



Elektrokardiogram a umelá inteligencia



ROBERT HATALA

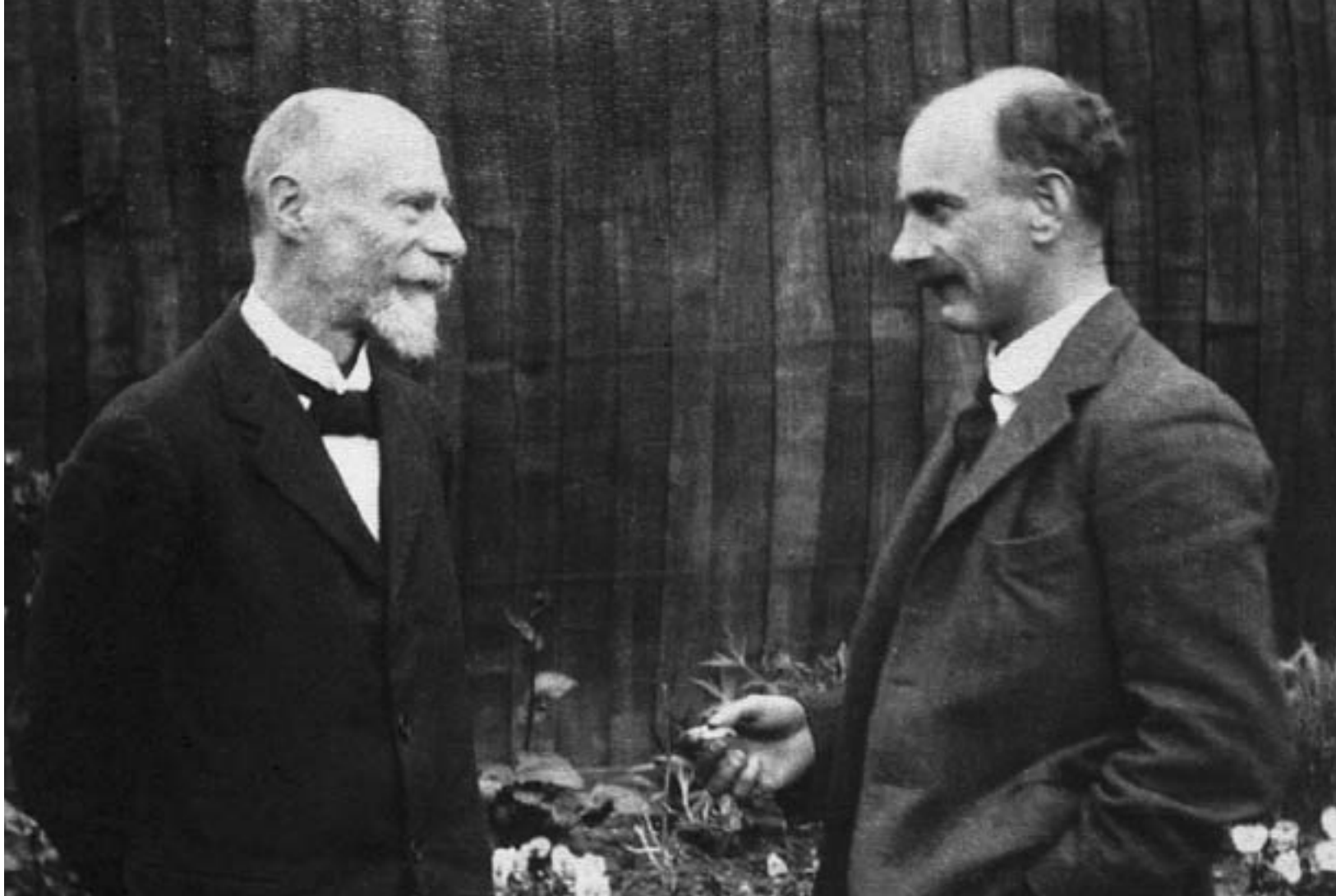
*Klinika kardiológie a angiológie
NÚSCH a.s. a LF SZU Bratislava*



Deklarace konfliktu zájmů

	Nemám konflikt zájmů	Mám konflikt zájmů	Specifikace konfliktu (vyjmenujte subjekty, firmy či instituce, se kterými Vaše spolupráce může vést ke konfliktu zájmů)
Zaměstnanecký poměr	X		
Vlastník / akcionář	X		
Konzultant	X		
Přednášková činnost	X		
Člen poradních sborů (advisory boards)	X		
Podpora výzkumu / granty	X		
Jiné honoráře (např. za klinické studie či registry)	X		

Willem Einthoven and Thomas Lewis Leiden, April 10th 1921



120 years of electrocardiography

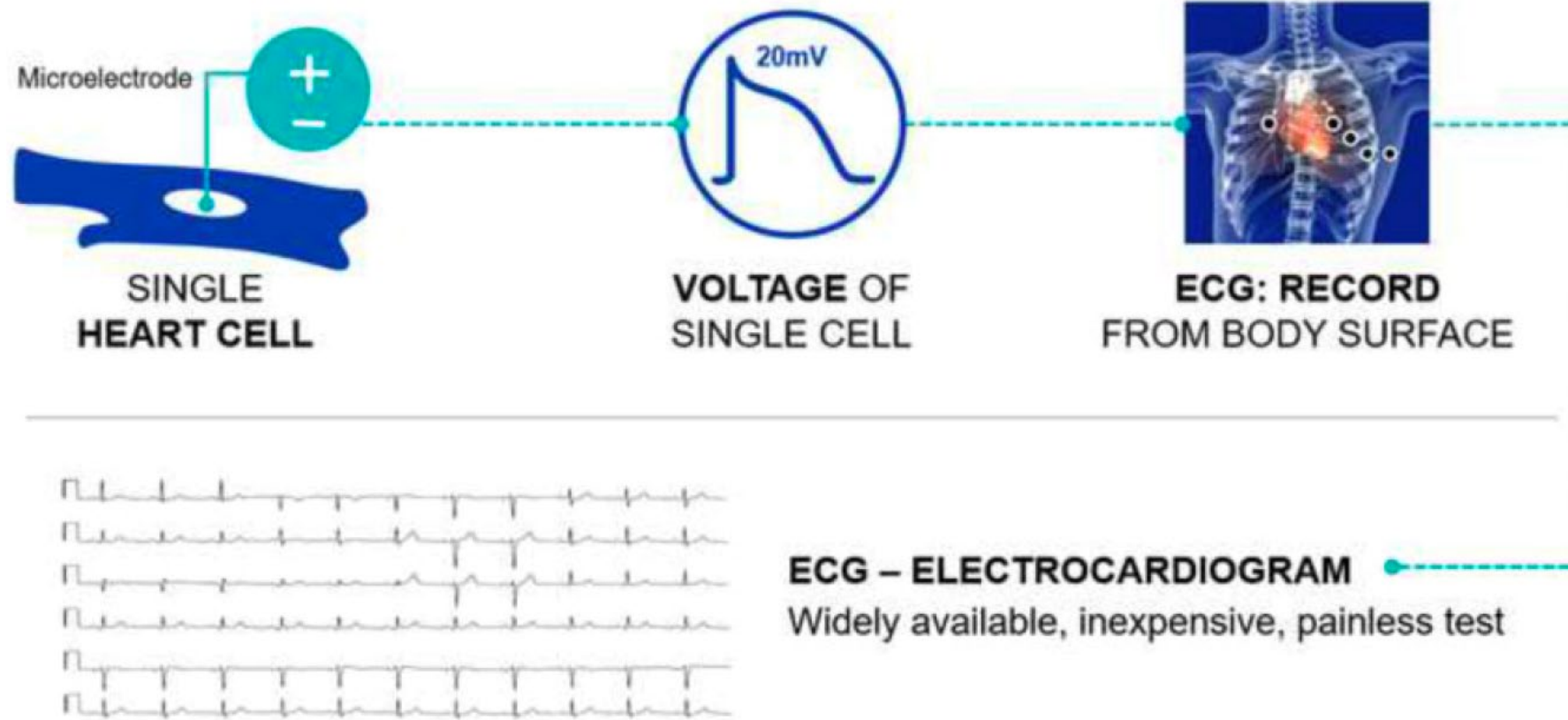
ECG – ubiquitous diagnostic tool in clinical medicine

- low cost, rapid, simple to obtain
- available at point-of-care

Inherent contradiction within electrocardiography

- Acquisition is standardized and highly reproducible
- Interpretation is highly variable - according to variable human expertise

EKG - bohatý zdroj fyziologických informácií, ktoré reflektujú stav srdca i iných systémov



Aims to eliminate the variability of ECG interpretation are not new

- **Computer based interpretation**
 - Applying predefined rules based on previous experience derived from experimental and clinical observation
- **Artificial intelligence**
 - Deep learning - neural networks
 - CNNs – **convolutional neural networks - „image recognition“**

ČÍM VIAC INFORMÁCII O ELEKTRICKOM POLI SRDCA
MÁME, TÝM LEPŠIE MÔŽEME DIAGNOSTIKOVAŤ
KARDIÁLNE OCHORENIA.

ALE: ODKIAĽ ČERPAŤ INFORMÁCIE?
EXTENZÍVNY VERSUS INTENZÍVNY PRÍSTUP

Computers for ECG interpretation

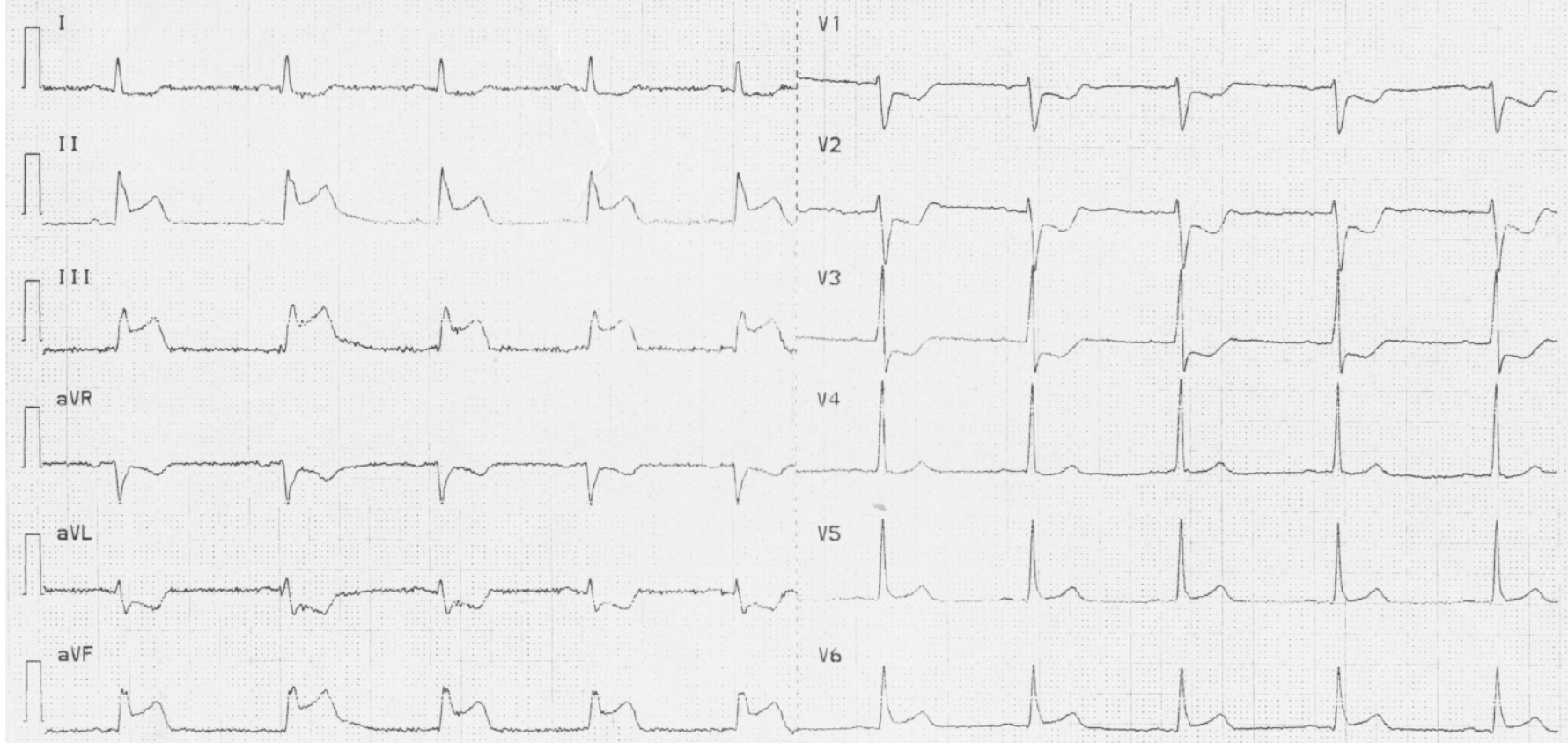
1. Mimicking human interpretation by applying predefined **rules** (traditional computer interpretation) = i.e. computer searches for digital counterparts of visible ecg changes.
2. **Artificial intelligence (AI)** - extracting information beyond the level of recognition by humans
3. **Expanding the number of leads** to increase precision.



Vent. Rate:	60 bpm
RR Interval:	988 ms
PR Interval:	168 ms
QRS Duration:	116 ms
QT Interval:	406 ms
QTc Interval:	406 ms
QT Dispersion:	56 ms
P-R-T AXIS:	-18° 66° 98°

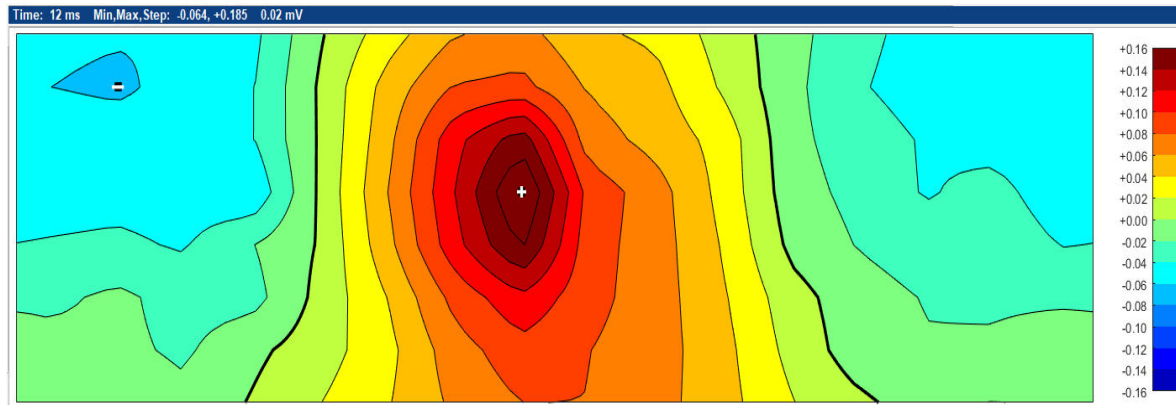
SINUS RHYTHM
 ** INTERPRETATION MADE WITHOUT KNOWING PATIENT'S GENDER/AGE
 Low maximum QRS spatial velocity
 Broad QRS in (V1 or V2) and (V5 or V6)
 INCOMPLETE LEFT BUNDLE BRANCH BLOCK
 Inferior ST elevation
 CONSIDER ACUTE INFERIOR INFARCTION
 Anterolateral ST-T changes
 STRONGLY SUGGESTS MYOCARDIAL INJURY/ISCHEMIA

