a conditional VAE was used to demonstrate the disentanglement between the label information and the style in a semi-supervised setting by providing an additional one-hot vector as an input to the generative model. The work in (Louizos et al. 2015) extended this idea by adding MMD to further encourage statistical independence. The work in both (Kingma et al. 2014) and (Louizos et al. 2015), however, requires a certain representations to be observed by condition-

ing the other representations on it. The work in (Kulkarni et al. 2015) explored the disentanglement of content and style in the domain of computer graphics and produced impressive result benefiting from unique training procedure requiring strong supervision in designing the data.

Background

Variational Autoencoder

The variational autoencoder (VAE) (Kingma and Welling 2013), (Rezende, Mohamed, and Wierstra 2014) is a generative model where the data \mathbf{X} is generated by the generative distribution $p_{\theta}(\mathbf{X}|\mathbf{z})$ involving a set of latent variables \mathbf{z} , which follows a prior distribution of $p(\mathbf{z})$. This process can be defined as:

$$\mathbf{z} \sim p(\mathbf{z}); \mathbf{X} \sim p_{\theta}(\mathbf{X}|\mathbf{z})$$

Due to the intractability of posterior inference of latent variables, the parameter estimation of such graphical model is challenging. This has been tackled by using stochastic gradient variational bayes (SGVB) (Kingma and Welling 2013) framework. Here, the proposal distribution $q_{\phi}(\mathbf{z}|\mathbf{X})$ is used to approximate the true posterior $p(\mathbf{z}|\mathbf{X})$ using variational inference. Combining this inference network $q_{\phi}(\mathbf{z}|\mathbf{X})$ with a generative network $p_{\theta}(\mathbf{X}|\mathbf{z})$ would resemble autoencoding architecture and thus has been known as variational autoencoder. As such, the objective of VAE is to maximize the following variational lower bound with respect to parameters θ and ϕ :

$$\sum_{n=1}^{N} \log p_{\theta}(\mathbf{X}_{n}) \ge \sum_{n=1}^{N} \mathbb{E}_{q_{\phi}(\mathbf{z}_{n}|\mathbf{X}_{n})} [\log p_{\theta}(\mathbf{X}_{n}|\mathbf{z}_{n})] -KL[q_{\phi}(\mathbf{z}_{n}|\mathbf{X}_{n})||p(\mathbf{z}_{n})]$$
(1)

This objective function (1) can be interpreted as minimizing a reconstruction error in the first term along with minimizing the KL-Divergence between the variational approximation of the posterior $q_{\phi}(\mathbf{z}|\mathbf{X})$ and the prior $p(\mathbf{z})$. The inference network $q_{\phi}(\mathbf{z}|\mathbf{X})$ and generative network $p_{\theta}(\mathbf{X}|\mathbf{z})$ are usually modeled with a neural network.

MMD

Maximum mean discrepancy (MMD) is a closed-form non-parametric two-sample test proposed by (Gretton et al. 2007). The purpose of the test involved in MMD is to determine if two samples are from different distributions. If we have two sets of samples $\mathbf{X} = \{\mathbf{x}_i\}_{i=1}^M$ and $\mathbf{Y} = \{\mathbf{y}_j\}_{j=1}^N$, then with the use of MMD we are able to determine if generating distribution $P_X = P_Y$. This is achieved by comparing the statistics between the two samples. A simple test could be the distance between empirical statistics $\phi(.)$ of the two samples:

$$\left\| \frac{1}{M} \sum_{i=1}^{M} \phi(\mathbf{x}_i) - \frac{1}{N} \sum_{j=1}^{N} \phi(\mathbf{y}_i) \right\|^2$$
 (2)

MMD (Gretton et al. 2007) is an estimator by expanding the above square such that kernel tricks can be applied.

