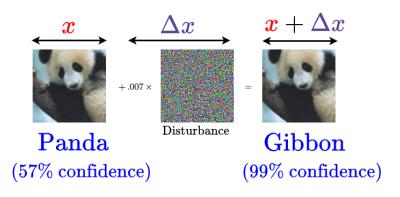
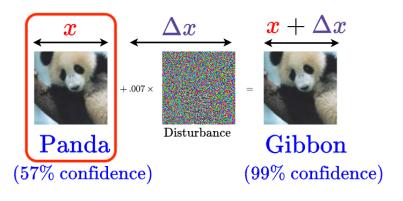
# Linear adversarial training, robustness in machine learning and applications to cardiology

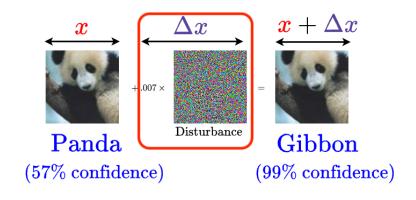
#### Antônio H. Ribeiro

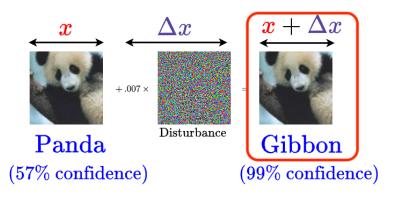
Uppsala University, Sweden

KTH, Royal Institute of Technology Stockholm, Sweden 2023









Adversarial training: Each training sample is modified by an adversary.

# Part I. Linear adversarial training

Regularization properties of adversarially-trained linear regression
Antônio H. Ribeiro, Dave Zachariah, Francis Bach, Thomas B. Schön.
NeurlPS (2023) - Spotlight

# Part II. Robustness of overparameterized models

Overparameterized Linear Regression under Adversarial Attack.

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Automatic diagnosis of the 12-lead ECG using a deep neural network Antônio H. Ribeiro , M.H. Ribeiro, Paixão, G.M.M. et al Nature Communications (2020)

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**▶** Linear regression:

$$\min_{\boldsymbol{\beta}} \sum_{i=1}^{\#train} (y_i - \boldsymbol{\beta}^{\top} \boldsymbol{x}_i)^2$$

**▶** Linear regression:

$$\min_{\beta} \sum_{i=1}^{\# train} (\underbrace{y_i}_{\text{observed}} - \underbrace{\beta^{\mathsf{T}} x_i}_{\text{linear prediction}})^2$$

► Linear regression:

$$\min_{\beta} \sum_{i=1}^{\# train} (y_i - \beta^\top x_i)^2$$

► Adversarial training in linear regression:

$$(y_i - \beta^\top (x_i + \Delta x_i))^2$$

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$$\sum_{i=1}^{\# train} \max_{\|\Delta x_i\| \le \delta} (y_i - (x_i + \Delta x_i)^\mathsf{T} \beta)^2$$

$$\sum_{i=1}^{\text{\#train}} \max_{\|\Delta x_i\| \le \delta} (y_i - (x_i + \Delta x_i)^\mathsf{T} \beta)^2$$

It can be rewritten as:

$$\sum_{i=1}^{\# train} \left( |\mathbf{y}_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}| + \delta \|\boldsymbol{\beta}\|_* \right)^2$$

where  $\|\cdot\|_*$  is the dual norm.

$$\sum_{i=1}^{\# train} \max_{\|\Delta x_i\|_{\infty} \le \delta} (y_i - (x_i + \Delta x_i)^{\mathsf{T}} \beta)^2$$

It can be rewritten as:

$$\sum_{i=1}^{\# train} \left( |y_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}| + \delta \|\boldsymbol{\beta}\|_1 \right)^2$$

where  $\|\cdot\|_1$  is the dual norm.

