



- **SIGNIFICATO PROGNOSTICO**
- **DEI MARKERS BIOUMORALI**
- **NELLO SCOMPENSO CARDIACO**

• **Antonio Bonora**

• **DAI Emergenza e Terapie Intensive**

• **Azienda Ospedaliera Universitaria di Verona**



Azienda Ospedaliera
Universitaria Integrata
Verona

•epidemiologia

- prevalenza nella popolazione adulta 1-2%
- Moster A, Hoes AW. Heart 2007

- prevalenza nella popolazione > 70: $\geq 10\%$
- Ponikowski P, Voors AA, Anker SD et al.
- Eur Heart J 2016

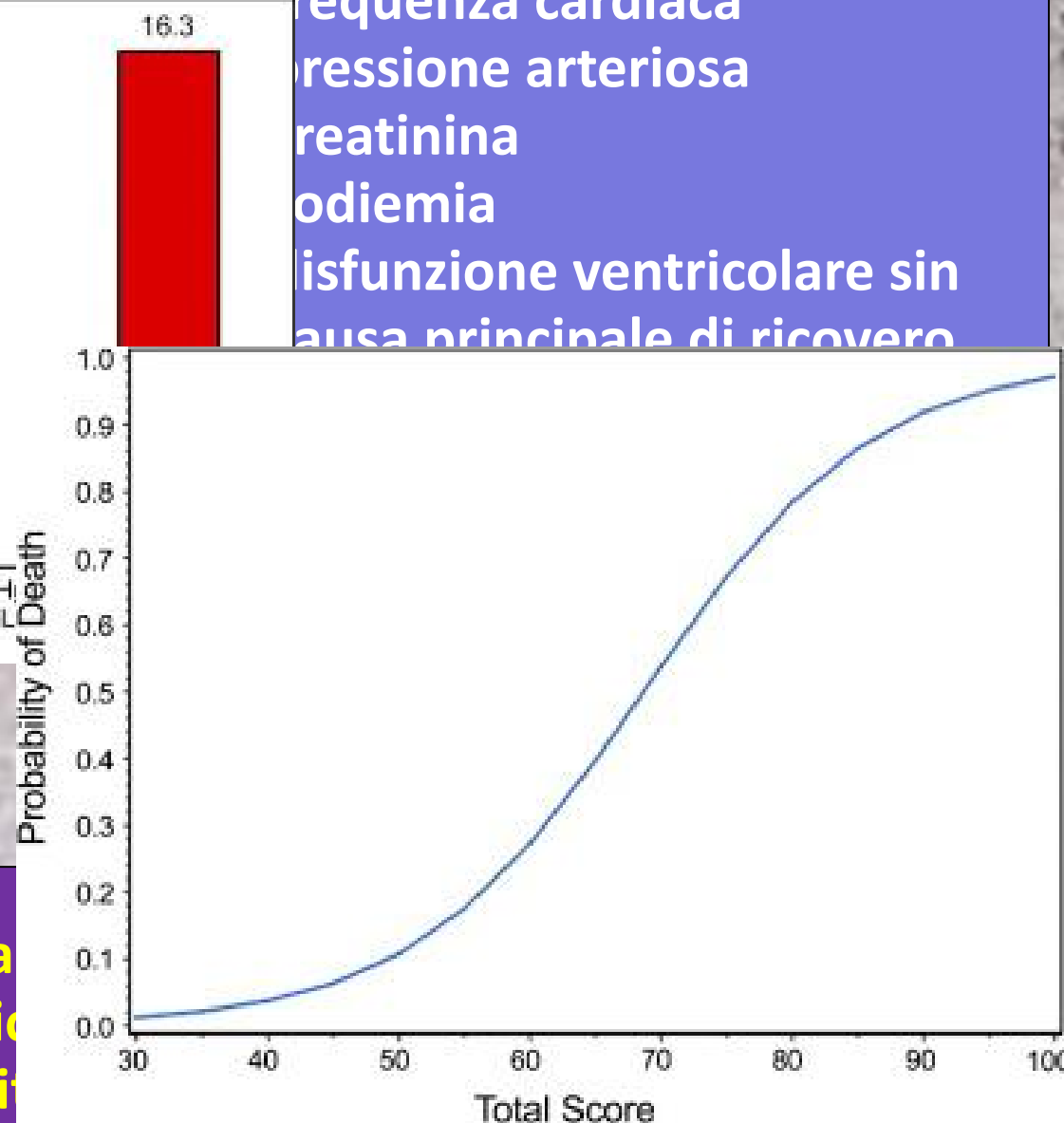
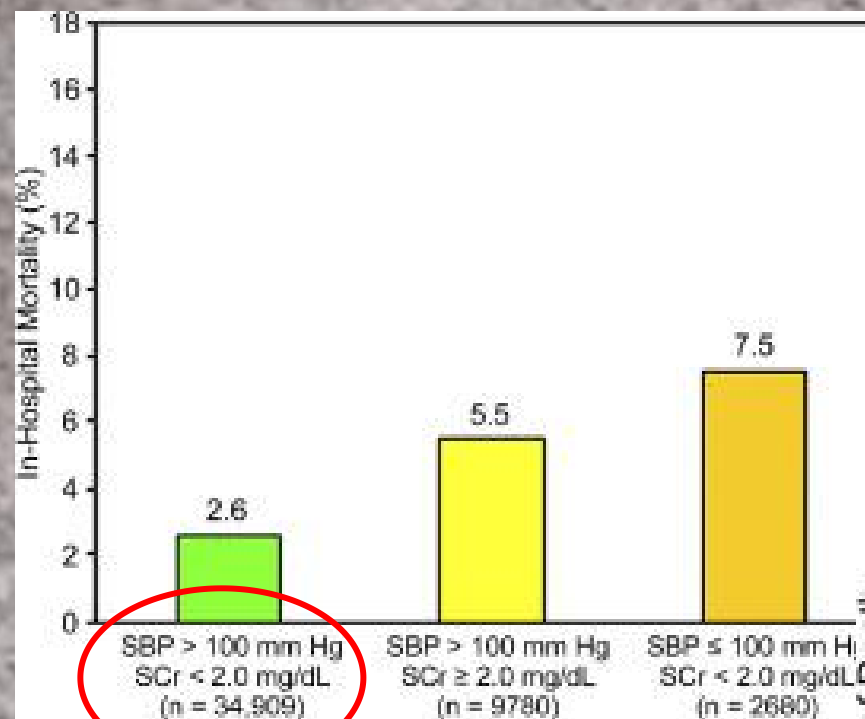
- oltre 1 milione di ricoveri all'anno
- Roger VL, Go AS, Lloyd-Jones DM. Circulation 2011