$$MMD[\mathbf{X}, \mathbf{Y}] = \frac{1}{M} \sum_{i=1}^{N} k(\mathbf{x}_i, \mathbf{x}_i)$$
$$-\frac{2}{MN} \sum_{i,j=1}^{M,N} k(\mathbf{x}_i, \mathbf{y}_i) + \frac{1}{N} \sum_{j=1}^{N} k(\mathbf{y}_i, \mathbf{y}_i)$$
(3)

The choice of ϕ in (2) determines what order of statistics we want to match. For instance, a simple identity function leads to matching the sample mean. When kernel trick is applied, the sample vectors are mapped into an infinite dimensional feature space and when this feature space corresponds to universal reproducing kernel hilbert space (RKHS), it is shown in (Gretton et al. 2012) that asymptotically, MMD is 0 if and only if $P_X = P_Y$. Therefore, minimizing MMD for a universal kernel like Gaussian kernel $k(\mathbf{x}, \mathbf{y}) = \mathbf{e}^{-\beta||\mathbf{x}-\mathbf{y}||^2}$ in (3) can be interpreted as matching or minimizing certain distance between weighted sums of all raw moments (Li, Swersky, and Zemel 2015) of P_X and P_Y .

Model

Conditional generative model

In this work, we introduce a conditional probabilistic model admitting two sources of variations: VT factor ${\bf v}$ to capture the general biophysical relationship between origins of ventricular activation and ECG data, and a latent representation ${\bf z}$ to capture subject-specific level of adjustments to the data involving factors such as heart and torso anatomy and characteristics of structural abnormality of the heart that are not explicitly observed. First, considering the task-specific factor ${\bf v}$ as an observed variable, we frame the conditional generative process as:

$$\mathbf{z} \sim p_{\theta}(\mathbf{z}); \mathbf{X} \sim p_{\theta}(\mathbf{X}|\mathbf{z}, \mathbf{v})$$

We model this conditional generative process in the variational autoencoder framework, where $p_{\theta}(\mathbf{X}|\mathbf{z},\mathbf{v})$ is a likelihood function described by a decoder network. The approximate posterior to infer subject-specific factor \mathbf{z} is modeled using an independent Gaussian distribution $q_{\phi}(\mathbf{z}|\mathbf{X},\mathbf{v}) = N(\mu, \sigma I)$. This is specified by an encoder network. The stochasticity in $q_{\phi}(\mathbf{z}|\mathbf{X},\mathbf{v})$ comes from both the data-distribution and the randomness of Gaussian distribution. Finally by choosing standard isotropic Gaussian prior $p(\mathbf{z}) = N(0, I)$, we can obtain the following lower bound:

$$\sum_{n=1}^{N} \log p_{\theta}(\mathbf{X_n}|\mathbf{v_n}) \ge \sum_{n=1}^{N} \mathbb{E}_{q_{\phi}(\mathbf{z_n}|\mathbf{X_n},\mathbf{v_n})} [\log p_{\theta}(\mathbf{X_n}|\mathbf{z_n},\mathbf{v_n})] - KL[q_{\phi}(\mathbf{z_n}|\mathbf{X_n},\mathbf{v_n})||p(\mathbf{z_n})] = \mathcal{F}(\phi,\theta;\mathbf{X}_n,\mathbf{v_n})$$
(4)

The objective in 4 gives us a simple conditional VAE similar to that used in (Louizos et al. 2015). However, unlike in (Louizos et al. 2015), neither v nor z in this work is observed. Therefore, in this work, v in (4) is obtained using

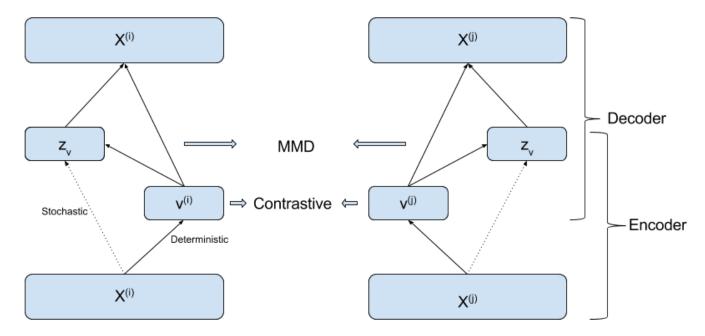


Figure 2: Illustrative diagram of the proposed network. The encoder takes a pair of input ECG data and outputs the VT factor \mathbf{v} and subject-specific factor \mathbf{z} . The decoder takes these disentangled factors as an input and reconstruct the ECG data.