Summary: ABNORMAL ECG * Unconfirmed Analysis

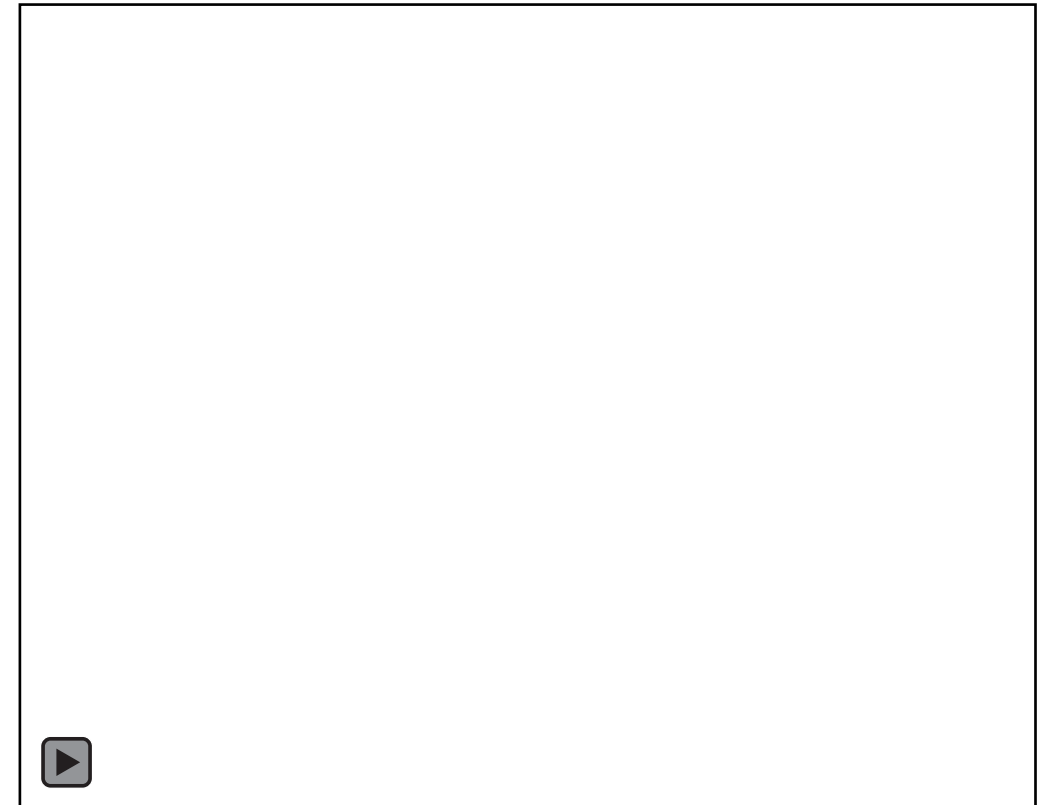


Noninvasive PVC localization

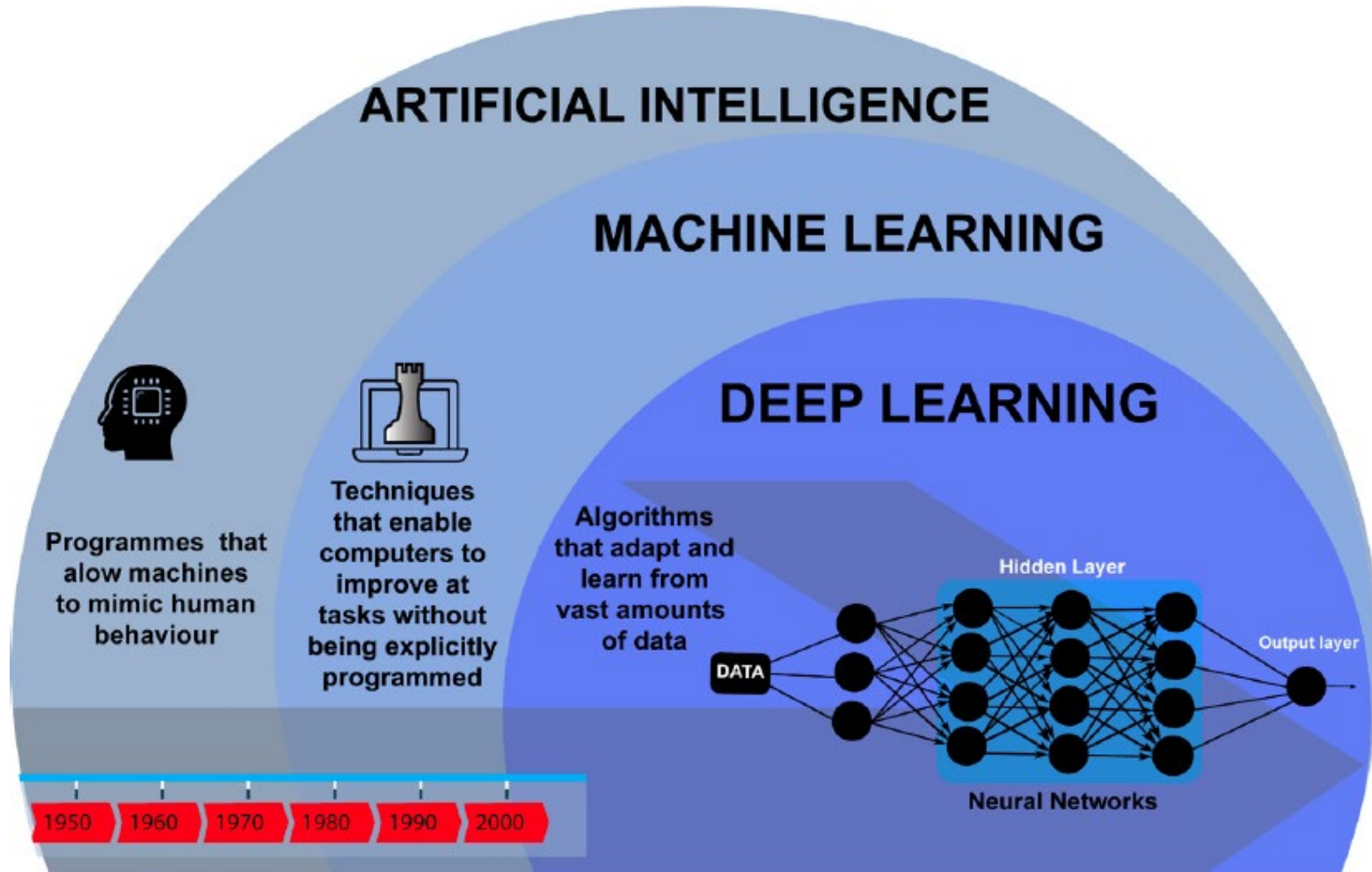
- Patient P034 with implanted pacemaker. During spontaneous activity the PVC focus could be localized in the basal lateral segment of the left ventricle.



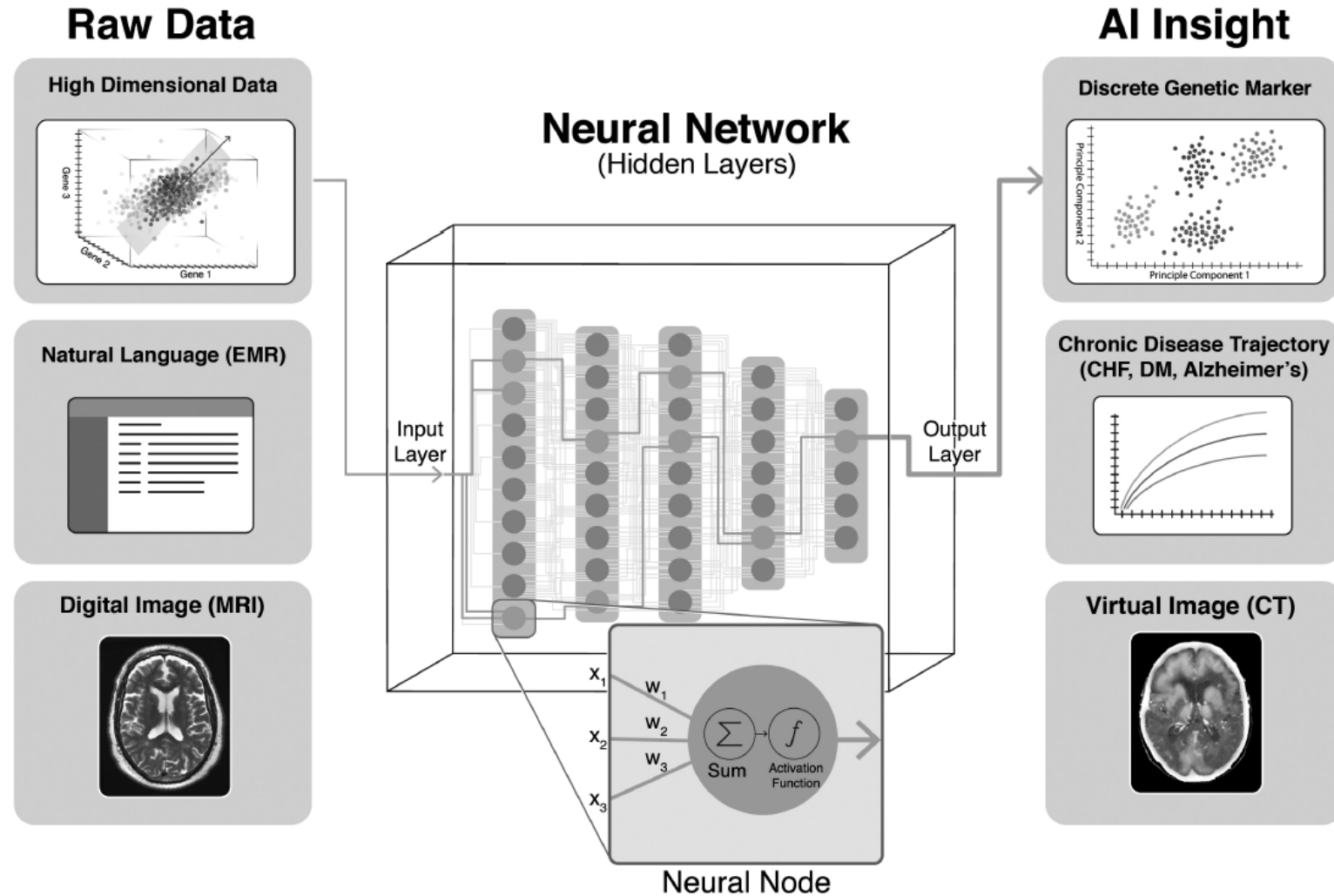
The PVC focus was localized from a BSPM 12 ms after the PVC onset.



SVET UMELEJ INTELEGENCIJE



Machine learning (AI) by an artificial neural network





Artificial intelligence

- ML aims to characterise **relationships among multiple variables** or features within data sets that are not readily discernible to humans and difficult to discover using traditional biostatistical methods
- ML can only yield **associations and correlations**; currently it cannot make definitive cause–effect inferences.
- The principal tasks of ML comprise: **classification** (e.g. distinguishing disease from non-disease); **prediction** (or regression) (e.g. estimating risk of future clinical events); and **discovery** (e.g. a new use for a drug or a new disease phenotype).

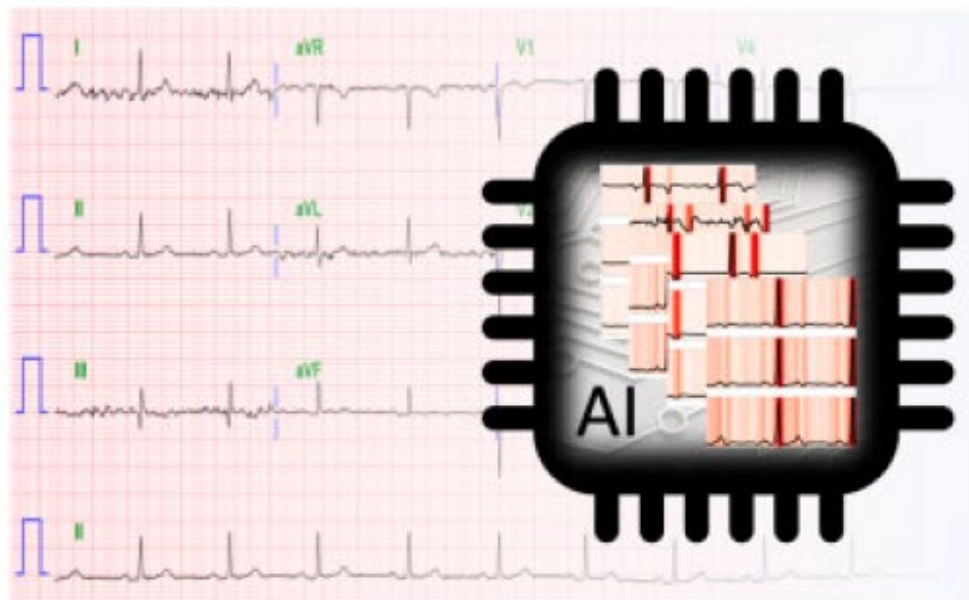
ECG = ideal target for applying AI and deep learning

- Widely available digital data
- Easily stored and transferred
- Huge datasets available
- Powerful computing readily available

STROJOVÉ UČENIE (MACHINE LEARNING)

- Strojové učenie, jeden z druhov umelej inteligencie, funguje na základe identifikácie vzorov v dostupných údajoch a následného uplatnenia týchto znalostí na nové údaje . Čím väčší je súbor údajov, tým ľahšie je možné odhaliť aj nepatrné súvislosti.
- Významné pokroky v týchto technológiách sa dosiahli použitím **veľkých súborov údajov (“big data”)** a **nevidanej výpočtovej sily**.

Umelá inteligencia v elektrokardiografii



Diagnosis

- Left ventricular systolic dysfunction
- Heart failure with preserved ejection fraction
- Aortic valve stenosis
- Mitral valve regurgitation
- Pulmonary hypertension
- Left ventricular hypertrophy
- Myocardial infarction with or without ST elevation
- Arrhythmia
- Hyperkalemia
- Anemia

Prediction

- Paroxysmal atrial fibrillation
- Patient deterioration and cardiac arrest
- Aortic valve stenosis
- Heart failure with preserved ejection fraction
- Mitral valve regurgitation

Agreement Between Artificial Neural Networks and Experienced Electrocardiographer on Electrocardiographic Diagnosis of Healed Myocardial Infarction

BO HEDÉN, MD, MATTIAS OHLSSON, PhD, RALF RITTNER, MSc, OLLE PAHLM, MD, PhD, WESLEY K. HAISTY, JR., MD,* CARSTEN PETERSON, PhD, LARS EDENBRANDT, MD, PhD

Lund, Sweden and Winston-Salem, North Carolina

Objectives. The purpose of this study was to compare the diagnoses of healed myocardial infarction made from the 12-lead electrocardiogram (ECG) by artificial neural networks and an experienced electrocardiographer.

Background. Artificial neural networks have proved of value in pattern recognition tasks. Studies of their utility in ECG interpretation have shown performance exceeding that of conventional ECG interpretation programs. The latter present verbal statements, often with an indication of the likelihood for a certain diagnosis, such as "possible left ventricular hypertrophy." A neural network presents its output as a numeric value between 0 and 1; however, these values can be interpreted as Bayesian probabilities.

Methods. The study was based on 351 healthy volunteers and 1,313 patients with a history of chest pain who had undergone

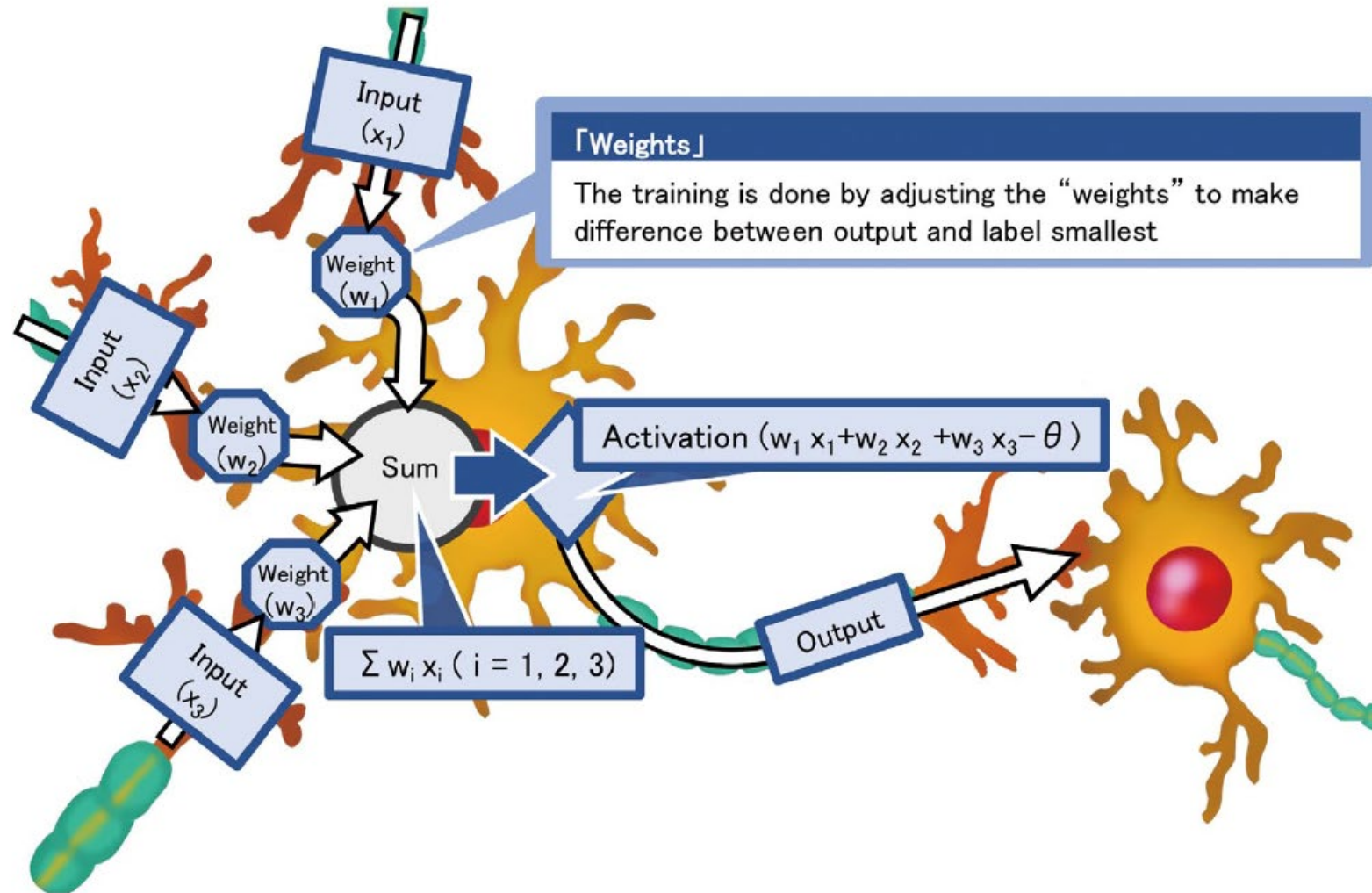
diagnostic cardiac catheterization. A 12-lead ECG was recorded in each subject. An expert electrocardiographer classified the ECGs in five different groups by estimating the probability of anterior myocardial infarction. Artificial neural networks were trained and tested to diagnose anterior myocardial infarction. The network outputs were divided into five groups by using the output values and four thresholds between 0 and 1.

Results. The neural networks diagnosed healed anterior myocardial infarctions at high levels of sensitivity and specificity. The network outputs were transformed to verbal statements, and the agreement between these probability estimates and those of an expert electrocardiographer was high.

