



One-class SVM to identify candidates to Rerference Genes based on the augment of RNA-seq data with Generative Adversarial Networks

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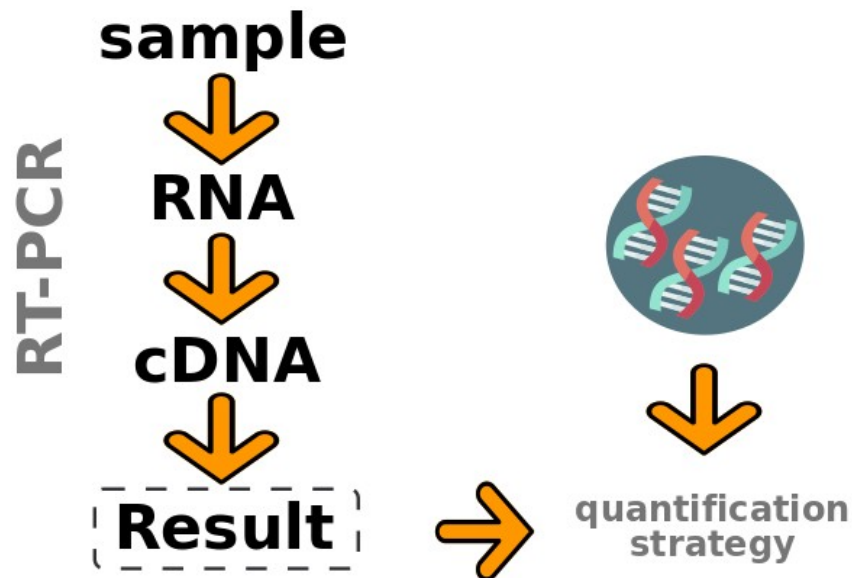
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- Introduction
- Proposed Method
- Experiments and Results
- Conclusions and Future Works

■ Reference Genes (RG)

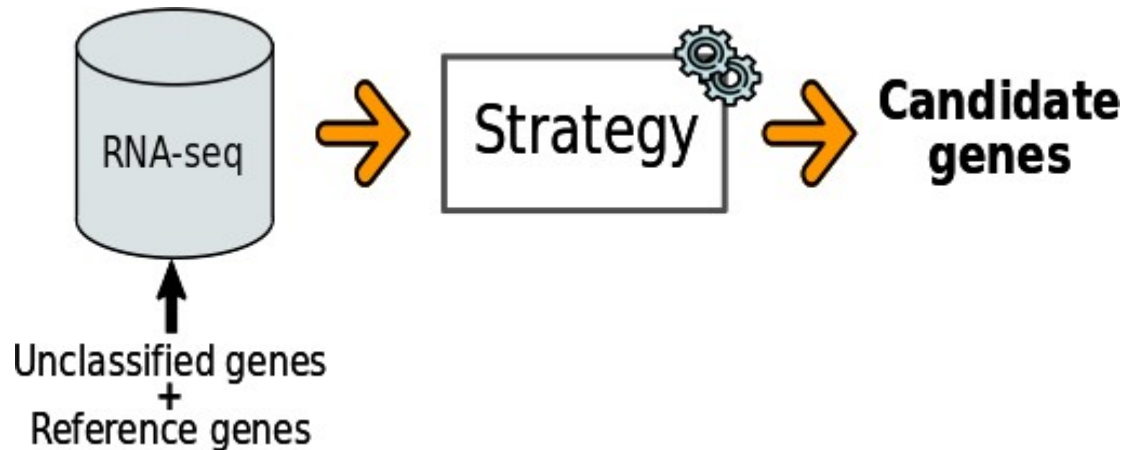
- Constitutive genes
- Expressed in all cells
- Used in internal controls (gene expression analysis)



- Normalize gene expression
- Demonstrates the variability and imperfections of the technology

Identify candidates for Reference Genes

- Pipeline:



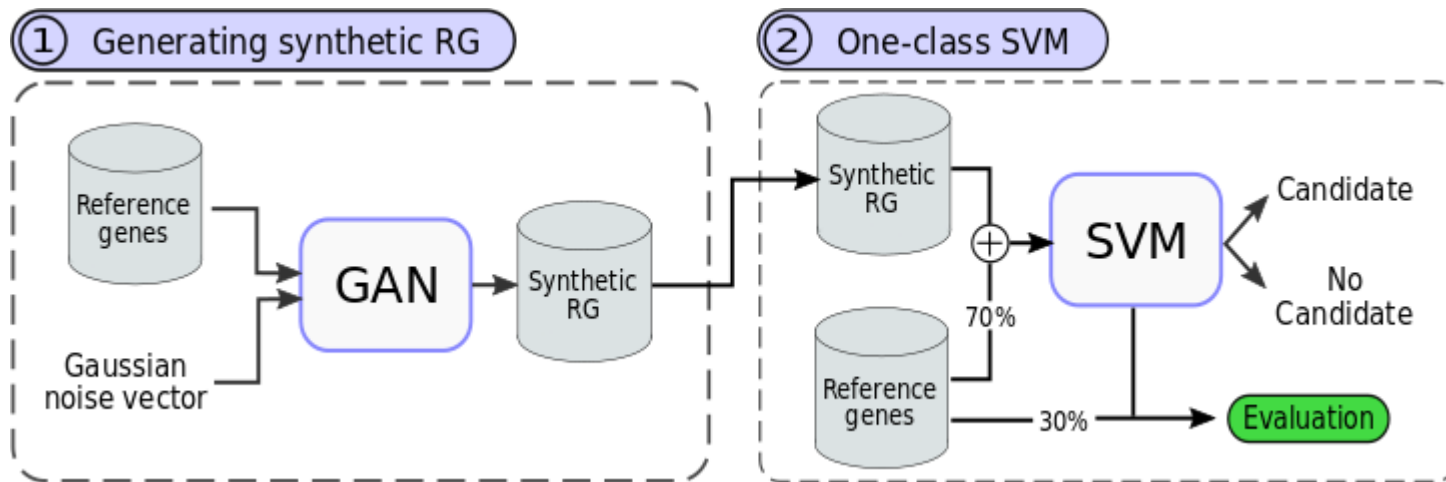
- Strategies:

- Based on clustering (Euclidean distance)
- Based on optimization algorithms

Proposed Method

■ Steps:

- 1) Generative Adversarial Networks to augment of RNA-seq data
- 2) One-Class SVM to select candidate genes



■ Initial step: data processing

Proposed Method

Data processing

■ Pipeline:

1) Normalization with RPKM:

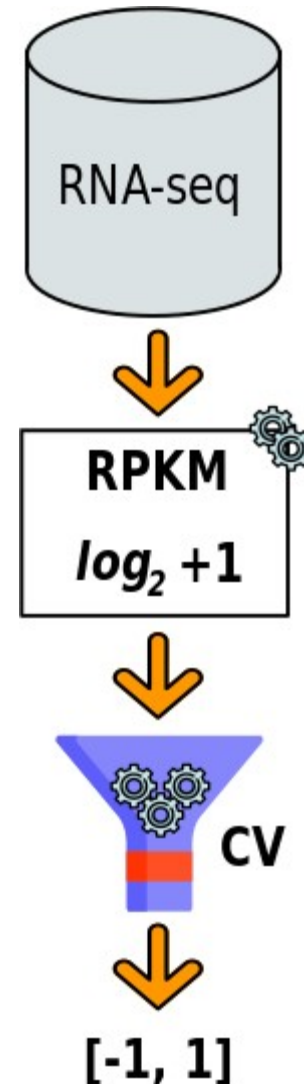
$$RPKM = \frac{numReads * 10^9}{geneLength * TMReads}$$

2) $\log_2 + 1$

3) Remove outliers based on the Coefficient of Variation (CV):