a separate deterministic encoder $q(\mathbf{v}|\mathbf{X})$ in which the only source of stochasticity comes from the data distribution. As such, the conditional likelihood in (4) can be expressed as:

$$\mathbf{X} \sim p_{\theta}(\mathbf{X}|\mathbf{z}, q(\mathbf{v}|\mathbf{X}))$$

Although the above model admits two separate sources of causal factors generating the ECG data, there is nothing preventing all of the information from flowing through the latent factor \mathbf{z} . In other words, the decoder could learn to ignore the VT factor \mathbf{v} . Below, we describe two regularization techniques added to the network to encourage the separation of the two factors and to learn a more discriminative task-specific representation.

Discriminative regularization

MMD

The work in (Li, Swersky, and Zemel 2015) and (Dziugaite, Roy, and Ghahramani 2015) used MMD measure to generate samples from an unknown distribution given i.i.d data, essentially, acting as a substitute of difficult adversarial training (Goodfellow et al. 2014). The idea in (Li, Swersky, and Zemel 2015) and (Dziugaite, Roy, and Ghahramani 2015) is to force the output of the generator network to match all orders of statistics of the training data. Motivated by this idea, we propose to match all order moments between marginal posterior distributions of subject-specific factors z which is conditioned upon VT factor v representing different segments in the heart:

$$MMD[\mathbf{z}_{\mathbf{v_i}}, \mathbf{z}_{\mathbf{v_j}}] = \left\| \mathbb{E}_{p(\mathbf{X}|\mathbf{v_i})} [\mathbb{E}_{q(\mathbf{z}|\mathbf{X},\mathbf{v_i})} [\phi(\mathbf{z}_{\mathbf{v_i}})]] - \mathbb{E}_{p(\mathbf{X}|\mathbf{v_j})} [\mathbb{E}_{q(\mathbf{z}|\mathbf{X},\mathbf{v_j})} [\phi(\mathbf{z}_{\mathbf{v_j}})]] \right\|^{2}$$
(5)

where $i \neq j$ and $k = \{1, ..., i, ..., j, ..., c\}$ represents c categories represented by specific factor \mathbf{v} .

Minimizing (5) can be interpreted as forcing the network to learn subject specific factor \mathbf{z} independent of VT factor \mathbf{v} . The MMD objective shown in (5) is differentiable when the kernel is differentiable. As such, we are using Gaussian kernels $k(\mathbf{x},\mathbf{y}) = \mathbf{e}^{-\beta||\mathbf{x}-\mathbf{y}||^2}$ whose gradient has a simple form and thus could be easily back-propagated through the inference network $q_{\phi}(\mathbf{z}|\mathbf{X},\mathbf{v})$. Bandwidth parameter β of kernel k(.) is considered as the hyper-parameter during training of the network.

Contrastive loss

MMD regularizer would be enough if the VT factor ${\bf v}$ is observed and the objective is to learn the subject-specific variations, similar to (Louizos et al. 2015) while learning fair representations. However, in this work, we are also interested in learning representation ${\bf v}$ informative for the clinical task at hand. To do so, we hypothesize that for the ECG data originating from nearby locations, the VT factor ${\bf v}$ should be similar regardless if the data are collected from the same patient; otherwise ${\bf v}$ should be different. In this way we are learning an embedding of VT factor ${\bf v}$ where similar data pairs are embedded nearby and dissimilar data pairs are far apart. To facilitate such pair-wise comparison, we propose an additional contrastive loss (Hadsell, Chopra, and LeCun