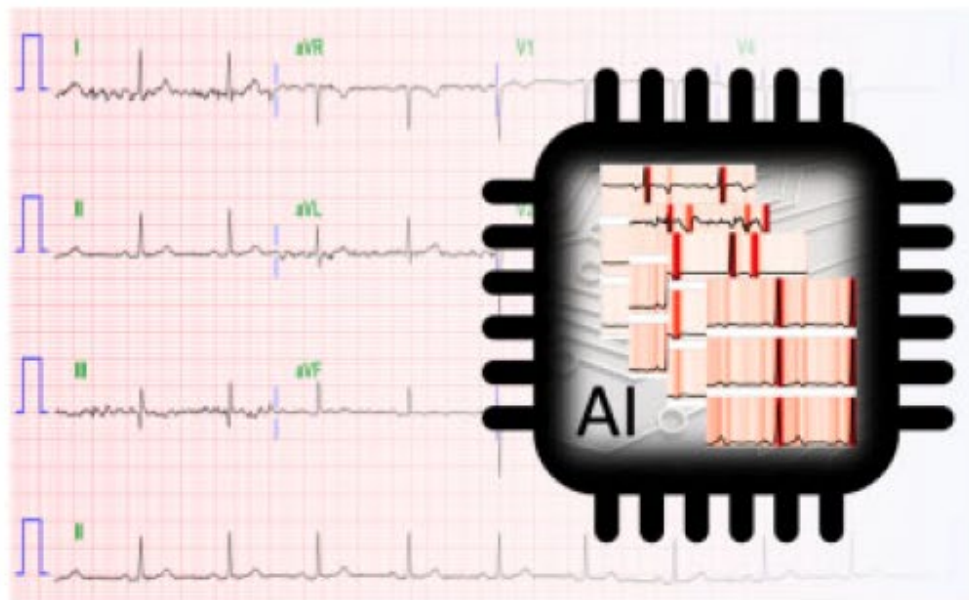
Conclusions. Artificial neural networks can be of value in automated interpretation of ECGs in the near future.

(J Am Coll Cardiol 1996;28:1012-6)

Neural network is constructed by multiple units of statistical model called “neuron” that simulates the function of neuron cells



Umelá inteligencia v elektrokardiografii



Diagnosis

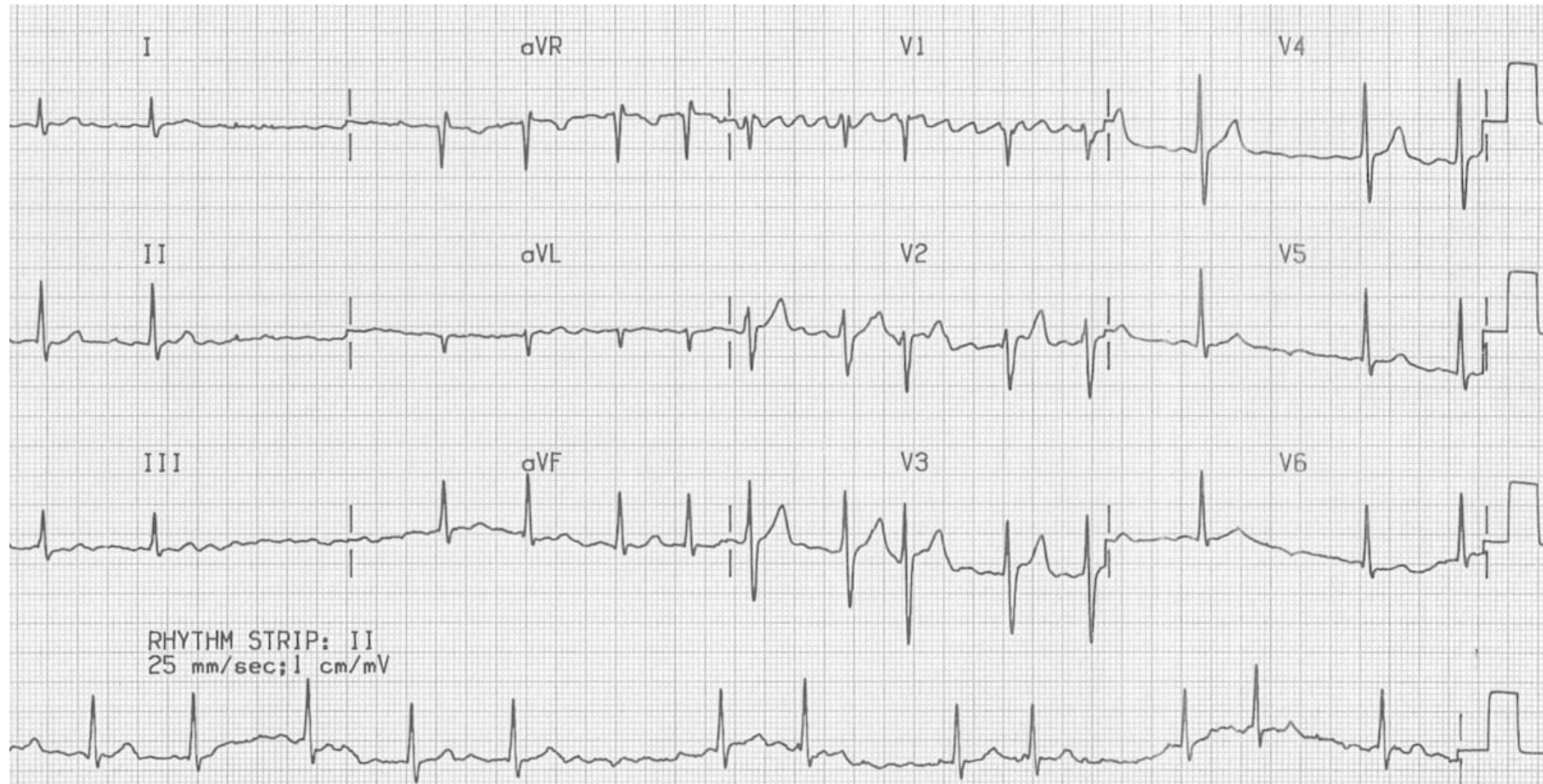
- Left ventricular systolic dysfunction
- Heart failure with preserved ejection fraction
- Aortic valve stenosis
- Mitral valve regurgitation
- Pulmonary hypertension
- Left ventricular hypertrophy
- Myocardial infarction with or without ST elevation
- Arrhythmia
- Hyperkalemia
- Anemia

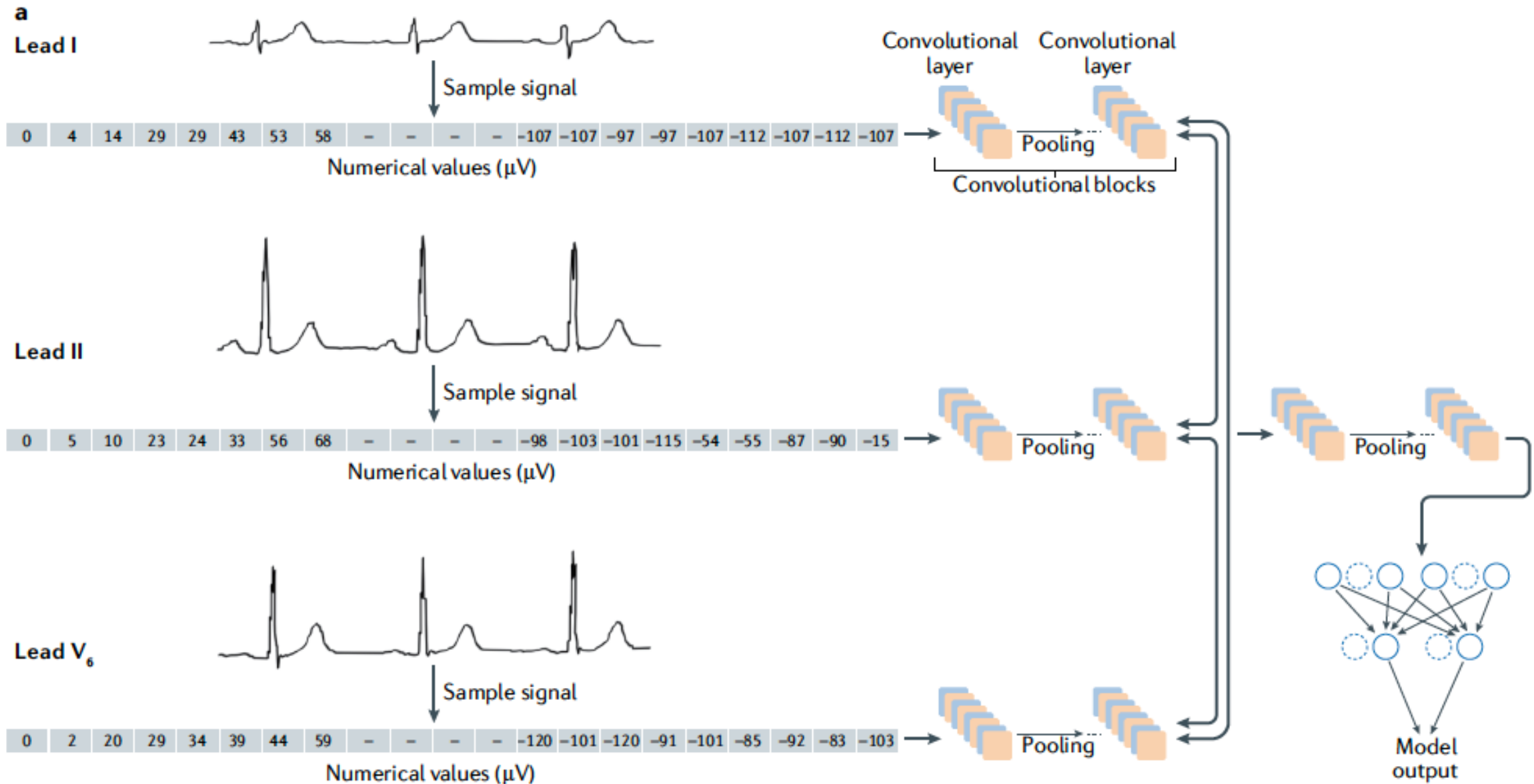
Prediction

- Paroxysmal atrial fibrillation
- Patient deterioration and cardiac arrest
- Aortic valve stenosis
- Heart failure with preserved ejection fraction
- Mitral valve regurgitation

ELEKTRICKÉ OCHORENIA SRDCA

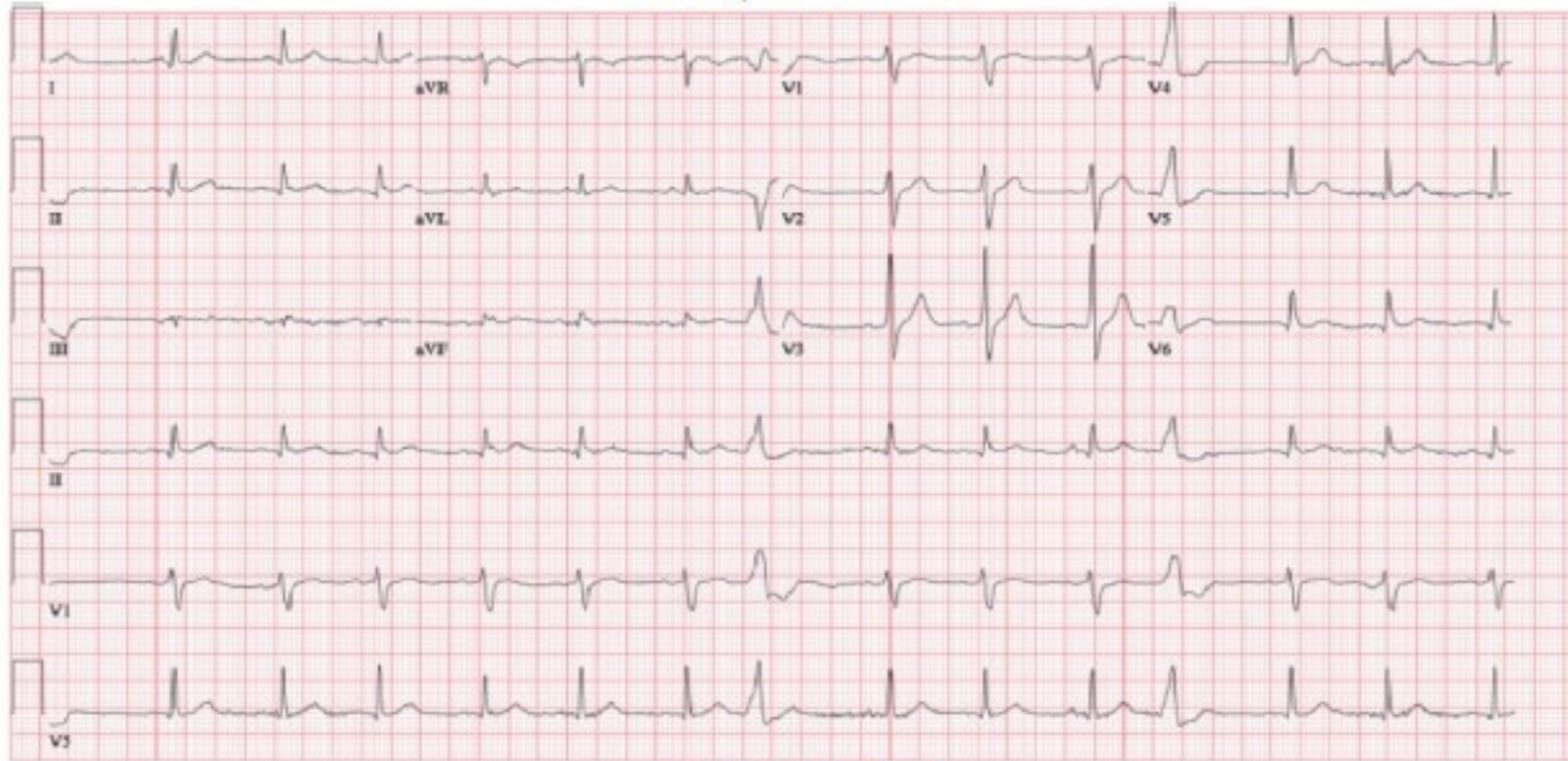
ATRIAL FIBRILLATION





b

ECG acquired over 10 s



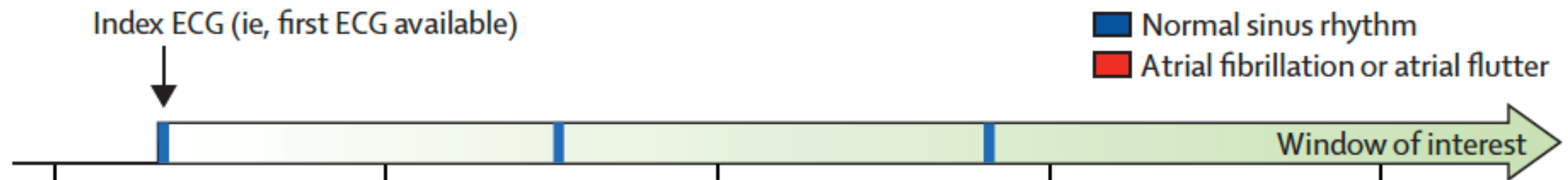
AI-ECG analysis

Atrial fibrillation detected over a longer period than the ECG

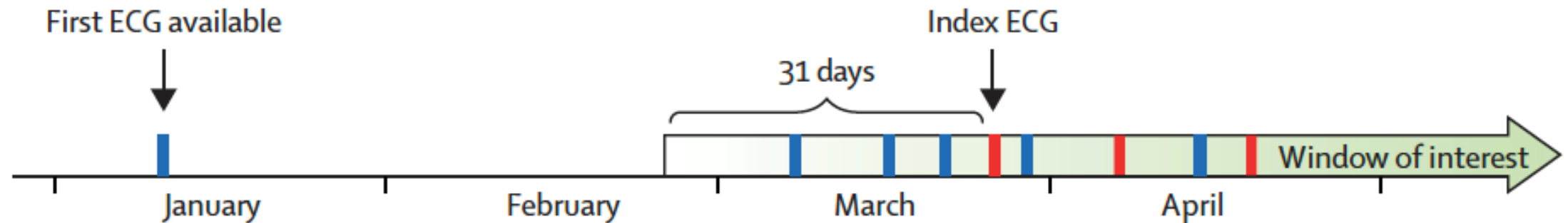
An artificial intelligence-enabled ECG algorithm for the identification of patients with atrial fibrillation during sinus rhythm: a retrospective analysis of outcome prediction