- mortalità ancora assai elevata
- Zile MR, Bennett TD, St John Sutton M et al. Circulation 2008
- Lassus JP, Siirilä-Waris K, Nieminen MS et al. Int J Cardiol 2013



•fattori predittivi

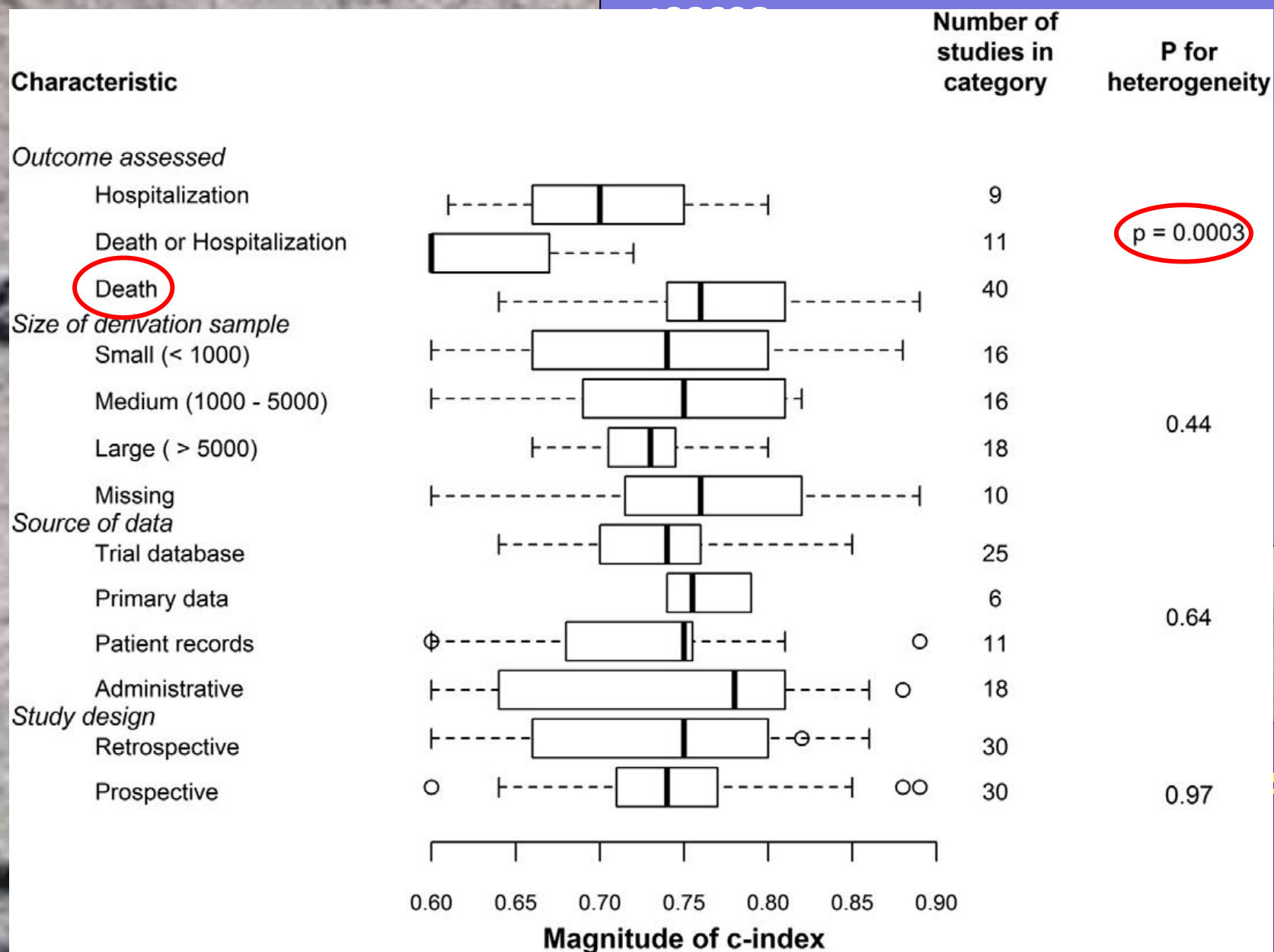
- età
- frequenza cardiaca
- pressione arteriosa
- creatinina
- sodiemia
- disfunzione ventricolare sinistra
- causa principale di ricovero



•Abra
Predic
hospit

•fattori predittivi

•età



•markers scompenso

- BNP
- ANP
- coceptina

- troponine
- fatty acid binding protein
- proteina (APO-1)

•RD
W

- selectina
- citochine
- proteina C-reattiva
- growth differantion factor
- mieloperossidasi

- azotemia
- creatinina
- cistatina C
- albuminuria

•RDW: un “nuovo” marker

A person in a white shirt and dark pants is swinging a large mallet at a target on a wall. The target is a red bullseye. The person is in the middle of the swing, with the mallet raised high. The background is a textured wall.