2006) as a weak supervision to the deterministic encoder. The contrastive loss is formulated as:

$$L_{c}(\mathbf{v}^{(i)}, \mathbf{v}^{(j)}) = e_{v}^{p} \frac{1}{2} \|\mathbf{v}^{(i)} - \mathbf{v}^{(j)}\|_{2}^{2} + (1 - e_{v}^{p}) \frac{1}{2} \max(0, \alpha - \|\mathbf{v}^{(i)} - \mathbf{v}^{(j)}\|_{2}^{2})$$
(6)

where e_v^p is 1 if the pair of the ECG data share the same VT exit label and 0 otherwise. Here α is the margin and the dissimilar pairs contribute to the loss function only if their distance is within this margin. We treat α as the hyper-parameter during training of the network.

To calculate both MMD and contrastive loss, we require a training architecture (presented in next section) which facilitate pair-wise comparison as shown in Fig. 2. Combining (4), (5) and (6), we get our combined objective as:

$$\mathcal{F}(\phi, \theta, \mathbf{X}_{n}, e_{v}^{p}) = \sum_{m_{i}, m_{j}}^{M} \left\{ \mathcal{F}(\phi, \theta; \mathbf{X}_{m_{i}}, \mathbf{v}_{m_{i}}) + \mathcal{F}(\phi, \theta; \mathbf{X}_{m_{j}}, \mathbf{v}_{m_{j}}) \right\} / 2$$

$$- \lambda_{1} \sum_{m_{i}, m_{j}; m_{i} \neq m_{j}}^{M} MMD[\mathbf{z}_{\mathbf{v}_{\mathbf{m}_{i}}}, \mathbf{z}_{\mathbf{v}_{\mathbf{m}_{j}}}]$$

$$- \lambda_{2} \sum_{m_{i}, m_{j}}^{M} L_{c}(\mathbf{v}^{(i)}, \mathbf{v}^{(j)})$$

$$(7)$$

where λ_1 and λ_2 are the non-negative weight. Here, the upper bound M in the summation refers to the total number of pairs presented to the networks as the input data.

Training architecture

We randomly generate M training pairs $X^p = (\mathbf{X}^{(i)}, \mathbf{X}^{(j)})$ from the training data ensuring different beats from the same pacing location are not paired together. Each pair is given a label e^p_v which is 1 if the pair of the ECG data share the same VT exit label and 0 otherwise. The architecture is inspired from *siamese* architecture (Chopra, Hadsell, and LeCun 2005) which consists of two copies of our proposed model sharing the same set of parameters. The input to the entire system is a pair of ECG data $(\mathbf{X}^{(i)}, \mathbf{X}^{(j)})$ and a label e^p_v . The output of the entire encoder can be represented as $\{(\mathbf{v}^{(i)}, \mathbf{v}^{(j)}), (\mathbf{z}_{v_i}, \mathbf{z}_{v_j})\}$ which is fed to the decoder network to calculate approximation of the ECG data as $(\tilde{\mathbf{X}}^{(i)}, \tilde{\mathbf{X}}^{(j)})$. The illustrative diagram of the network is shown in Fig. 2.

Experiments

We apply the proposed model to disentangle subject-specific variations present in the ECG data to learn the representation informative for the clinical task of localizing origin of ventricular tachycardia. Following the setup in (Yokokawa et al. 2012), we divide the left-ventricular surface into ten segments. In this setting, the VT localization process becomes a 10-class classification problem.

The Data

Data are collected from 39 patients during routine pacemapping procedures as the patients underwent ablation of scar-related VT. The database includes 15-second 12lead ECG recordings recorded from 1012 distinctive pacing sites on the left-ventricular (LV) endocardium, all identified on an intro-operative electroanatomical mapping system (CARTO3) with known coordinates. All ECG data are processed with noise removal and baseline correction using an open-source software. As illustrated in Fig. 3, manual selection and extraction of QRS complexes are carried out by student trainees to avoid motion artifacts, ectopic beats, and non-capture beats. The final input signal is in the form of one ORS beat from 12 leads, down-sampled in time to 100×12 in size. Because multiple quality beats can be extracted from each ECG recording, we obtain in-total 16848 sets of ECG data and the corresponding sites of pacing in the form of a segment label. Using these data, we try to learn to predict the origin of the ventricular activation from the ECG data, where the origin is in the form of one of the ten LV segments. When applied to VT, the predicted origin of ventricular activation corresponds to the site where the VT circuit exits the scar.