 $\blacktriangleright$   $\ell_{\infty}$ -adversarial attacks:

$$\sum_{i=1}^{\# train} \left( |\mathbf{y}_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}| + \delta \|\boldsymbol{\beta}\|_1 \right)^2$$

Lasso:

$$\sum_{i=1}^{\# train} \left( |\mathbf{y}_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}| \right)^2 + \lambda \|\boldsymbol{\beta}\|_1.$$

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▶  $\ell_{\infty}$ -adversarial attacks:

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#### Main results:

#1. Map  $\lambda \leftrightarrow \delta$  for which they yield the same result.

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 $\blacktriangleright$   $\ell_{\infty}$ -adversarial attacks:

$$\sum_{i=1}^{\# \operatorname{train}} \left( |y_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}| + \|\boldsymbol{\delta}\| \boldsymbol{\beta}\|_1 \right)^2$$

Lasso:

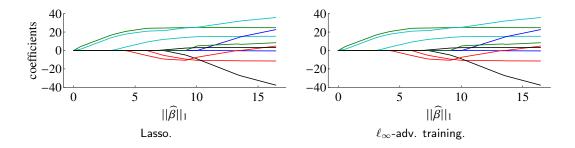
$$\sum_{i=1}^{\# train} \left( |\mathbf{y}_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta}| \right)^2 + |\boldsymbol{\lambda} || \boldsymbol{\beta} ||_1.$$

#### Main results:

- #1. Map  $\lambda \leftrightarrow \delta$  for which they yield the same result.
- #2. More parameters than data: abrupt transition into interpolation.
- #3. **Optimal choice** of  $\delta$  independent on noise level.

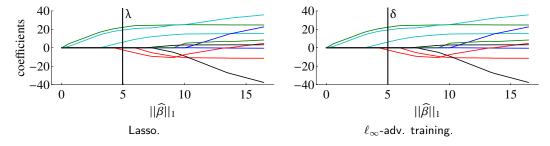
# # 1. Equivalence with Lasso

Map  $\lambda \leftrightarrow \delta$  for which they yield the same result.



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Map  $\lambda \leftrightarrow \delta$  for which they yield the same result.



The that yield the same result are not necessarily the same, i.e.:  $\delta \neq \lambda$ 

Lasso: transition only in the limit

$$\lambda o 0^+ \Rightarrow \sum_{i=1}^{\# train} \left( y_i - x_i^\mathsf{T} \beta \right)^2 o 0$$

Lasso: transition only in the limit

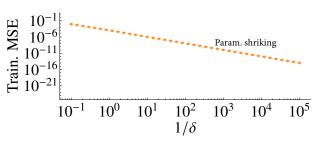
$$\lambda \to 0^+ \Rightarrow \sum_{i=1}^{\# train} \left( \mathbf{y}_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta} \right)^2 \to 0$$

$$\delta \in (0, \text{threshold}] \Rightarrow \sum_{i=1}^{\# train} \left( y_i - \mathbf{x}_i^\mathsf{T} \boldsymbol{\beta} \right)^2 = 0$$

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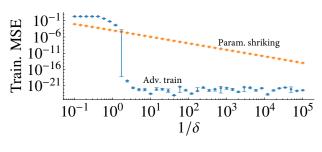
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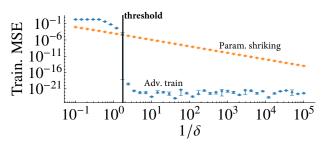
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# # 2. Equivalence with minimum norm interpolator

For  $\delta \in (0, \mathrm{threshold}]$ , the minimum-norm interpolator is the solution to adversarial training.

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For  $\delta \in (0, \mathrm{threshold}]$ , the minimum-norm interpolator is the solution to adversarial training.

#### Relevance

Connect adversarial training with double descent and benign overfitting

## # 3. Invariance to noise levels

To obtain near-oracle performance.