•L'RDW (red blood cell distribution width) è indice della variabilità del volume eritrocitario, definita convenzionalmente anisocitosi

•Salvagno GL, Sanchis-Gomar F, Picanza A, Lippi G. Red blood cell distribution width: A simple parameter with multiple clinical applications. Crit Rev Clin Lab Sci 2015

•Lippi G, Mattiuzzi C, Cervellin G. Learning more and spending less with neglected laboratory parameters: the paradigmatic case of red blood cell distribution width. Acta Biomed 2017

•RDW: marker

p

•STUDI COM

•43

INSERITI

ETANALISI

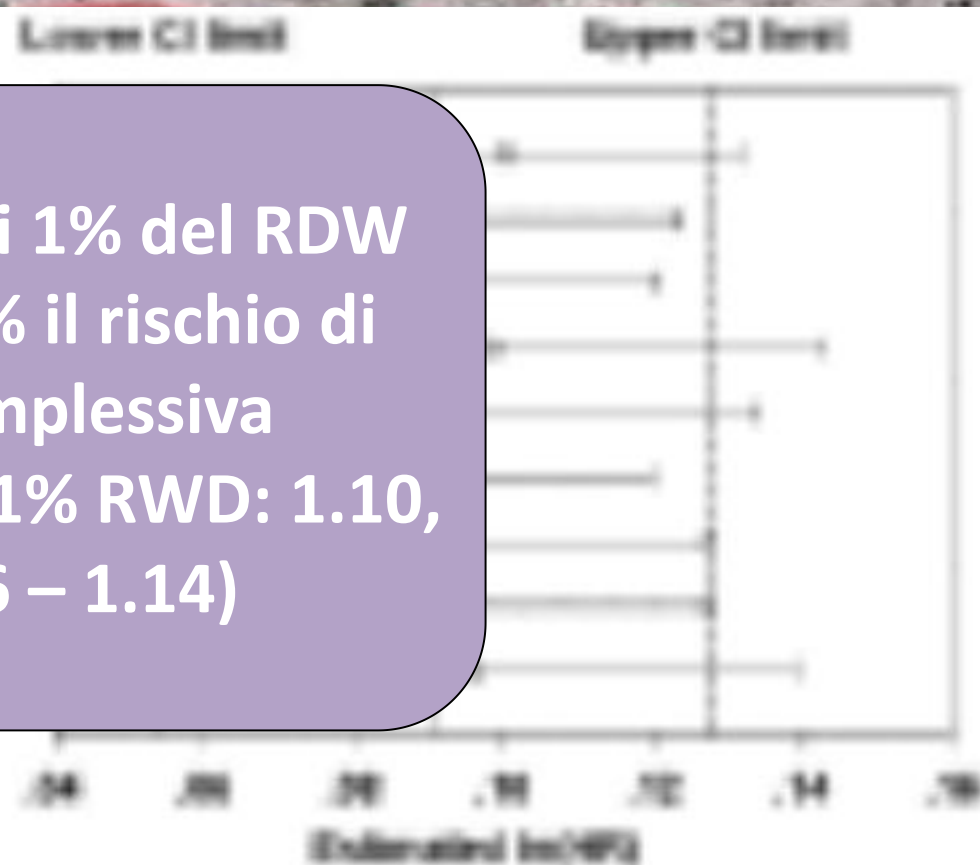
9

- mortalità complessiva
- mortalità per cause cardiache
 - ricoveri complessivi
- ricoveri per scompenso cardiaco
 - mortalità a 1 anno per
 - scompenso cardiaco

•Huang YL, Hu ZD, Liu SJ et al. Prognostic Value of Red Blood Cell Distribution Width for Patients with Heart Failure: A Systematic Review and Meta-Analysis of Cohort Studies. PLoS One 2014