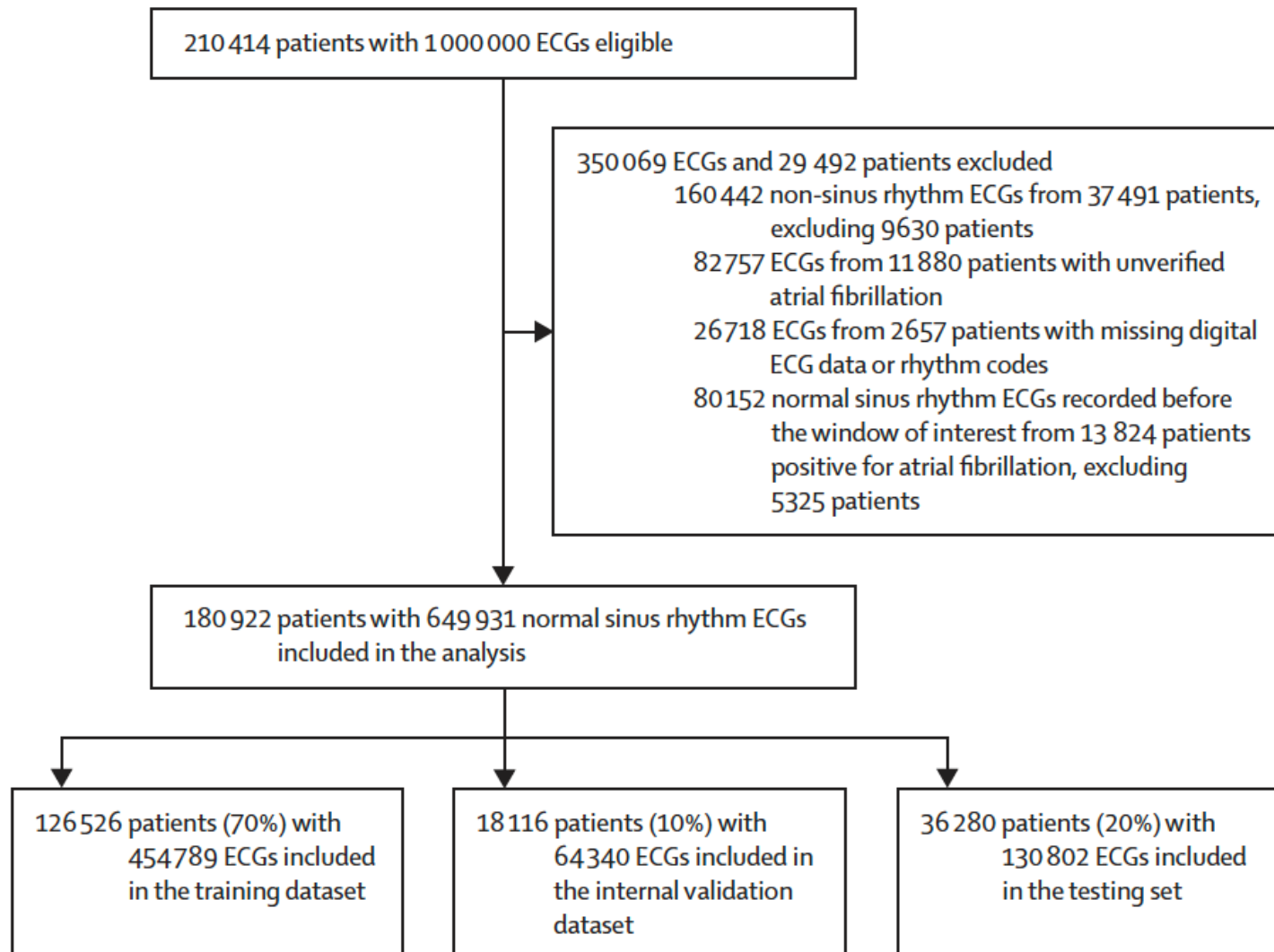
Zachil Attia*, Peter A Noseworthy*, Francisco Lopez-Jimenez, Samuel J Asirvatham, Abhishek J Deshmukh, Bernard J Gersh, Rickey E Carter, Xiaoxi Yao, Alejandro A Rabinstein, Brad J Erickson, Suraj Kapa, Paul A Friedman

Patient with no atrial fibrillation rhythms recorded



Patient with at least one atrial fibrillation rhythm recorded





An artificial intelligence-enabled ECG algorithm for the identification of patients with atrial fibrillation during sinus rhythm: a retrospective analysis of outcome prediction

Zachil Attia*, Peter A Noseworthy*, Francisco Lopez-Jimenez, Samuel J Asirvatham, Abhishek J Deshmukh, Bernard J Gersh, Rickey E Carter, Xiaoxi Yao, Alejandro A Rabinstein, Brad J Erickson, Suraj Kapa, Paul A Friedman

An artificial intelligence-enabled ECG algorithm for the identification of patients with atrial fibrillation during sinus rhythm: a retrospective analysis of outcome prediction

Zachi I Attia*, Peter A Noseworthy*, Francisco Lopez-Jimenez, Samuel J Asirvatham, Abhishek J Deshmukh, Bernard J Gersh, Rickey E Carter, Xiaoxi Yao, Alejandro A Rabinstein, Brad J Erickson, Suraj Kapa, Paul A Friedman

	AUC	Sensitivity	Specificity	F1 score	Accuracy
Main analysis	0.87 (0.86–0.88)	79.0% (77.5–80.4)	79.5% (79.0–79.9)	39.2% (38.1–40.3)	79.4% (79.0–79.9)
Secondary analysis	0.90 (0.90–0.91)	82.3% (80.9–83.6)	83.4% (83.0–83.8)	45.4% (44.2–46.5)	83.3% (83.0–83.7)




Data in parentheses are 95% CIs. In the main analysis, only the score of the first normal sinus rhythm ECG in the window of interest was used. In the secondary analysis, the highest score for all ECGs done in the first month of the window of interest was used. AUC=area under the curve. ECG=electrocardiograph.

“Torsade de Points” Tachycardia in long QT syndrome



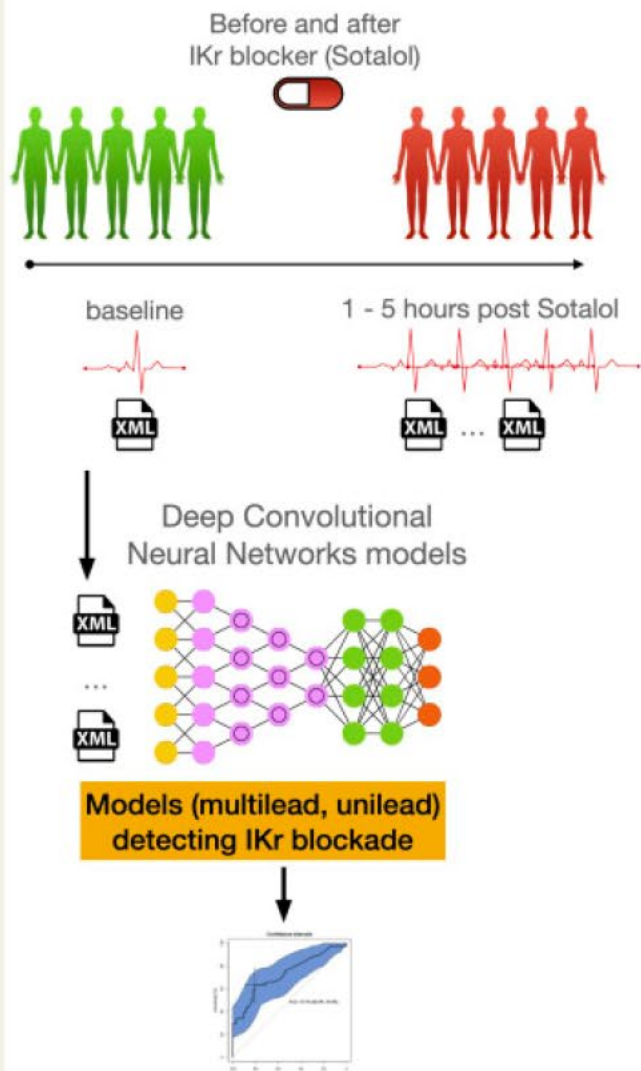
LONG QT SYNDROME

Figure 2: Characteristics of LQTS 1–3

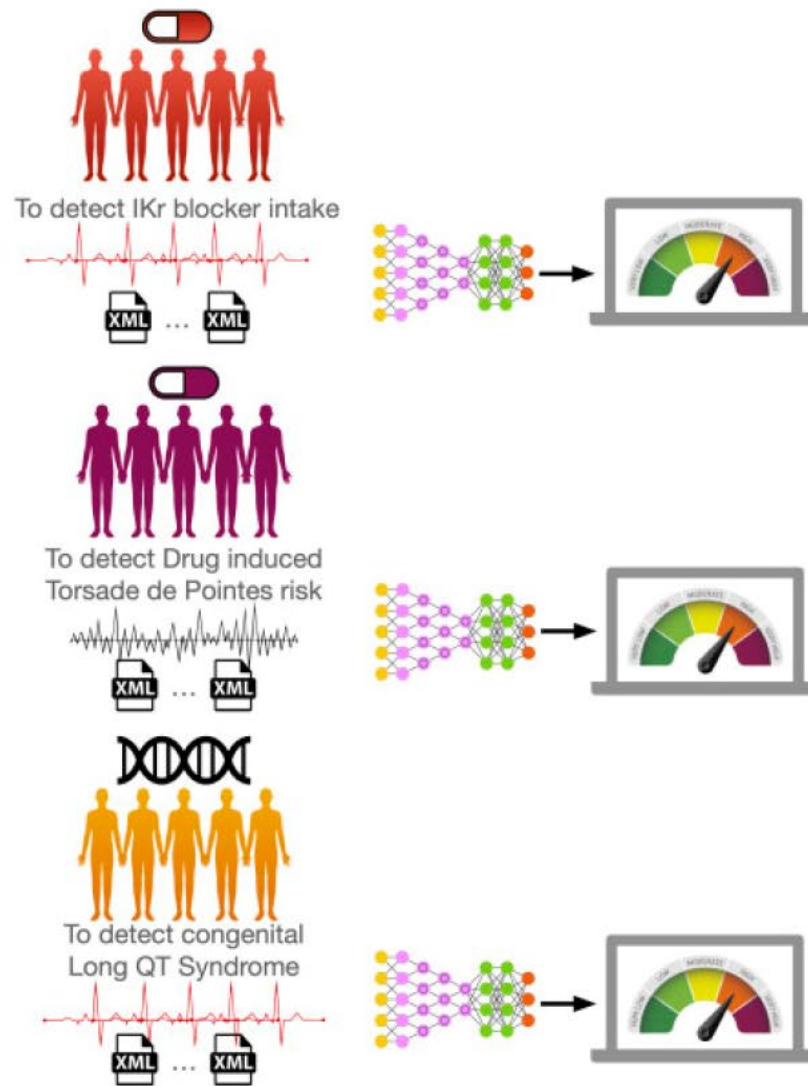
Type	Current	Functional Effect	Frequency Among LQTS	ECG	Triggers Lethal Cardiac Event	Penetrance*
LQTS1	K	↓	30%-35%		Exercise (68%) Emotional stress (14%) Sleep, response (9%) Others (19%)	62%
LQTS2	K	↓	25%-30%		Exercise (29%) Emotional stress (49%) Sleep, response (22%)	75%
LQTS3	Na	↑	5%-10%		Exercise (4%) Emotional stress (12%) Sleep, response (64%) Others (20%)	90%

Source: Adapted from Rev Esp Cardiol. 2007;60(7):739-5. Published with permission of Elsevier España.

Training artificial intelligence models

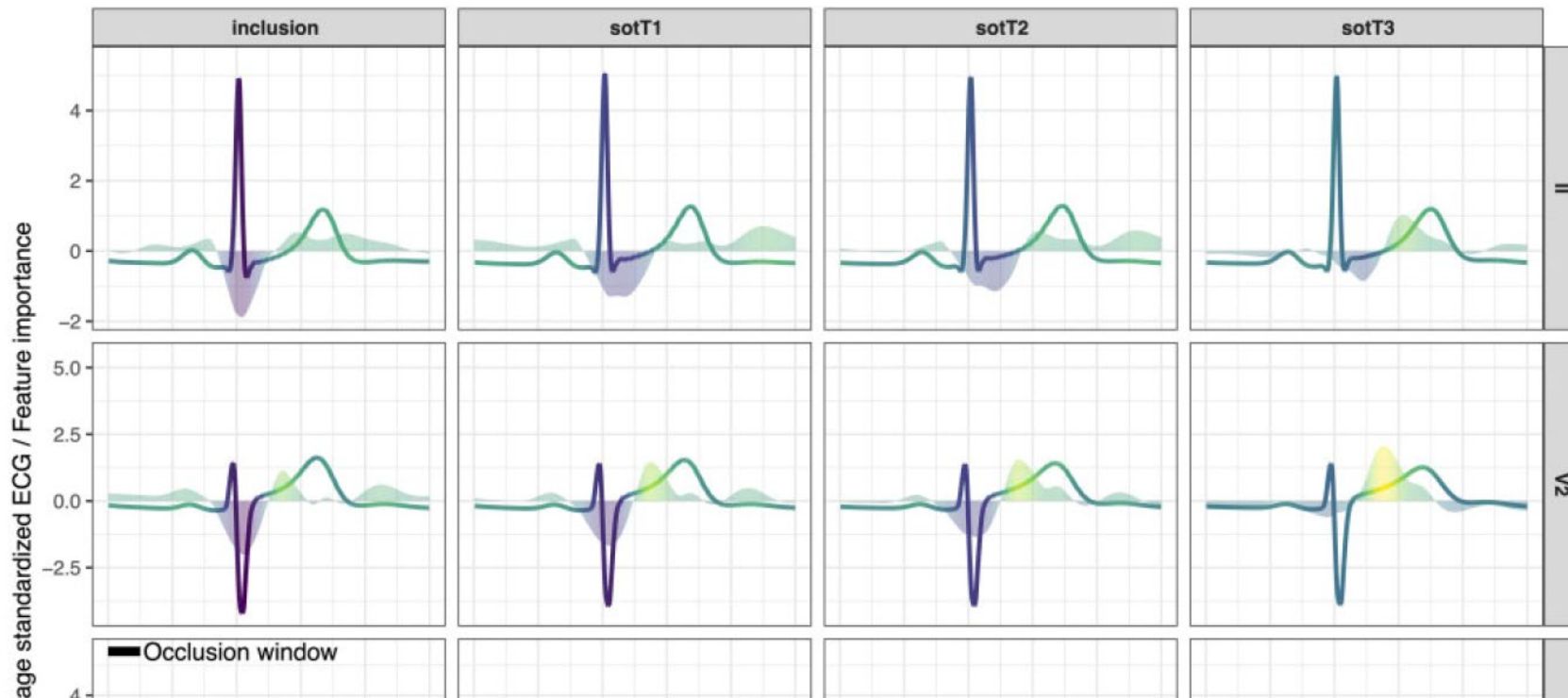


Applying artificial intelligence models



Deep learning analysis of electrocardiogram for risk prediction of drug-induced arrhythmias and diagnosis of long QT syndrome





Profil významnosti jednotlivých charakteristik pre identifikáciu ekg stopy sotalolu (feature importance profile)

CNN sa “naučila” rozoznávať ekg zmeny (často subtilne, presahujúce schopnosť ľudského posúdenia) indukované sotalolom ako modelom blokády kanála Ikr. To je rozhodujúci mechanizmus predĺženia intervalu QT, čo predisponuje ku vzniku TdP.

Mapa senzitivity pre určenie ekg segmentu rozhodujúceho pre dg. aortálnej stenózy



Use of Artificial Intelligence and Deep Neural Networks in Evaluation of Patients With Electrocardiographically Concealed Long QT Syndrome From the Surface 12-Lead Electrocardiogram

J. Martijn Bos, MD, PhD; Zachi I. Attia, PhD; David E. Albert, MD; Peter A. Noseworthy, MD; Paul A. Friedman, MD; Michael J. Ackerman, MD, PhD

- the AI-ECG was found to distinguish patients with electrocardiographically concealed LQTS
- provide a nearly 80% accurate **pregenetic** test anticipation of LQTS genotype status.
- For a cut-off of QTc > 500ms, a strong diagnostic and risk marker for the likelihood of LQTS, the area under the curve (AUC) was 0.97, with sensitivity and specificity of 80.0%, and 94.4%, respectively, indicating strong utility as a **screening method**.

DYSFUNKCIA ĽAVEJ KOMORY

A QRS SCORING SYSTEM FOR ASSESSING LEFT VENTRICULAR FUNCTION AFTER MYOCARDIAL INFARCTION

SEBASTIAN T. PALMERI, M.D., DAVID G. HARRISON, M.D., FREDERICK R. COBB, M.D.,
KENNETH G. MORRIS, M.D., FRANK E. HARRELL, PH.D., RAYMOND E. IDEKER, M.D., PH.D.,
RONALD H. SELVESTER, M.D., AND GALEN S. WAGNER, M.D.

Abstract A QRS scoring system for estimating the size of a myocardial infarct was evaluated in 55 patients who did not have left ventricular hypertrophy or conduction abnormalities. Serial 12-lead surface electrocardiograms were scored according to a 29-point system based on the duration of Q and R waves and on the ratios of R-to-Q amplitude and R-to-S amplitude. The scores were proportional to the severity of wall-motion abnormalities, which was determined by radionuclide blood-pool scanning and which correlated inversely with the radionuclide-

determined left ventricular ejection fraction (LVEF). A score >3 was 93 per cent sensitive and 88 per cent specific for both severe regional dyssynergy and major depression of the global LVEF. The following equation was used to estimate the LVEF from the QRS score: $LVEF (\%) = 60 - (3 \times QRS \text{ score})$.

After acute myocardial infarction, an electrocardiogram can provide important indirect quantitative information about left ventricular function. (N Engl J Med. 1982; 306:4-9.)