$$CV = \frac{\sigma}{\mu}$$

4) Data scaled between $[-1, 1]$

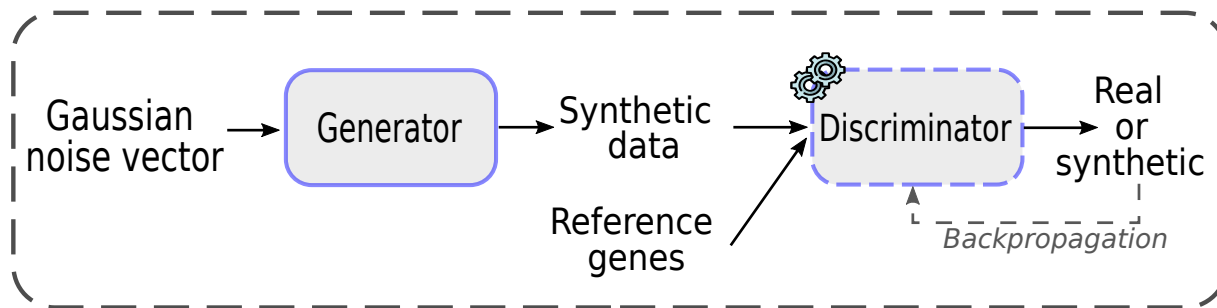


Proposed Method

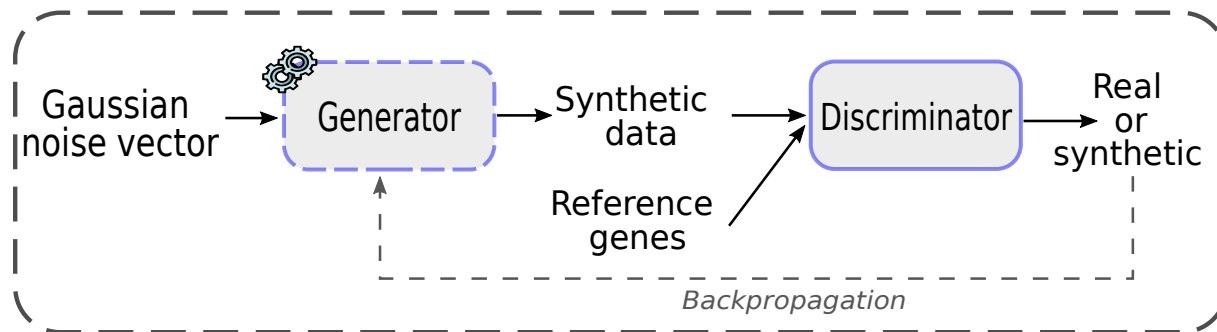
Generative Adversarial Networks (GAN)

■ Steps:

1) Training the Discriminator network and freezing their weights

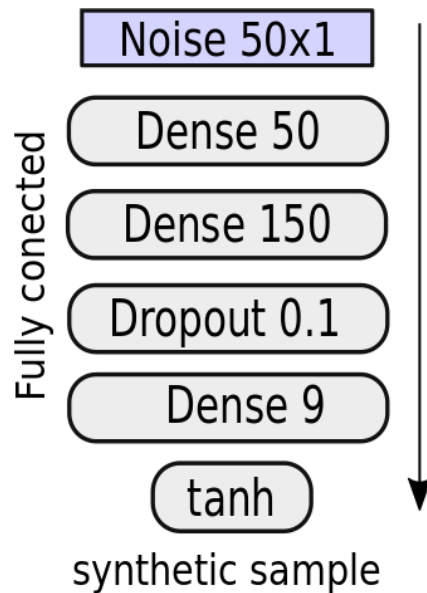


2) Training the Generator network

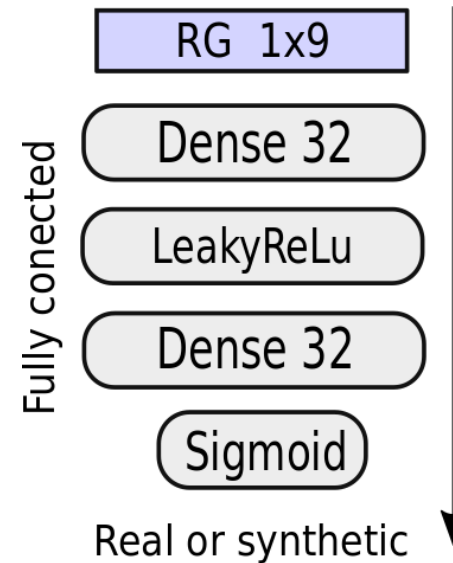


Proposed architecture

- Generator network



- Discriminator network



- A normal distribution $N(0,1)$ as a noise vector
- Stochastic Gradient Descent to compute the gradients

Evaluation

- A proposed Similarity metric $\mathbf{S(x,x')}$ to evaluate the performance of the GAN:

$$S(x, x') = \sum_i^m \sum_j^{n_g} \sum_k^{n_f} \frac{|x_i^{(k)} - x'_j{}^{(k)}|}{n_f n_g m} + |0.5 - \frac{1}{n_g} \sum_j^{n_g} \hat{y}_j|$$

- \hat{y} : Class predicted by D network for a synthetic gene
- x' : Set of synthetic genes generated by the G network
- x : Set of Reference Genes
- m : Number of Reference Genes
- n_g : Number of synthetic genes
- n_f : Number of features (gene expression)