Lasso:

$$\lambda \propto \sigma \sqrt{\log(\# extit{params})/\# extit{train}}$$

 $ightharpoonup \ell_{\infty}$ -adversarial attack:

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 $\triangleright$   $\ell_{\infty}$ -adversarial attack:

$$\delta \propto \sqrt{\log(\#params)/\#train}$$

#### Data model

$$y = \underbrace{\mathbf{x}^{\top} \boldsymbol{\beta}^{*}}_{\text{signal}} + \underbrace{\boldsymbol{\sigma}}_{\text{noise std.}} \boldsymbol{\varepsilon}.$$

# Regularization properties of adversarially-trained linear regression

#### Additional results:

 $\triangleright$   $\ell_2$ -adv. attacks and ridge regression.

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# Regularization properties of adversarially-trained linear regression

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# Regularization properties of adversarially-trained linear regression

#### Additional results:

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- ▶ Connection to **robust regression** and  $\sqrt{Lasso}$ .

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# Part I. Linear adversarial training

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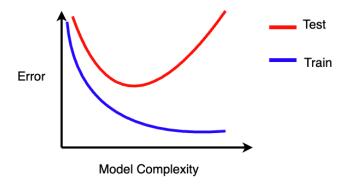
Overparameterized Linear Regression under Adversarial Attack. Antônio H. Ribeiro, Thomas B. Schön. IEEE Transactions on Signal Processing (2023)

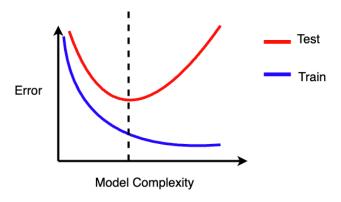
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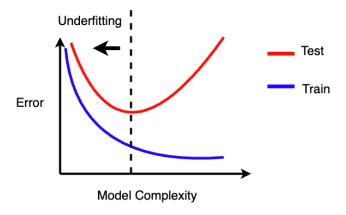
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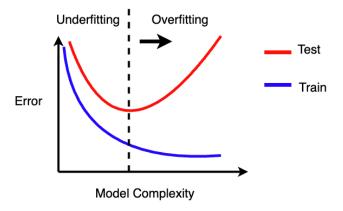
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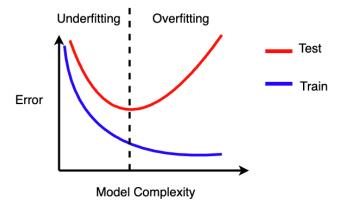
Nature Communications (2020)







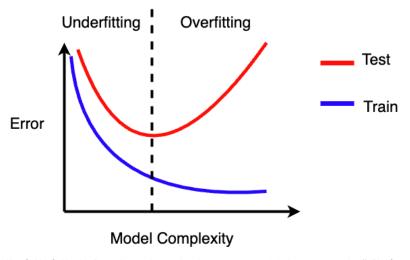




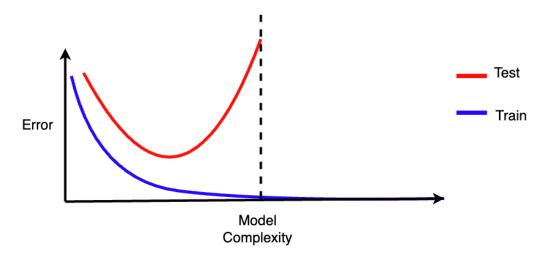
## Generalization of deep neural networks



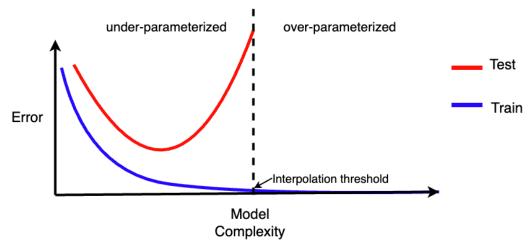
C. Zhang, S. Bengio, M. Hardt, B. Recht, and O. Vinyals. Understanding deep learning requires rethinking generalization. ICLR, 2017