Experimental setup

To evaluate the proposed model, the entire dataset is split into training, validation and test set as follows: 10292 from 22 patients, 3017 from 5 patients, and 3539 from 12 patients. Note that the patients in the test and validation sets are not included in the training set. For each encoder, we have two hidden layers of 800 and 600 units. Similarly for the decoder, we have two hidden layers of size 600 and 800 units. Different sets of hyperparameters including the size of VT factor v, the size of subject-specific factor z, the margin for contrastive regularization α and the bandwidth parameter β of the MMD are tuned on validation set. The scaling parameter λ_1 and λ_2 are set to 1. Rectified linear units (ReLU) were used for the non-linear activation in both encoder and decoder network. The training data is divided into minibatches and to normalize the data in each mini-batch, batch normalization (Ioffe and Szegedy 2015) layers are used in both encoder and decoder network. The optimization of the objective function was done with stochastic optimizer Adam (Kingma and Ba 2014). Finally the classification task of the learned factor was performed using a simple linear classifier to ensure classifier itself does not aid in boosting the classification performance. The codes associated with this work are available at https://github.com/Prasanna1991/ MMD VAE.

Results

We evaluate the performance of our model by first performing classification by associating the learned VT factor into different segments in the left-ventricular region of the heart. We also perform experiments to evaluate and analyze the disentanglement abilities of the network.

Classification

The results for classification of VT factor v into pre-defined ten segments for proposed model along with other different

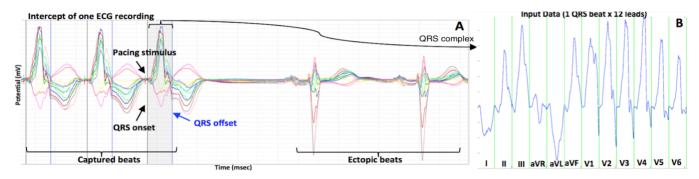


Figure 3: (A) Illustration of experimental data and processing. ECG recordings are pre-processed for extraction of noise-free QRS complex. (B) The final data which is formed by concatenating the extracted QRS complex from each lead.

Architecture	Segment classification	
	(in %)	
QRS Integral	45.28	
VAE	52.37	
VAE+MMD	53.17	
Proposed	55.29	

Table 1: Segment classification accuracy of the proposed method versus three comparison methods. Classification accuracy is reported in percentage on separately held out test dataset.

models are presented in Table 1. The localization accuracy in terms of classification accuracy of the proposed model is compared against commonly used prescribed features of 120-ms QRS Integral (Sapp et al. 2012), standard VAE without disentangling the two representations and finally with standard VAE with MMD regularization where disentanglement between subject-specific factor and task-specific VT factor is achieved by matching all order of moments between the marginal posterior distributions of subject-specific factor conditioned on task-specific VT factors representing different ventricular segments of the heart. To be consistent with the network architecture, all models are implemented with the same siamese architecture. The results are presented in Table 1 where we can see an improvement as we go from feature based approach to deep network in the form of standard VAE. Further improvement is observed with VAE+MMD model where model is encouraged to separate VT factor v with subject-specific factor z. Finally, classification is further improved with the proposed model where contrastive regularizer is added for the learning of VT factor v on top of VAE+MMD model.