Evaluation

- A proposed **$E(\mathbf{x}')$** metric to select the best sample of synthetic data:

$$E(x') = \frac{1}{n_g} \sum_j^{n_g} \left[CV(x'_j) + \frac{1 - D(x'_j)}{D(x'_j)} \right]$$

- CV : Coefficient of variation
- x' : Set of synthetic genes generated by the G network
- $n_g = 300$: Number of synthetic genes
- **Other metrics:** Binary Cross-Entropy, Precision score

One-Class SVM

- Implements the RBF kernel (Gaussian kernel)
- Recall score to evaluate the performance of the One-Class SVM:

$$Recall = \frac{TP}{TP + FN}$$

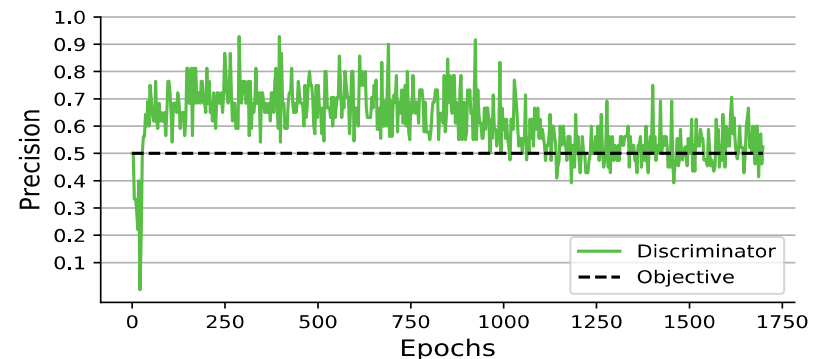
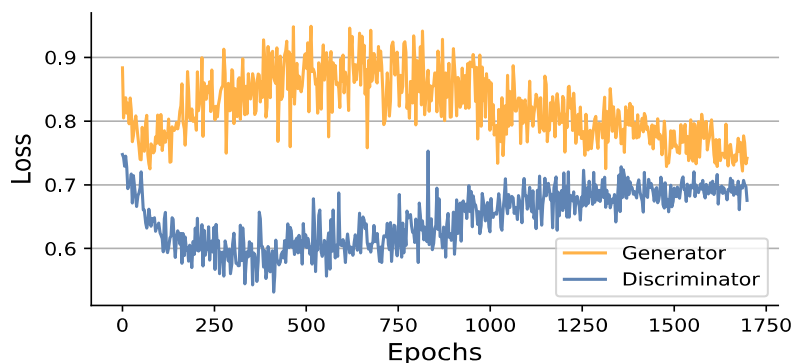
- TP : True Positives
- FN : False Negatives
- Recall score allows to measure the ability of the classifier to find all positive samples (RG)
- A Recall score close to one indicates that the classifier has a good performance

Experiments and Results

- The *Escherichia coli* MG1655 dataset was used
- Parameters used for training the GAN architecture

Parameter	Generator	Discriminator
Optimizer	SGD	SGD
Learning rate	0.00015	0.001
Decay rate	0.00015/1700	0.001/1700
Momentum	0.92	0.9
Epochs	1700	1700