•RDW: marker prognostico

- un incremento di 1% del RDW aumenta del 10% il rischio di mortalità complessiva
- (adjusted HR per 1% RDW: 1.10, 95% CI 1.06 – 1.14)



•Huang YL, Hu ZD, Liu SJ et al. Prognostic Value of Red Blood Cell Distribution Width for Patients with Heart Failure: A Systematic Review and Meta-Analysis of Cohort Studies. PLoS One 2014

•RDW: marker prognostico

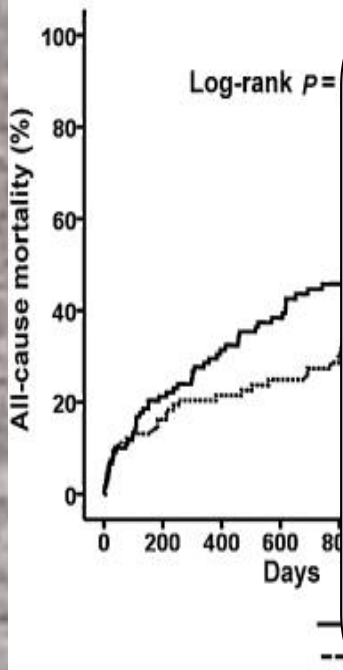
- 229 pazienti
- età media 76 ± 12
- NYHA III e IV

- RDW basale 15 ± 1.9
 - > 14.5 52.4%
 - < 14.5 47.6%
- RDW dimissione 15.4 ± 2.5
 - > RDW 36% paz
 - < RDW 64% paz
- Variazione media $0.0 \pm 1.2\%$

•Uemura Y, Shibata R, Takemoto K et al. Elevation of red blood cell distribution width during hospitalization predicts mortality in patients with acute decompensated heart failure. J Cardiol 2016

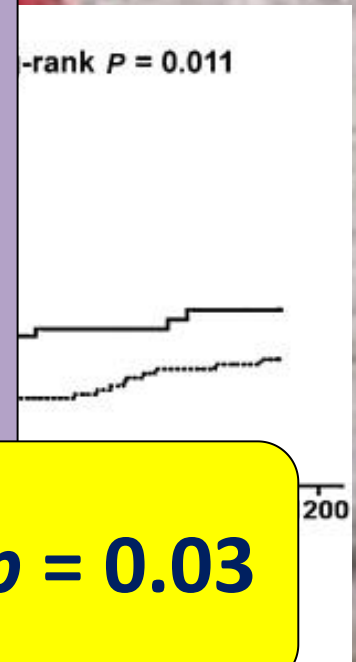
•RDW: marker

prognostic



- la variazione del RDW appare correlata alla mortalità totale
- (HR 1.310, 95% CI 1.141 – 1.505)
- e per cause cardiache
- (HR 1.250, 95% CI 1.037 – 1.508)

• $p = 0.001$



• $p = 0.03$

•Uemura Y, Shibata R, Takemoto K et al. Elevation of red blood cell distribution width during hospitalization predicts mortality in patients with acute decompensated heart failure. J Cardiol 2016



- Early in-hospital variation of red blood cell distribution width predicts mortality in patients with acute heart failure

- Turcato G, Zorzi E, Prati D, Ricci G, Bonora A, Zannoni M, Maccagnani A, Salvagno GL, Sanchis-Gomar F, Lippi G
•Int J Cardiol 2017