Analyzing disentanglement

To gain an insight into the effect of factor disentangling, we performed different experiments on both causal factors to the ECG data. First, we examine if information of one factor is removed from the representation learned for other factor, i.e., whether information about VT origin is removed from the patient-specific variation, and vice versa. To do so, we attempt to use the learned representation for inter-subject variations z to classify the location of the VT, and use the

factor	VT location	patient ID
	(in %)	(in %)
v	55.29	15.23
$\overline{\mathbf{z}}$	17.19	32.33
random-chance	10	4.5

Table 2: Classification accuracy when one factor is associated with label of the other factor. Classification accuracy involving subject-specific factor **z** is reported on train dataset.

learned representation for VT location v to recognize the patient ID. Both of them are compared with random chance as shown in Table 2. As already seen in Table 1, VT factor v is informative for the specific task of localizing the origin of ventricular activation. In comparison, Table 2 demonstrates that the subject-specific factor z has minimal information about the specific-task. On the other hand, subject-specific factor z is much better in associating with different patient ID present in training data compared to VT factor v.

Further, we present visualization of ECG signals used and generated from the experiments in Fig. 4. For the purpose of analyzing disentanglement we generate different ECG signals by setting z=N(0,I) as shown in Fig. 4 (C). For VT factor ${\bf v}$ we use the representation learned from $q({\bf v}|{\bf X})$ where ${\bf X}$ represents different ECG signals with the same VT label. Although the ECG data comes from different subjects, because of the same VT factor, the generated ECG signal looks alike in all three cases. However, there still exist variations and to demonstrate such variations, we present the detailed view by selecting one of the lead (lead aVR) among 12-leads in Fig. 4 (D) where maximum variations can be observed.

Discussion

The results presented in this work are obtained when the models only see 22 patients during training, a very limited sample size for the purpose of disentangling inter-subject variations. Moreover, as the data were collected from actual pace-mapping procedures during ablation, available data on each patient typically covers only a specific region of the heart that is within and around the myocardial scar. The networks within the models are constructed with a simple

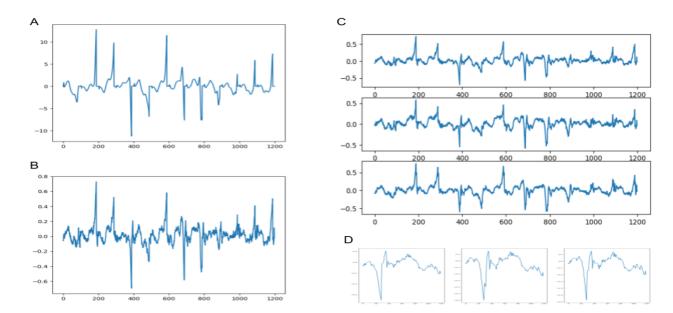


Figure 4: (A) Original ECG as presented to the model. (B) Reconstructed ECG. (C) Generated ECG signal by setting $\mathbf{z} = N(0, I)$ and \mathbf{v} from different ECG signals with the same VT labels. (D) Lead aVR of the generated ECG as shown in (C) for a more detailed view.

multilayer perceptron (MLP) and using powerful feature extractor like convolutional neural network (CNN) (Rajpurkar et al. 2017) within the encoder network and/or the decoder network may further improve the performance. The ability to separate the subject-specific factor with the factor informative for clinical task at hand, at the presence of a limited dataset, however, does demonstrate the feasibility of the proposed model.

Conclusion

The paper investigates the first use of deep generative models to disentangle inter-subject variations within the ECG data from the representation informative for the clinical task at hand. The feasibility of the proposed framework is demonstrated by a novel yet important clinical task to localize the VT exit which may enhance the efficiency and efficacy of ablation therapies. The paper presents a conditional variational autoencoder to extract the subject-specific adjustments to the ECG data, conditioned on task-specific representations learned from a deterministic encoder. To encourage independence between these representations, kernel based MMD measure is used. The learning of task-specific representations from a deterministic encoder is performed using weakly supervised contrastive regularization.

Acknowledgement

This work is supported in part by the National Institutes of Health [No: R21HL125998] and the National Science Foundation [No: ACI-1350374

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