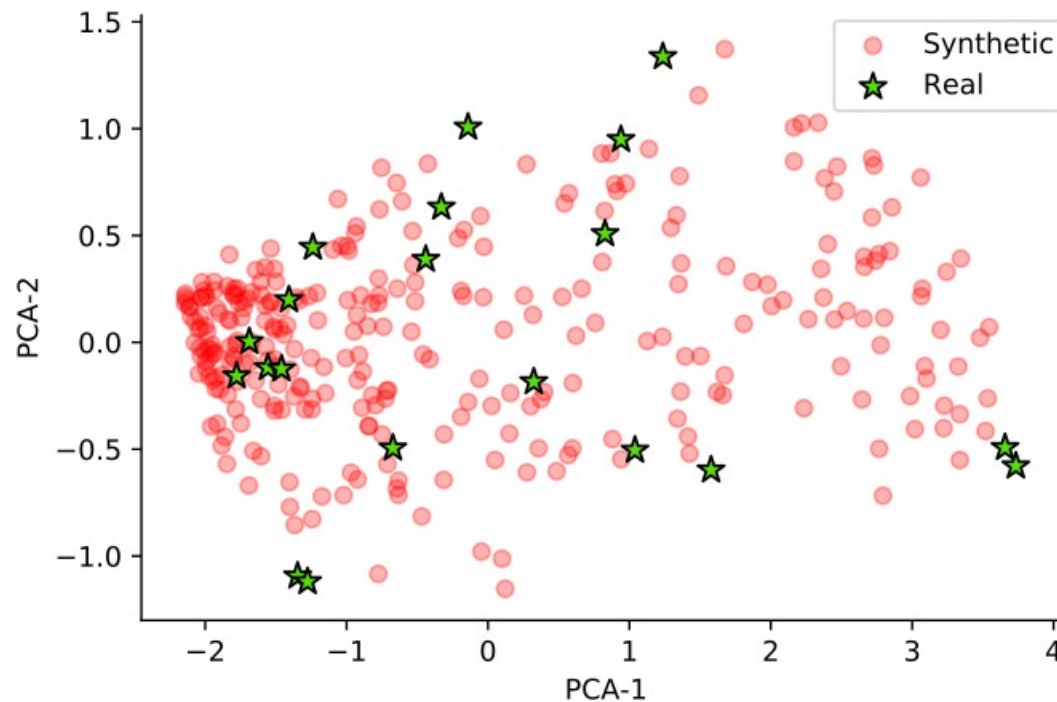
- Convergence process



Experiments and Results

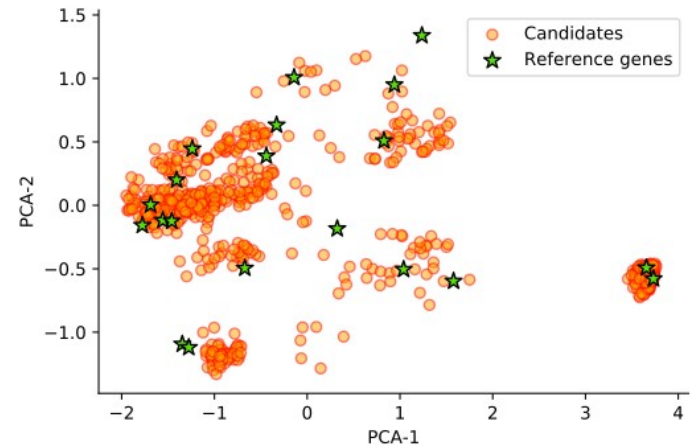
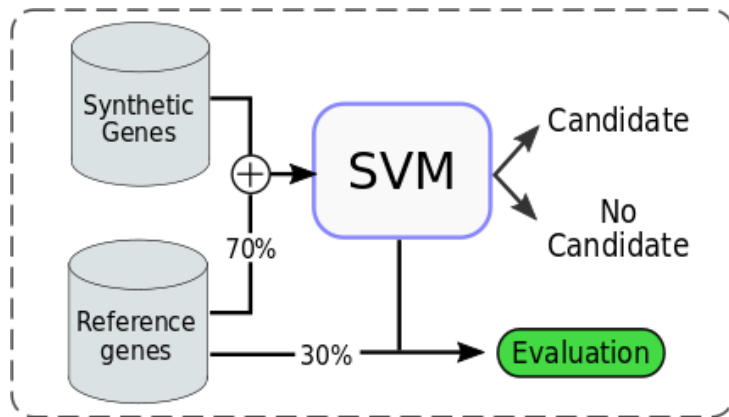
■ Selection of synthetic genes

- We generated 5000 sets of 300 synthetic genes and selected the set with the smallest value of $E(\mathbf{x}')$



Experiments and Results

■ Selection of candidate genes



■ Recall score on test and training data

Metrics	Training data		Test data	
	Augmented data	Reference Genes (70%)	Reference Genes (30%)	Synthetic data
<i>Recall</i>	98.40%	92.85%	85.71%	98.26%

■ We selected the 11 most relevant candidate genes

Experiments and Results

- Comparing results with augmented and unaugmented data:

Reference Genes	<i>Recall score</i>	
	Training data	Test data
Augmented	98.40%	85.71%
Unaugmented	85.71%	66.66%

- With augmented dataset, we improved recall score of proposed classifier by 19%

- With the proposed method we were able to identify 807 possible candidate genes from a total of 4170 unclassified genes from the *Escherichia coli* MG1655 dataset.
- Augmenting the set of RG we can increase the performance of the One-class svm classifier
- Code is being maintained on GitHub:
https://github.com/ejrueda/20_ICCSA_RG_GAN

- Other GAN architecture can be tested
- A more extensive hyperparameter tuning could also be applied in the training of the GAN architecture
- Other One-class classifiers can be tested (Ex: GAN)



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Thank you!

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