COHORT OF 84 PATIENTS

Table 1. Criteria for Determining Point Score in the QRS Scoring System.⁹

LEAD	CRITERIA (NO. OF POINTS)		MAXIMUM POINTS
	DURATION	AMPLITUDE RATIO	
I	Q > 30 msec (1)	R/Q < 1 (1)	2
II	Q > 40 msec (2) Q > 30 msec (1)		2
aVL	Q > 30 msec (1)	R/Q < 1 (1)	2
aVF	Q > 50 msec (3) Q > 40 msec (2) Q > 30 msec (1)	R/Q < 1 (2) R/Q < 2 (1)	5
V ₁	Any Q (1) R > 50 msec (2) R > 40 msec (1)	R/S > 1 (1)	4
V ₂	Any Q or R < 20 msec (1) R > 60 msec (2) R > 50 msec (1)	R/S > 1.5 (1)	4
V ₃	Any Q or R < 30 msec (1)		1
V ₄	Q > 20 msec (1)	R/Q or R/S < 0.5 (2) R/Q or R/S < 1 (1)	3
V ₅	Q > 30 msec (1)	R/Q or R/S < 1 (2) R/Q or R/S < 2 (1)	3
V ₆	Q > 30 msec (1)	R/Q or R/S < 1 (2) R/Q or R/S < 3 (1)	3

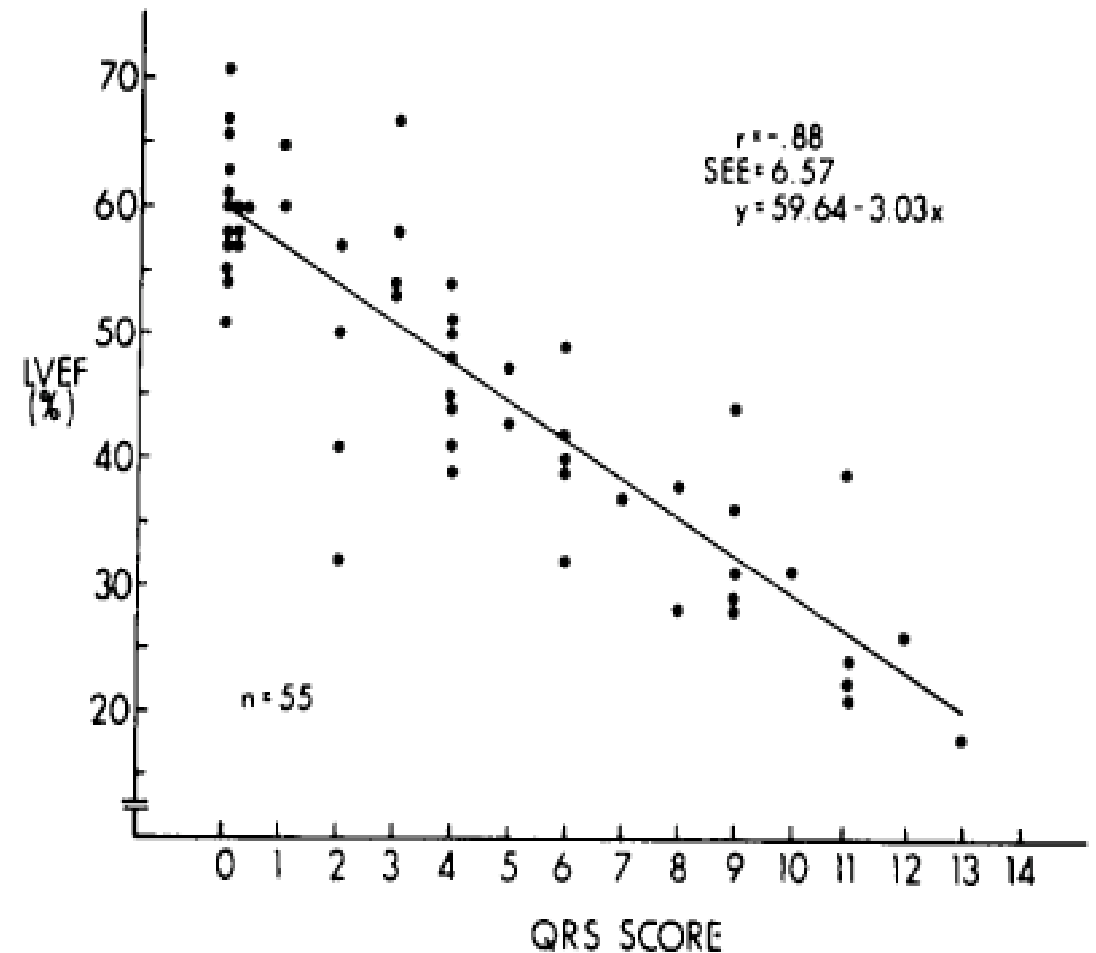
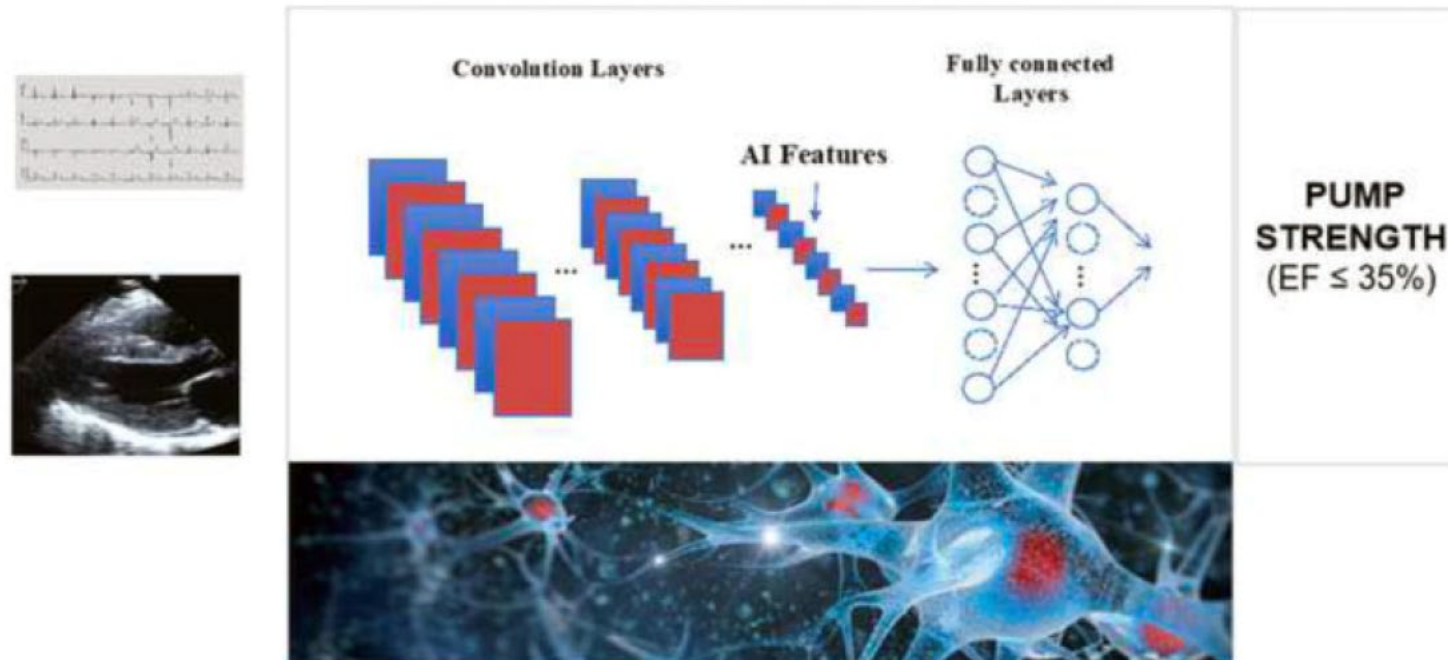


Figure 1. Correlation between QRS Score and LVEF (Determined by Radionuclide Angiography) Three Weeks after Infarction.

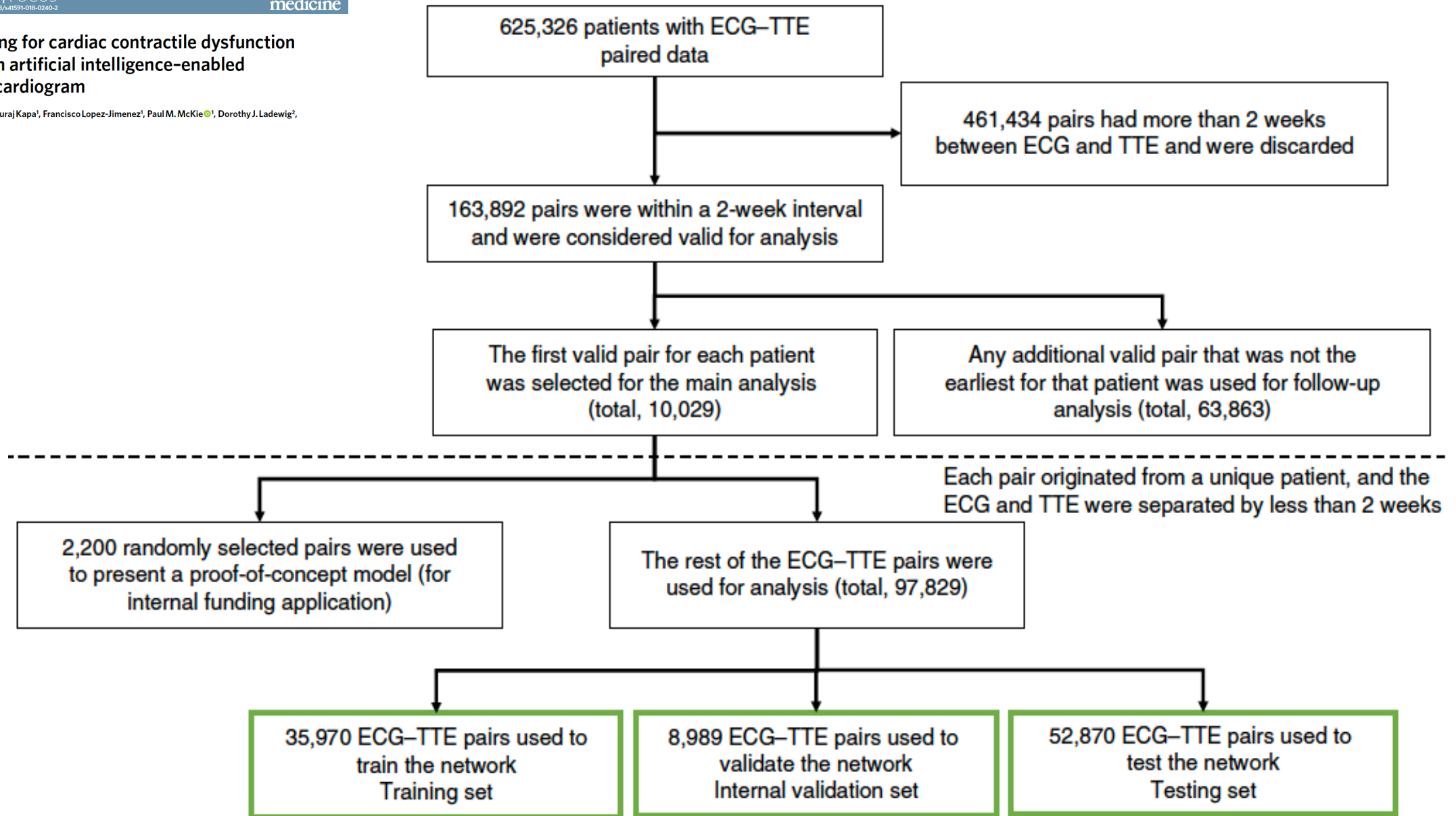
Konvolučná neuronálna sieť (CNN) s “učením pod dohľadom” (supervised learning)

ROBUST DIGITAL WAREHOUSE OF MEDICAL INFORMATION CONVOLUTIONAL NEURAL NETWORK



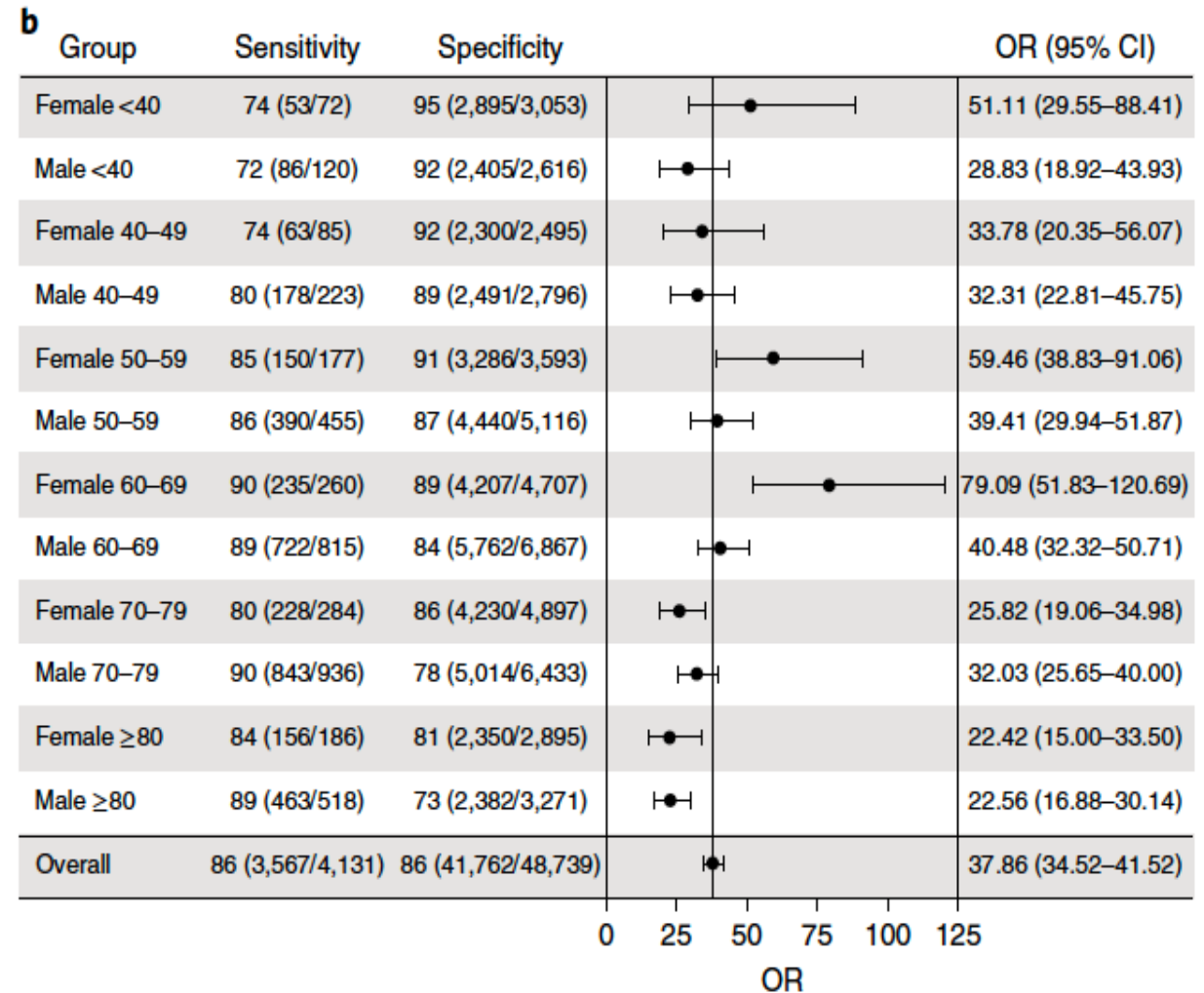
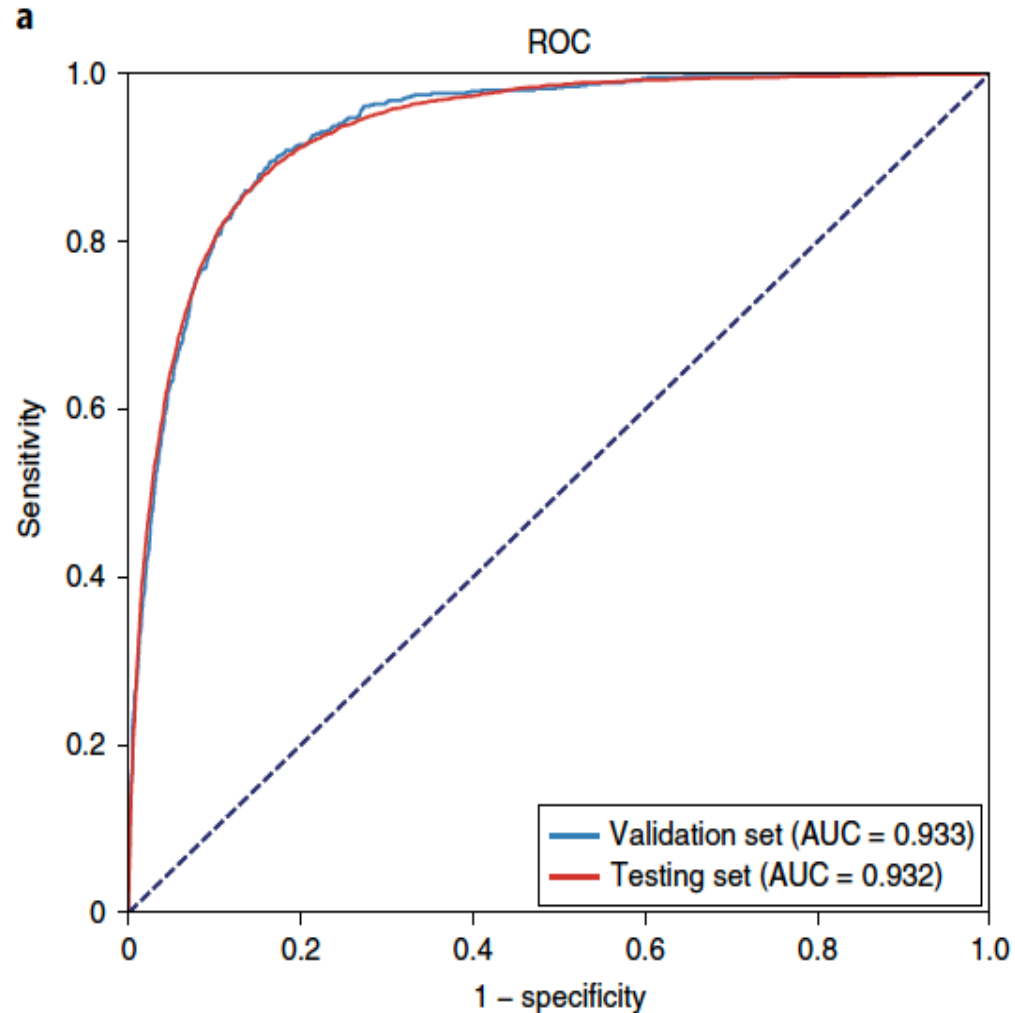
Screening for cardiac contractile dysfunction using an artificial intelligence-enabled electrocardiogram