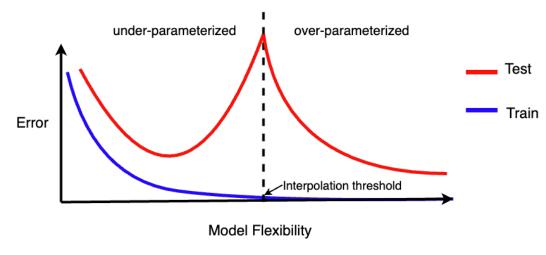
M. Belkin, D. Hsu, S. Ma, S. Mandal. Reconciling modern machine learning practice and the bias-variance trade-off. PNAS, 2020.



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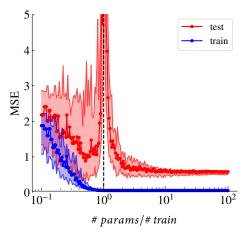
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#### Double-descent

Ph.D. seminar course: The unreasonable effectiveness of overparameterized machine learning models (3 hp), 2021

#### Double-descent

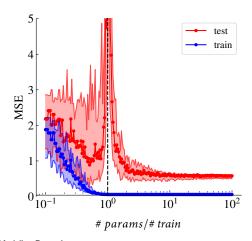
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Beyond Occam's Razor in System Identification: Double-Descent when Modeling Dynamics Antônio H. Ribeiro, Johannes N. Hendriks, Adrian G. Wills, Thomas B. Schön. IFAC Symposium on System Identification (SYSID), 2021. Honorable mention: Young author award

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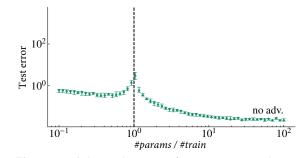
#### Deep networks for system identification: a Survey

Gianluigi Pillonetto, Aleksandr Aravkin, Daniel Gedon, Lennart Ljung, Antonio H. Ribeiro, Thomas Bo Schön. Automatica (Provisionally accepted), 2023.

Given a **test point**  $(x_0, y_0)$ , the error is:

no adversary

$$(\mathbf{y}_0 - \mathbf{\beta}^{\top} \mathbf{x}_0)^2$$



**Figure:** Adv. risk. min.  $\ell_2$ -norm interpolator

Overparameterized Linear Regression under Adversarial Attack.

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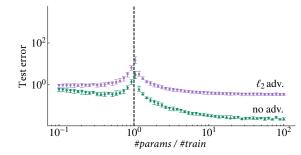
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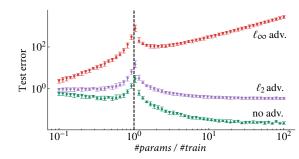
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# Overparameterized Linear Regression under Adversarial Attack

#### Interpretation

Minimum  $\ell_2$ -norm interpolation  $\Leftrightarrow \ell_2$ -adversarial training. (Result #2, Part I)

# Overparameterized Linear Regression under Adversarial Attack

#### Interpretation

Minimum  $\ell_2$ -norm interpolation  $\Leftrightarrow \ell_2$ -adversarial training. (Result #2, Part I)

#### Analysis:

- Assimptotic results showing the phenomena
- Non-asymptotic results: concentration inequalities

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JEEF Transactions on Signal Processing (2023)

## Part III. Automatic ECG analysis

# The electrocardiogram (ECG) exam

#### Cardiovascular diseases:

ightharpoonup  $\approx$ 18 million deaths in 2019 (32%).

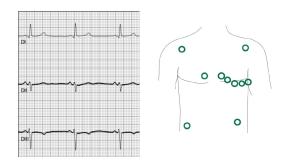
# The electrocardiogram (ECG) exam

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The ECG is the major diagnostic tool.

- ► Low-cost, safe and non-invasive
- Can detect arrhythmias, myocardial infarction, cardiomyopathy...



Left: ECG signal Right: Electrode placement.

## Computational electrocardiography



Figure Automated ECG interpretation Glasgow (1971).

Macfarlane, P.W.; Kennedy, J. "Automated ECG Interpretation—A Brief History from High Expectations to Deepest Networks." Hearts 2021.

### The transition into end-to-end learning

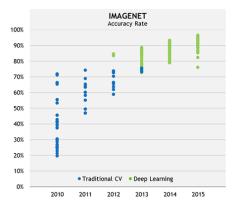


Figure: Accuracy on Imagenet as models transitioned from feature extraction to end-to-end.

J. Deng, W. Dong, R. Socher, L.-J. Li, K. Li, and L. Fei-Fei, "Imagenet: A large-scale hierarchical image database," CVPR (2009)

# Telehealth and automatic diagnosis



Figure: State of Minas Gerais

#### Telehealth and automatic diagnosis

#### Telehealth Center of Minas Gerais

- ▶ 1100 municipalities
- ightharpoonup > 3500 ECGs per day

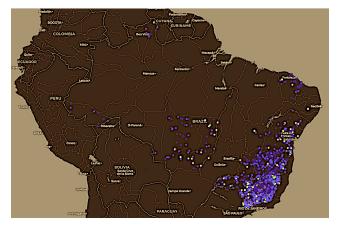
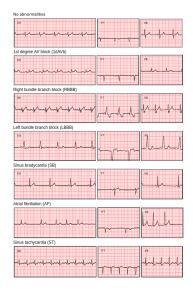


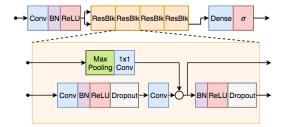
Figure: Municipalities assisted by the telehealth center

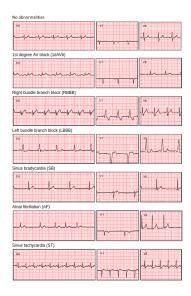
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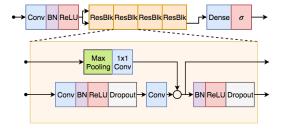


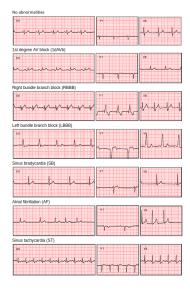
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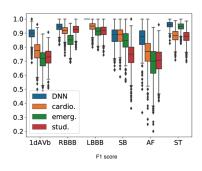




## Automatic diagnosis of the ECG (cont.)

Result: Deep neural network (DNN) performs at least as well as experts

cardio.  $\to$  4th year cardiology residents emerg.  $\to$  3rd year emergency residents stud.  $\to$  5th year Medical students

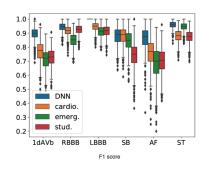


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► Goal: Improve the accuracy

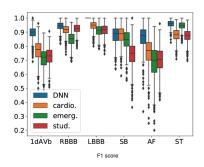


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Goal: Improve the accuracy assist more patients

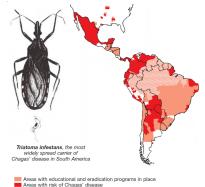


#### Three directions

- 1. Automatic diagnosis;
- 2. Screening;
- 3. Prognosis.

## Screening for Chagas disease from the ECG using deep neural networks

- **6** million people infected.
- Diagnosed with **blood test**.
- Early diagnosis and treatment halt progression.
- Low detection rates

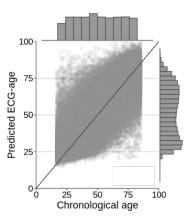


Areas with risk of Chagas' disease ### Areas with limited data on Chagas' disease and no eradication programs