Zachil I. Attia¹, Suraj Kapa¹, Francisco Lopez-Jimenez¹, Paul M. McKie¹, Dorothy J. Ladewig²

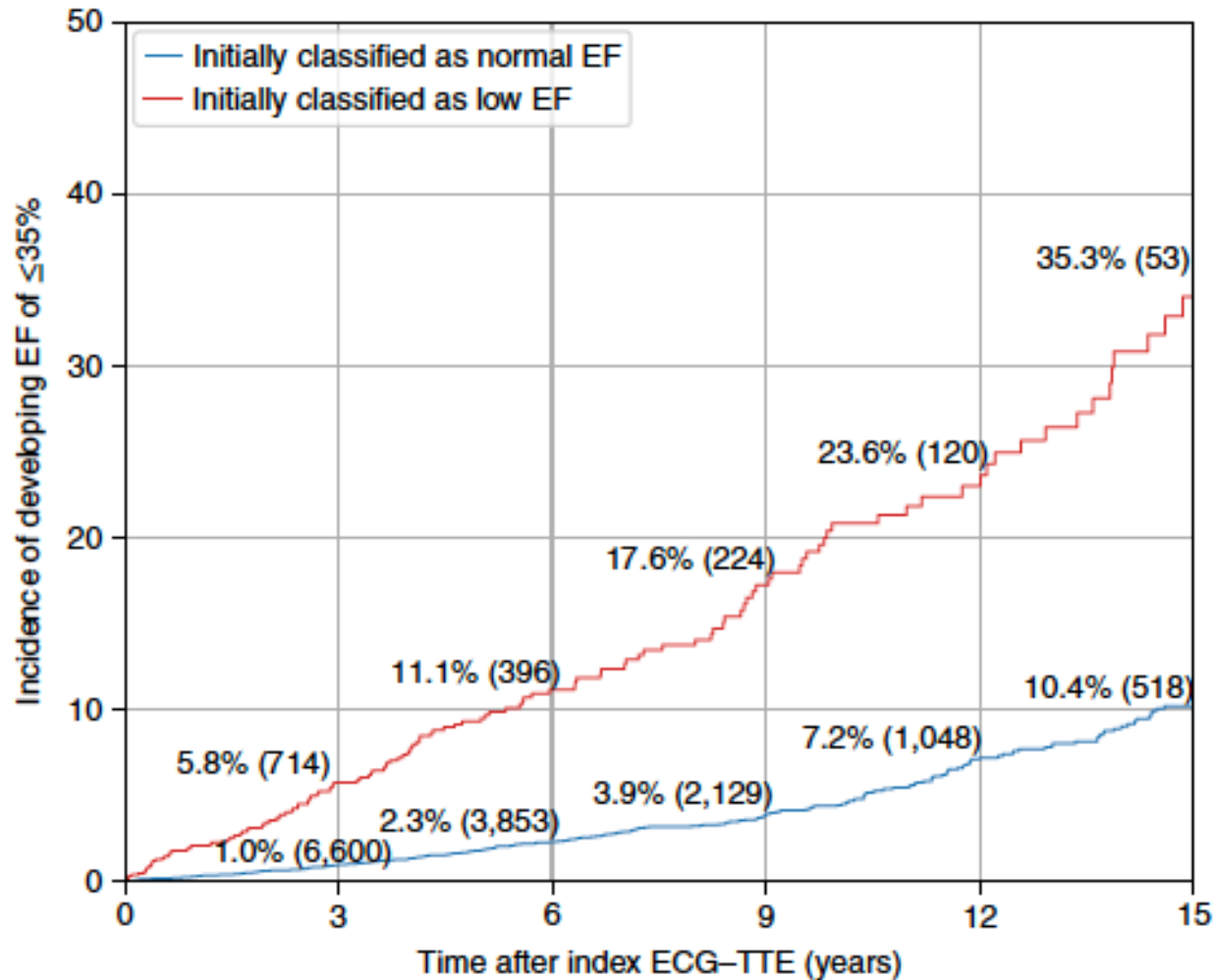


Screening for cardiac contractile dysfunction using an artificial intelligence-enabled electrocardiogram

Zachi I. Attia¹, Suraj Kapa¹, Francisco Lopez-Jimenez², Paul M. McKie¹, Dorothy J. Ladewig²



Long-term incidence of developing an EF of $\leq 35\%$ in patients with an initially normal EF stratified by AI classification



Screening for cardiac contractile dysfunction using an artificial intelligence-enabled electrocardiogram

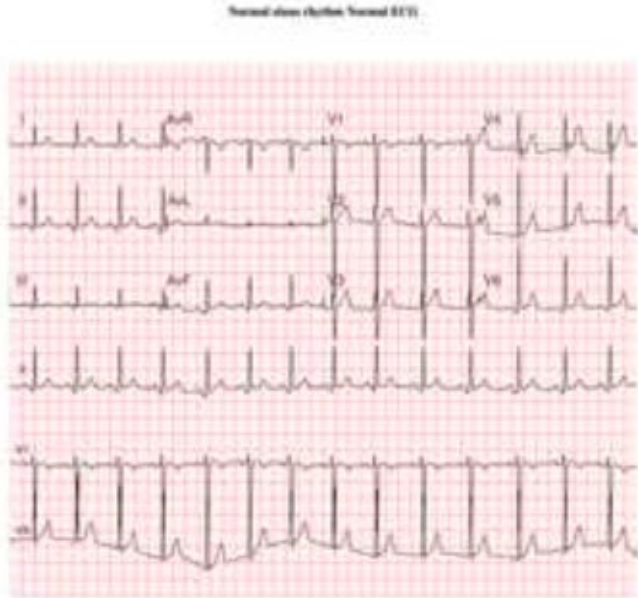
Zachi I. Attia¹, Suraj Kapa¹, Francisco Lopez-Jimenez¹, Paul M. McKie¹, Dorothy J. Ladewig²,

HR for developing low LVEF with abnormal AI-ECG = 4.1 (CI 3.3-5.0)

Applying AI permits the ECG to serve as a powerful tool:

- to screen for heart failure with LV dysfunction
- to identify individuals at increased risk for its development in the future.

AI ECG: Positive for Low EF

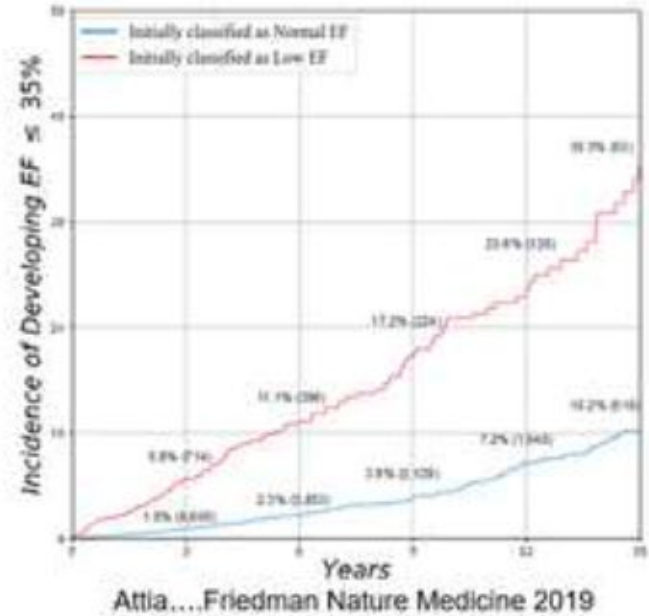


Echo Ejection Fraction: 50%
Apparent False Positive

AI ECG: Positive for Low EF



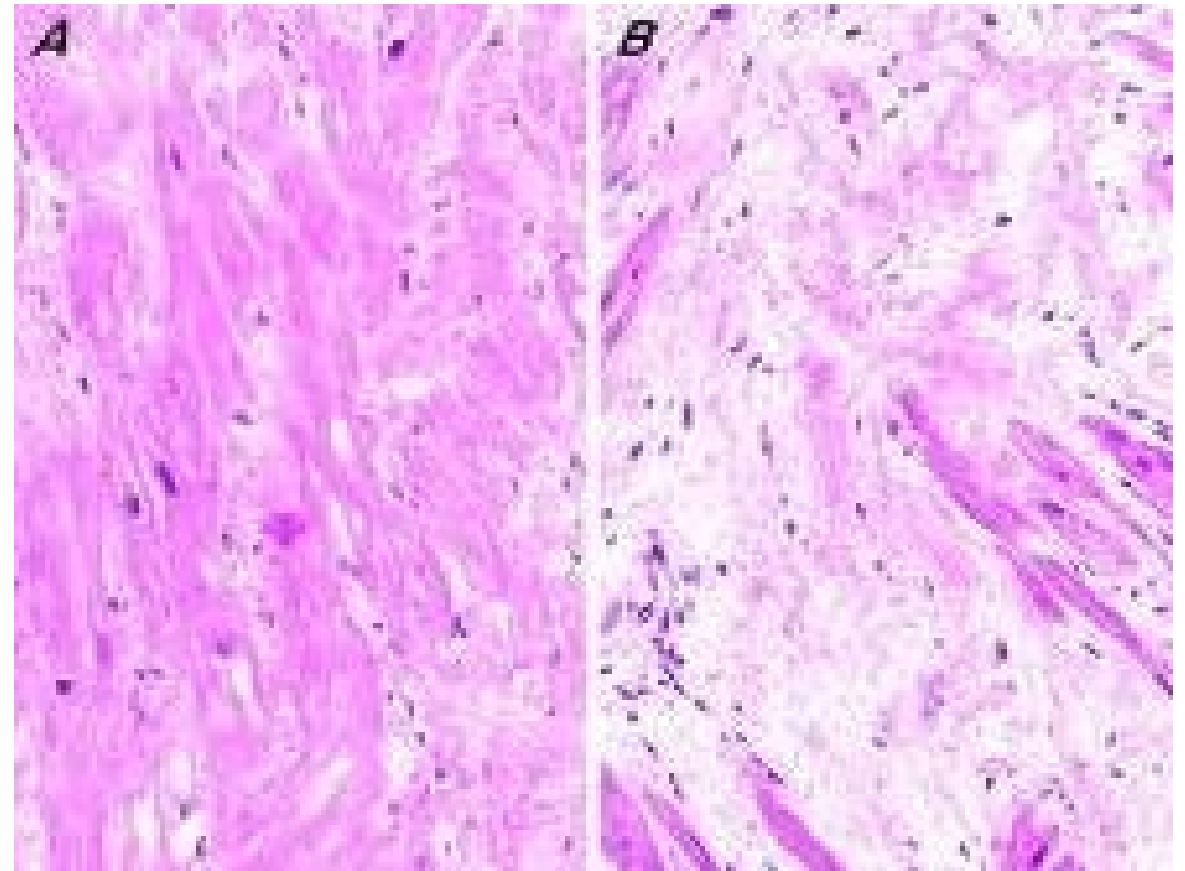
Echo Ejection Fraction 5 years later (age 33) : **31%**



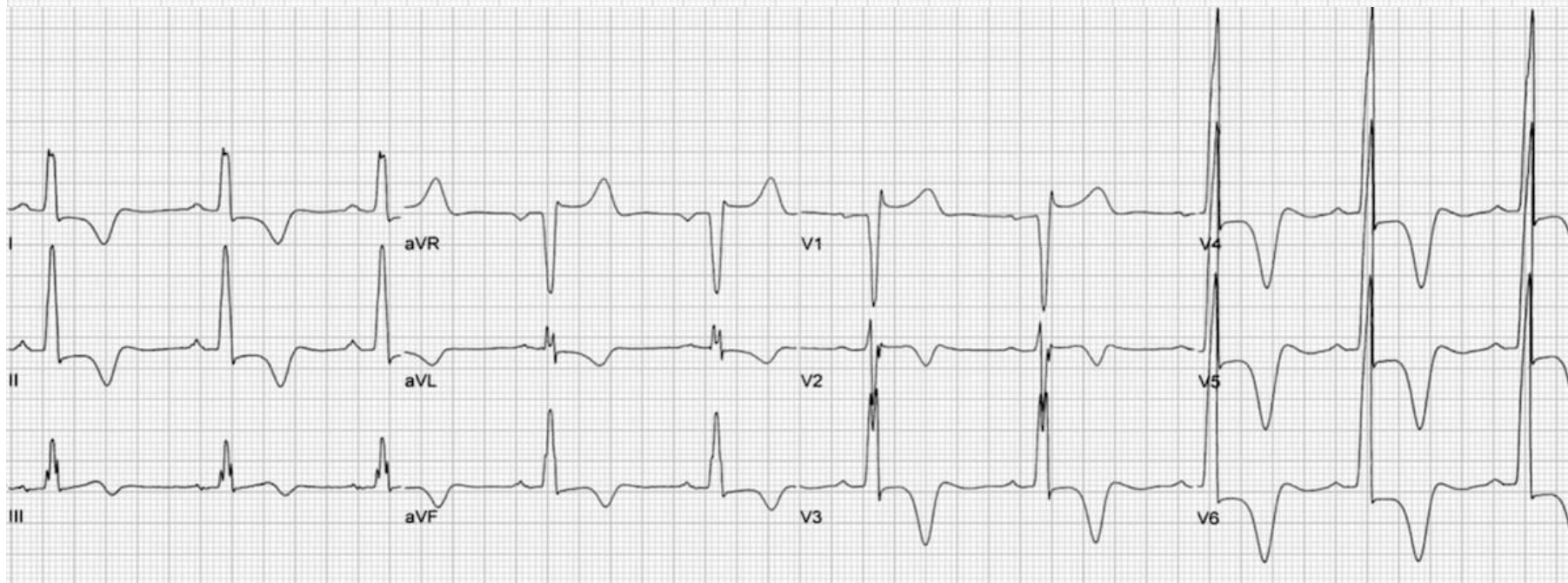
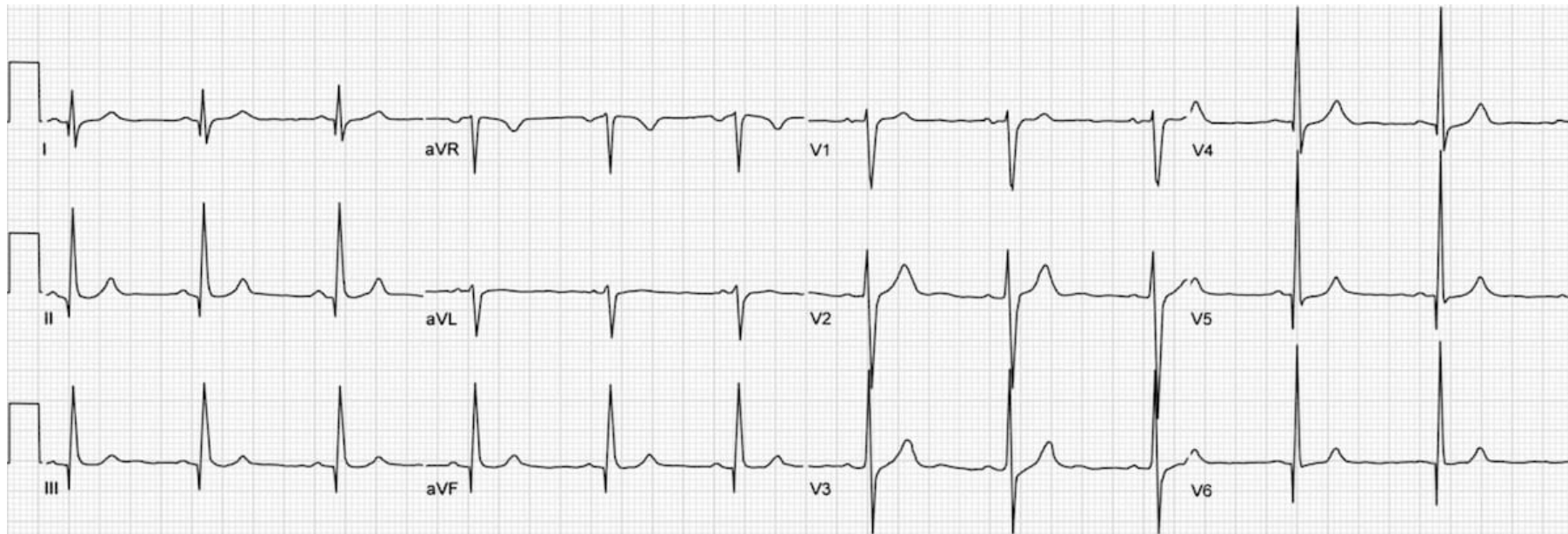
AI disease “previvor:”
 disease detected
 before manifest