#### Screening for Chagas disease from the electrocardiogram using a deep neural network

Carl Jidling, Daniel Gedon, Thomas B. Schön, Claudia Di Lorenzo Oliveira, Clareci Silva Cardoso, Ariela Mota Ferreira, Luana Giatti, Sandhi Maria Barreto, Ester C. Sabino, Antônio L. P. Ribeiro, Antônio H. Ribeiro Plos Neglected Tropical Diseases (2023)

# ECG predicted-age

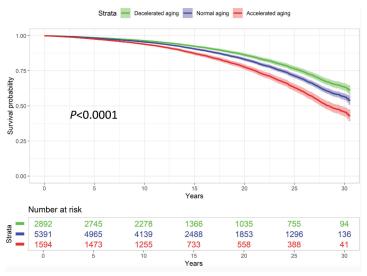


Deep neural network estimated electrocardiographic-age as a mortality predictor

Emilly M. Lima\*, Antônio H. Ribeiro\*, Gabriela MM Paixão\*, et. al. Equal contribution

Nature Communications (2021)

#### Risk predictor of cardiovascular events



Electrocardiographic Age Predicts Cardiovascular Events in Community: The Framingham Heart Study

Luisa C C Brant, **Antônio H Ribeiro**, Marcelo M Pinto-Filho, et. al. *Circulation: Cardiovascular Quality and Outcomes* (2023)

Interpretability Attempt to draw real electrocardiographic knowledge.

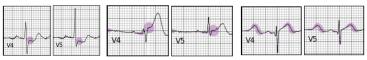


Figure: Grad-CAM plots. (Left) STEMI. (Middle) STEMI. (Right) NSTEMI.

Development and validation of deep learning ECG-based prediction of myocardial infarction in emergency department patients.

S. Gustafsson, D. Gedon, E. Lampa, Antônio H. Ribeiro, M. J. Holzmann, T. B. Schön, J. Sundström.

Scientific Reports (2022)

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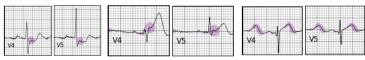


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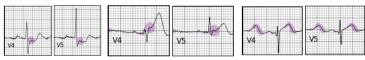


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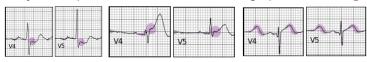


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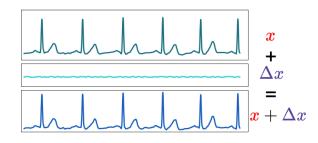
Robustness. Ability to work in real situations.

ML algorithms don't need to be really interpreable to be useful in clinical practice.

But they need to be robust!

#### Adversarial attacks in ECGs

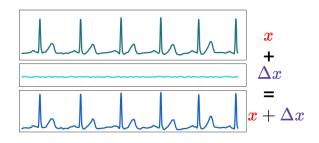
Normal (Probability = 0.99)



Han, X., Hu, Y., Foschini, L. et al. Deep learning models for electrocardiograms are susceptible to adversarial attacks. Nature Medicine. (2020)

#### Adversarial attacks in ECGs

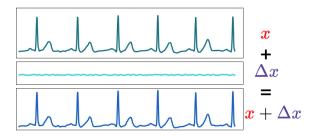
- Normal (Probability = 0.99)
- $ightharpoonup \|\Delta x\| < \delta$



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#### Adversarial attacks in ECGs

- Normal (Probability = 0.99)
- $ightharpoonup \|\Delta x\| < \delta$
- $x + \Delta x \rightarrow \widetilde{y}$ : **AFib** (Probability = 1.00)



Han, X., Hu, Y., Foschini, L. et al.Deep learning models for electrocardiograms are susceptible to adversarial attacks. Nature Medicine. (2020)

#### Conclusion

- ► Large-scale models have great potential for medicine (and critical applications).
- Robustness is a major challenge.
- Adversarial attacks framework allows for analysis of worst-case scenarios.
- Linear models for insight and analysis.
- Adversarially-trained linear regression is a competitive regression method.

#### Thank you!

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