ŠTRUKTURÁLNE OCHORENIA SRDCA

Patologic findings in HCMP



Maron BJ et al. NEJM 1987

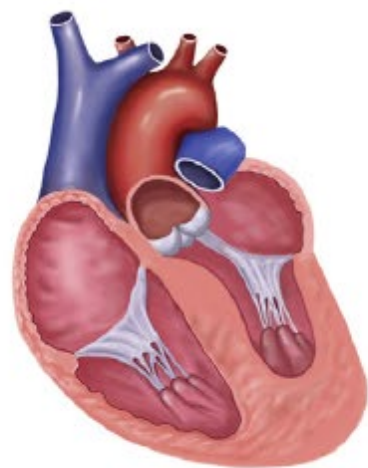


ECG in HCM

Study Population

Convolutional Neural Network Development for HCM Detection by ECG

Model Performance Characteristics



ECGs on 3,060 Patients With
HCM and 63,941 Controls

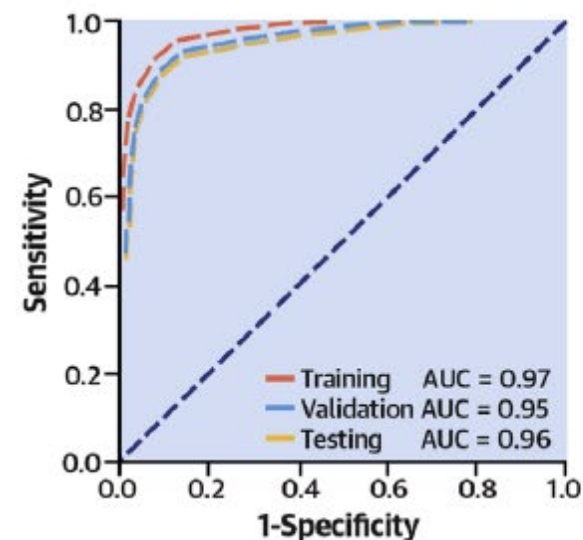


Three Non-overlapping Cohorts:

Training
N = 46,901

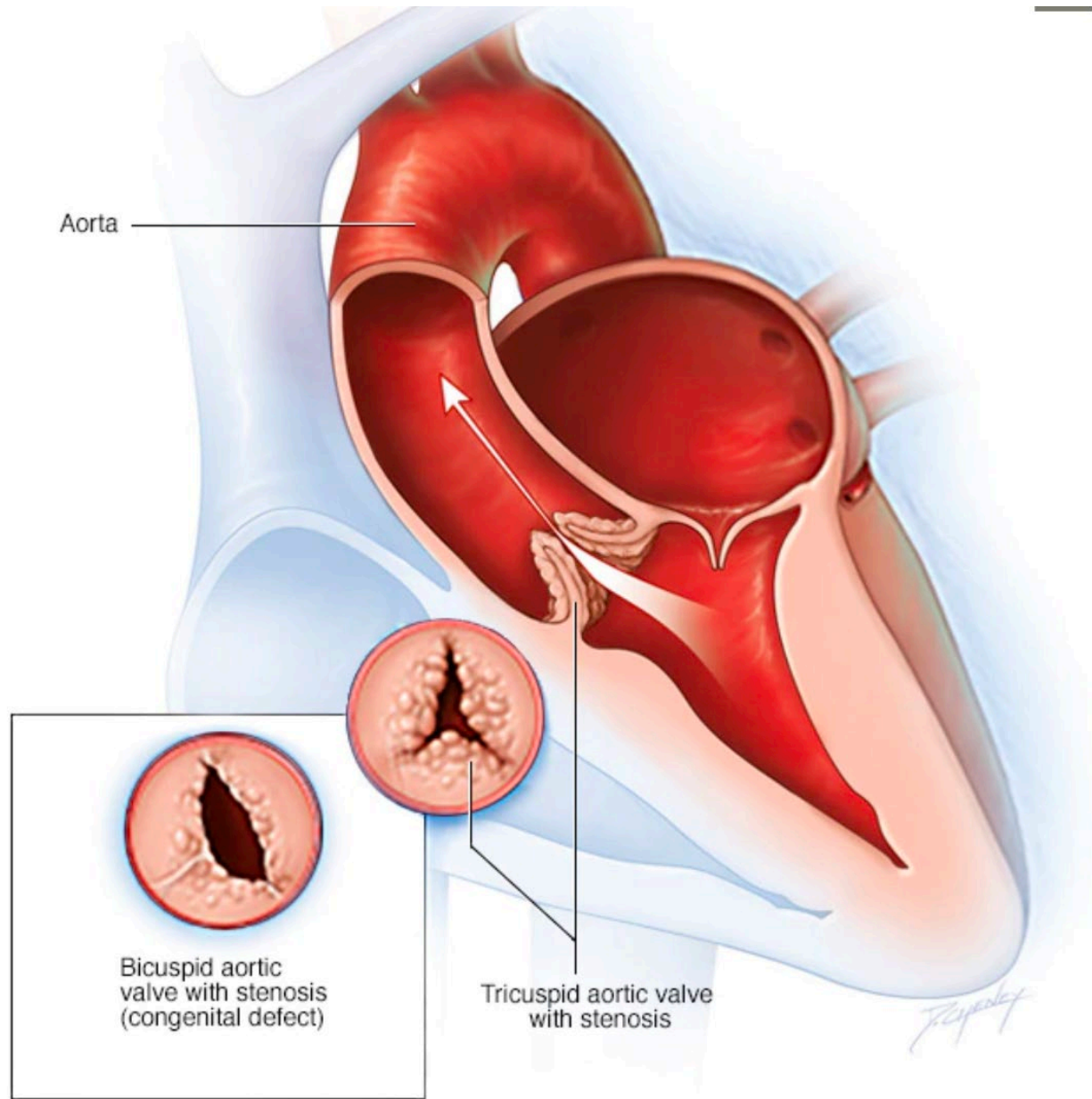
Validation
N = 6,700

Testing
N = 13,400

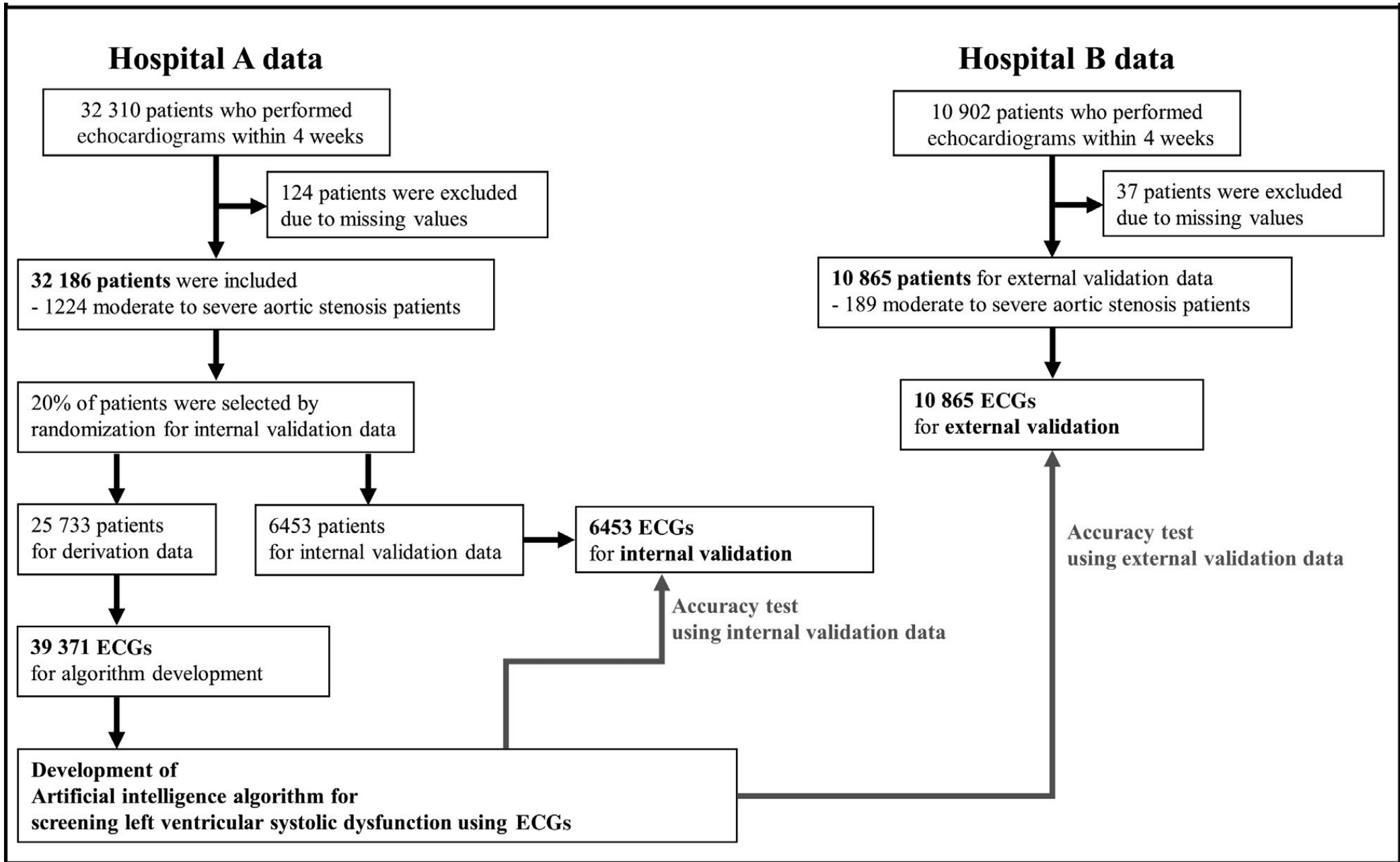


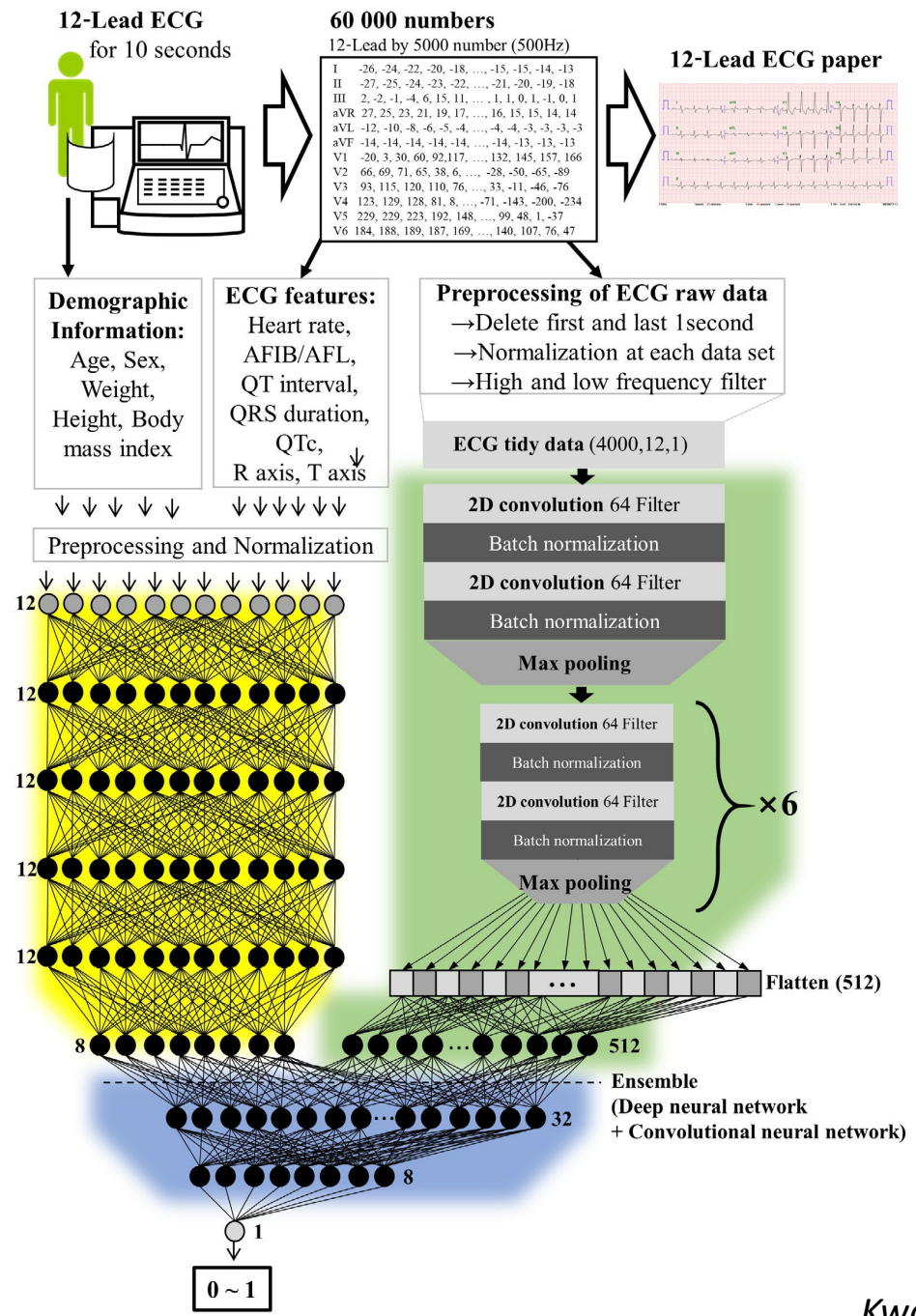
Sensitivity 87%
Specificity 91%
Positive Predictive Value 31%
Negative Predictive Value 99%

Subgroup	Sensitivity	Specificity
Age <40 Years	95%	92%
LVH by ECG	97%	68%
"Normal" ECG	93%	87%



AORTIC STENOSIS





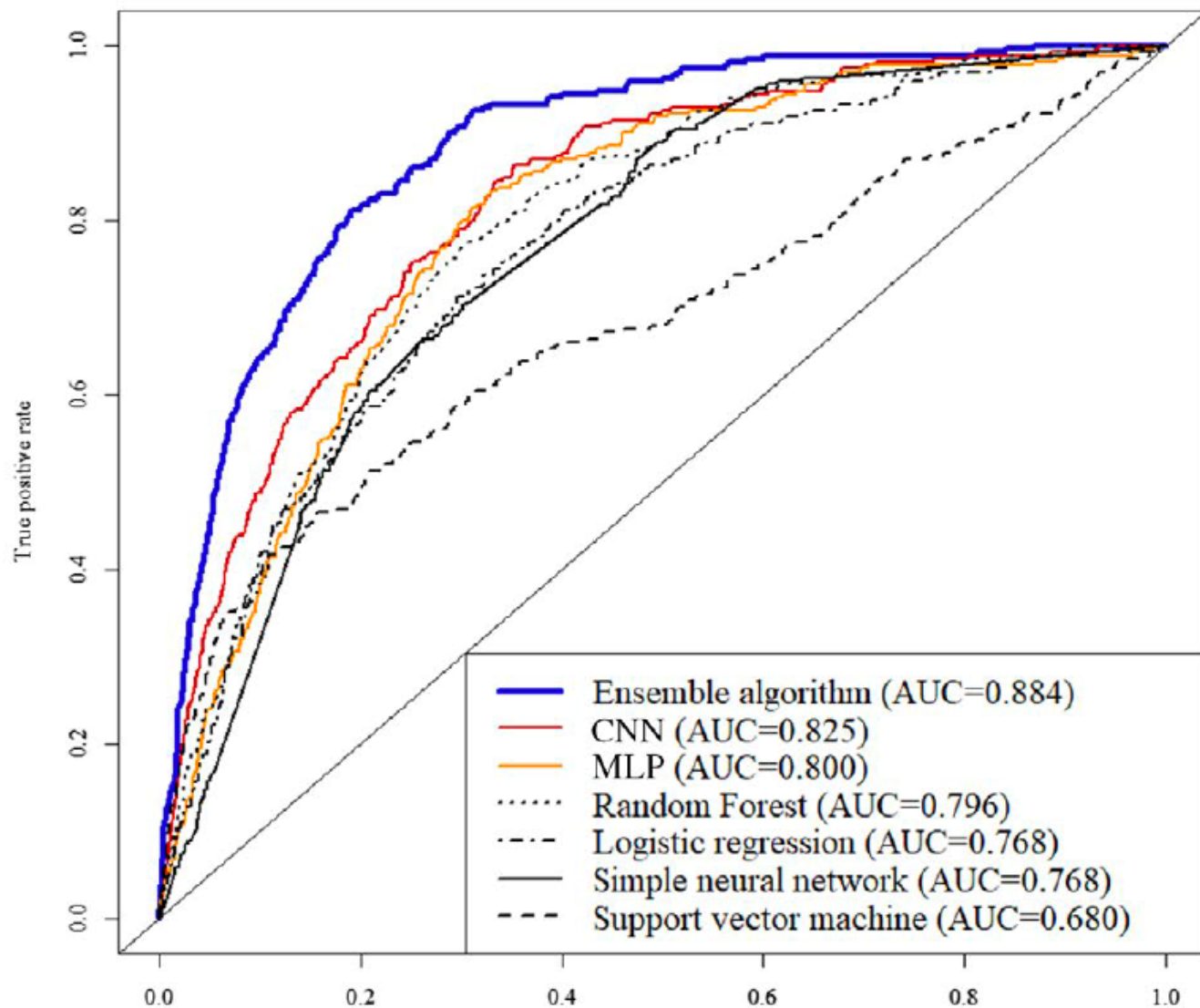
ORIGINAL RESEARCH

Deep Learning–Based Algorithm for Detecting Aortic Stenosis Using Electrocardiography

Joon-Myoung Kwon, MD, MS*; Soo Youn Lee, MD, MS*; Ki-Hyun Jeon, MD, MS; Yeha Lee, PhD; Kyung-Hee Kim, MD, PhD; Jinsik Park, MD, PhD; Byung-Hee Oh, MD, PhD; Myong-Mook Lee, MD, PhD

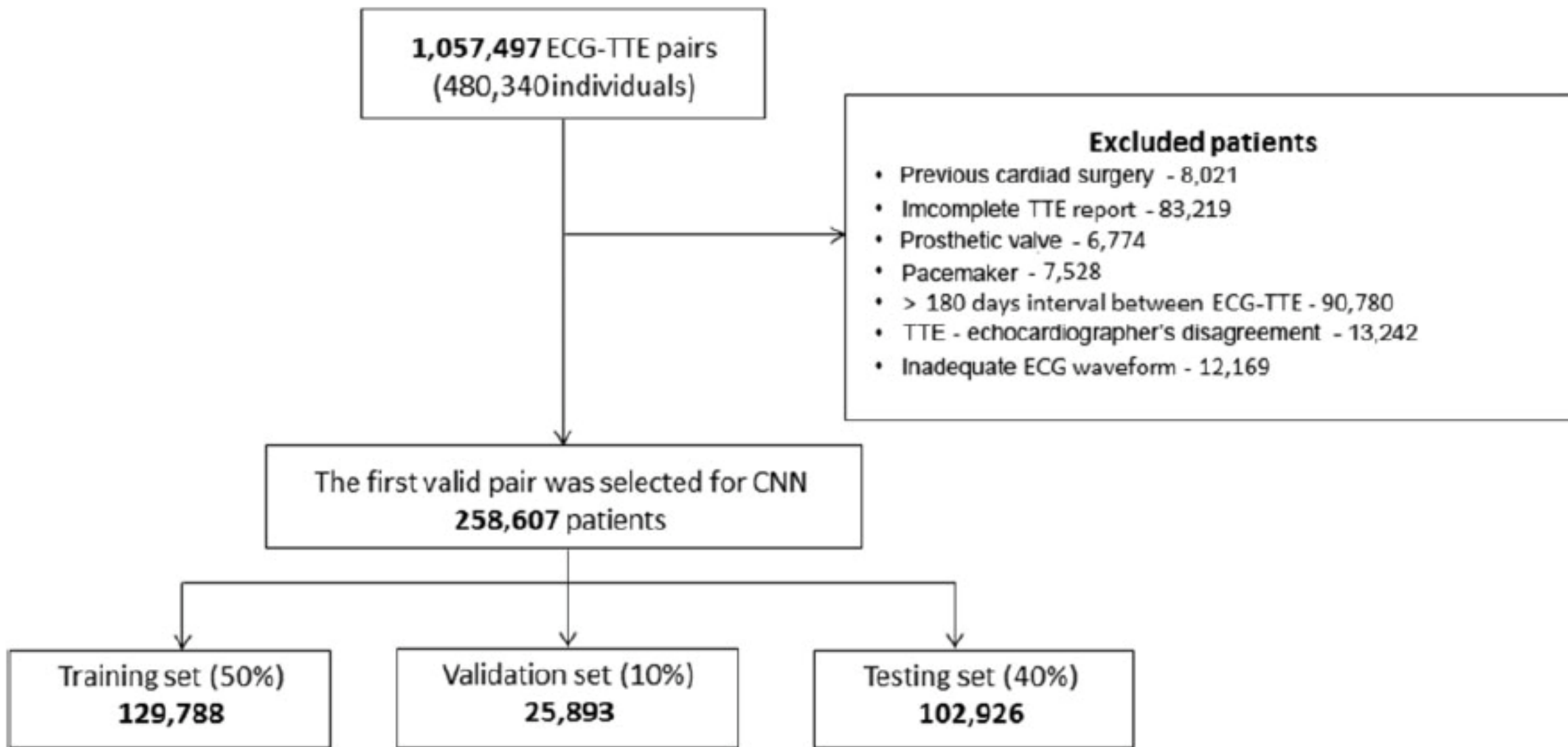
Algoritmus AI a EKG

ROC curve of internal validation



Výkon algoritmu AI pre detekciu aortálnej stenózy

ROC krivka je graf, ktorý popisuje kvalitu binárneho klasifikátora v závislosti na nastavení jeho klasifikačného prahu.



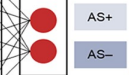
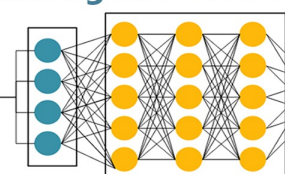
258,607 Patients

ECG-Echo pairs



50% Training and 10% Validation

12 lead ECG waveform fed as input



Illustrated Convolutional Neural Network to predict AS from ECG
We used DenseNet architecture with 62 convolutional layers and a fully connected output layer

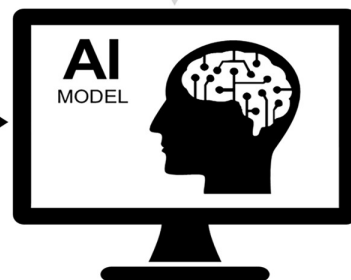
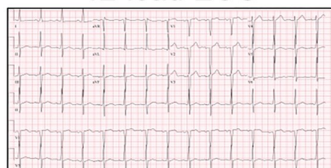
40% Testing

102,926 patients

Model	AUC	Sensitivity	Specificity
1	0.85	78.0%	74.0%
2	0.87	78.0%	79.8%
3	0.90	75.0%	88.0%

1: ECG only
2: ECG + Age & Sex
3: Model 2 for non-hypertensive patients

12-lead ECG

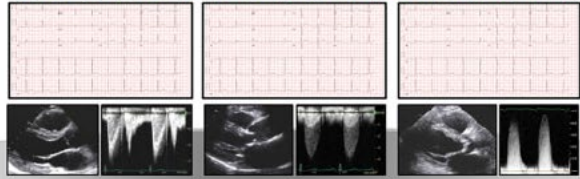


Aortic Stenosis

- Yes
- No

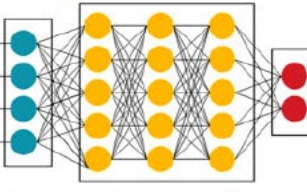
AI-ECG pre skrining aortálnej stenózy pomocou konvolučných neuronálnych sietí (CNN)

258,607 Patients
ECG-Echo pairs



50% Training and 10% Validation

12 lead ECG waveform fed as input



AS+

AS-

Illustrated Convolutional Neural Network to predict AS from ECG
We used DenseNet architecture with 62 convolutional layers and a fully connected output layer

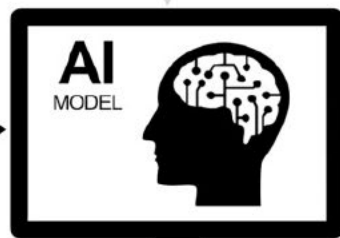
40% Testing

102,926 patients

Model	AUC	Sensitivity	Specificity
1	0.85	78.0%	74.0%
2	0.87	78.0%	79.8%
3	0.90	75.0%	88.0%

- 1: ECG only
- 2: ECG + Age & Sex
- 3: Model 2 for non-hypertensive patients

12-lead ECG

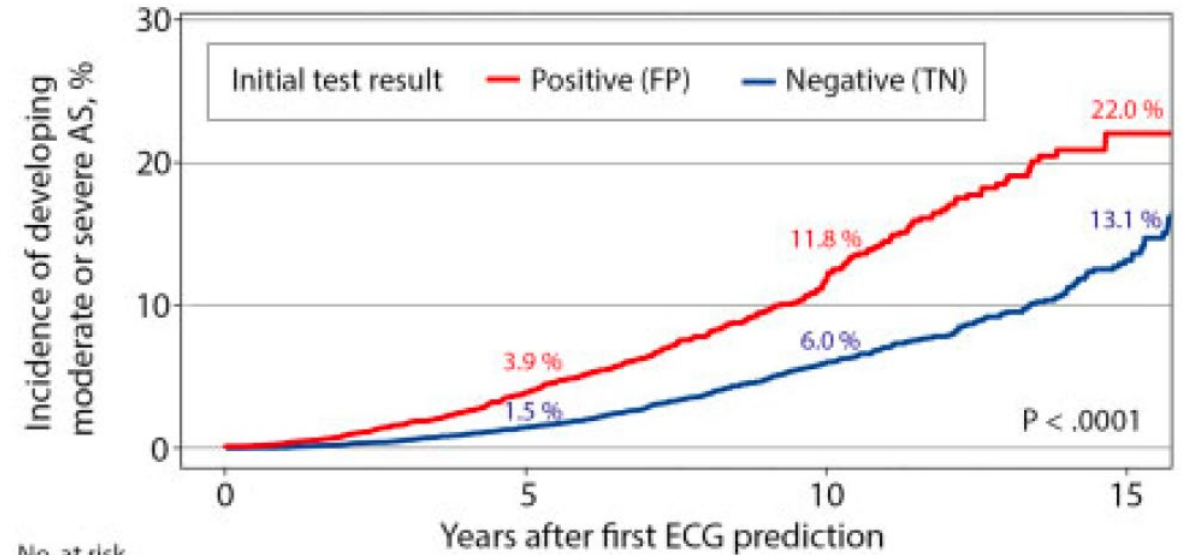


Aortic Stenosis

- Yes
- No

Electrocardiogram screening for aortic valve stenosis using artificial intelligence

Michal Cohen-Shelly¹, Zachi I. Attia¹, Paul A. Friedman¹, Saki Ito¹,



No. at risk

	0	5	10	15
Positive (FP)	8474	2880	793	102
Negative (TN)	20718	7969	2654	382

Positive predictive value was low at 10.5%, but negative predictive value was 98.9%.

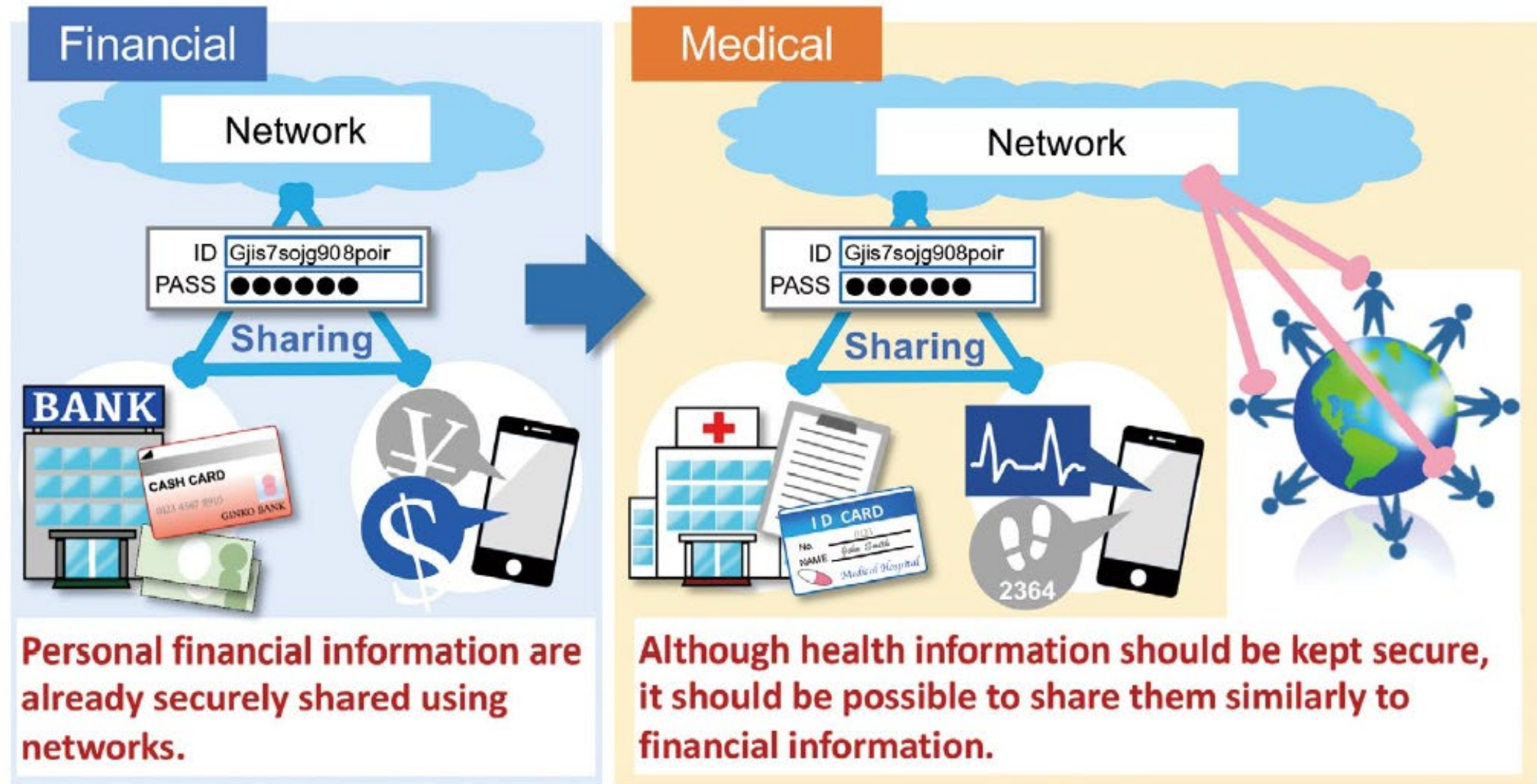
NEKARDIÁLNE OCHORENIA

Development and Validation of a Deep-Learning Model to Screen for Hyperkalemia From the Electrocardiogram

Conner D. Galloway, MSc; Alexander V. Valys, BS; Jacqueline B. Shreibati, MD; Daniel L. Treiman, BS;
Frank L. Petterson, PhD; Vivek P. Gundotra; David E. Albert, MD; Zachi I. Attia, MSc; Rickey E. Carter, PhD;
Samuel J. Asirvatham, MD; Michael J. Ackerman, MD, PhD; Peter A. Noseworthy, MD;
John J. Dillon, MD; Paul A. Friedman, MD

- Using 2 leads of an ECG acquired from patients with chronic kidney disease, a DL detected elevated potassium with an AUC of 0.853 to 0.883 and a sensitivity of 88.9% to 91.3%.
- The application of AI to the ECG may enable screening for hyperkalemia (HR for death with hyperkalemia as high as 13!)

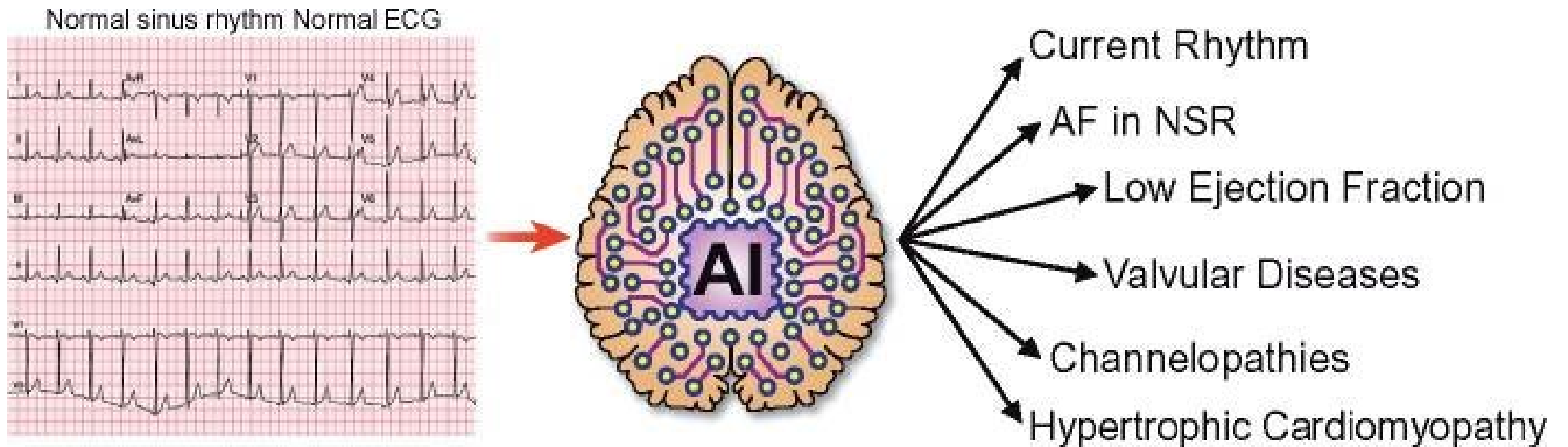
AI and personal data protection



AI heralds a new era in expanding the role of ECG

- **ECG is rich** on relevant pathophysiologic information which remained largely undiscovered
- **Deep learning** allows for identifying both "**visible and invisible**" contents of the ECG.
- **Two important advantages offered:**
 - **compensate for lack of expertise** in ECG interpretation by non-specialist
 - **detecting early stages** of various disease processes incl. non-cardiac diseases not discernable otherwise
- **Potential to diagnose and predict**
 - later manifestation of the disease (aortic stenosis, AFIB, HCM)
 - disease with intermittent phenotype manifestation (paroxysmal arrhythmias)
 - ECG as biomarker for systemic diseases (i.e. NPV for +SARS-CoV2 - 99% ...)

AI V KARDIOLÓGII



Umelá inteligencia (AI) a EKG

- AI otvára pre EKG netušený, v podstate nadľudský potenciál pre diagnostiku KV ochorení
- Schopnosť učiť sa z obrovského objemu dát (bez potreby súčasnej znalosti ich biologických mechanizmov) vytvára možnosť diagnostikovať aj nekardiálne ochorenia /cirhóza pečene, COVID-19,...)
- Vzhľadom k širokej dostupnosti EKG a jeho získania pomocou “nositeľných! senzorov (“wearables”) môže byť AI cestou k “demokratizácii” - t.j. univerzálnej prístupnosti pokročilej kardiovaskulárnej medicíny



A little learning is a dangerous thing;
drink **deep**, or taste not the Pierian
spring:
there shallow draughts intoxicate the
brain,
and drinking largely sobers us again.

Alexander Pope
"An Essay on Criticism", (1709)

ĎAKUJEM ZA POZORNOST!



**THANK YOU FOR YOUR TIME AND
ATTENTION !**