•popolazione di

- CRITERI DI
- ESCLUSIONE

- 1242 pazienti osservati in D5A per scompenso cardiaco acuto
- gen – dic 2015

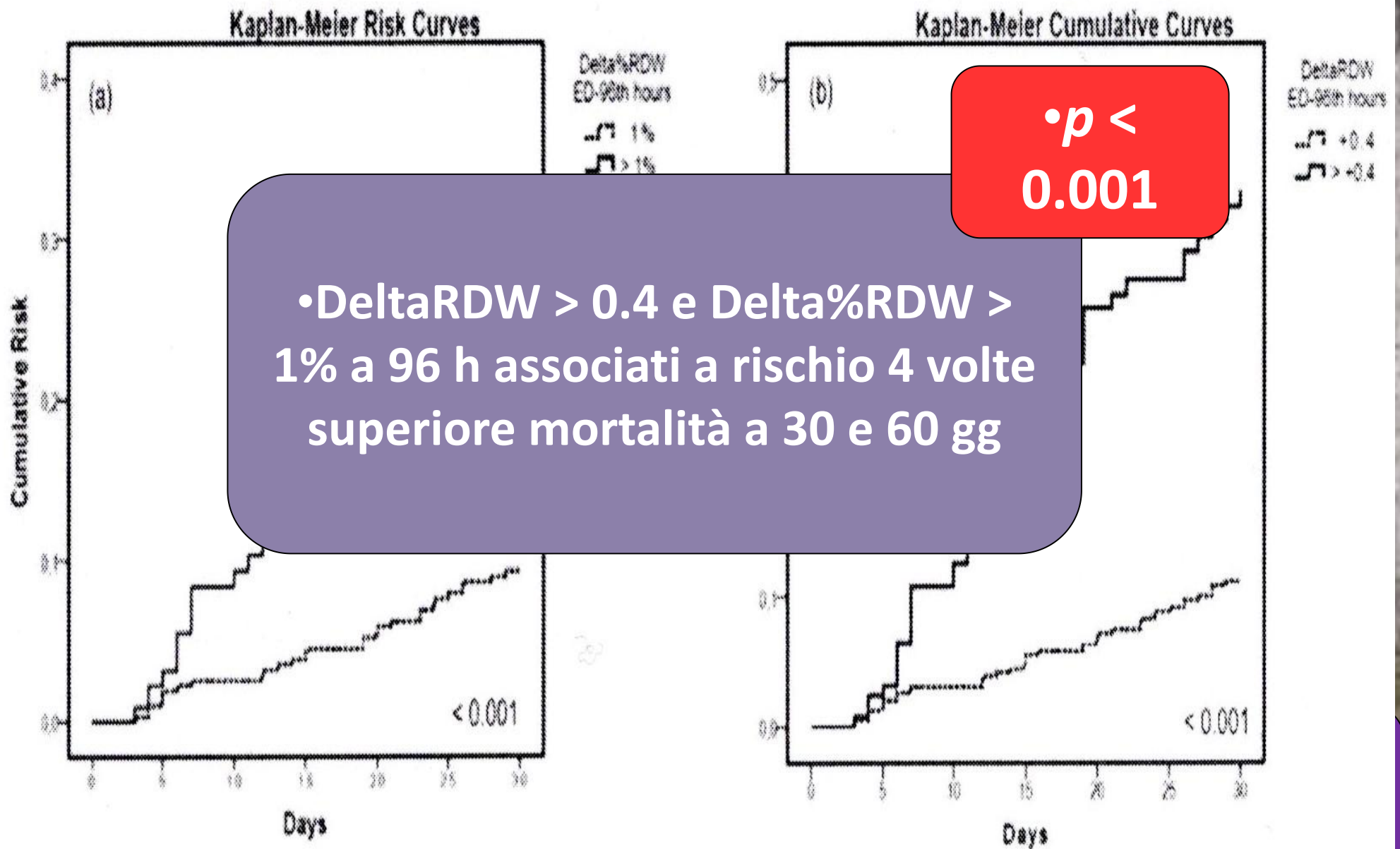
- età < 65 anni
- sintomi da > 5 gg
- diagnosi di infezione polmonare
- diagnosi di SCA
- patologia neoplastica
- aspettativa di vita < 6 mesi
- anemia severa

•Turcato G, Zorzi L, Frati D et al. Leucocyte blood cell distribution width predicts mortality in acute heart failure. Int J Cardiol 2015

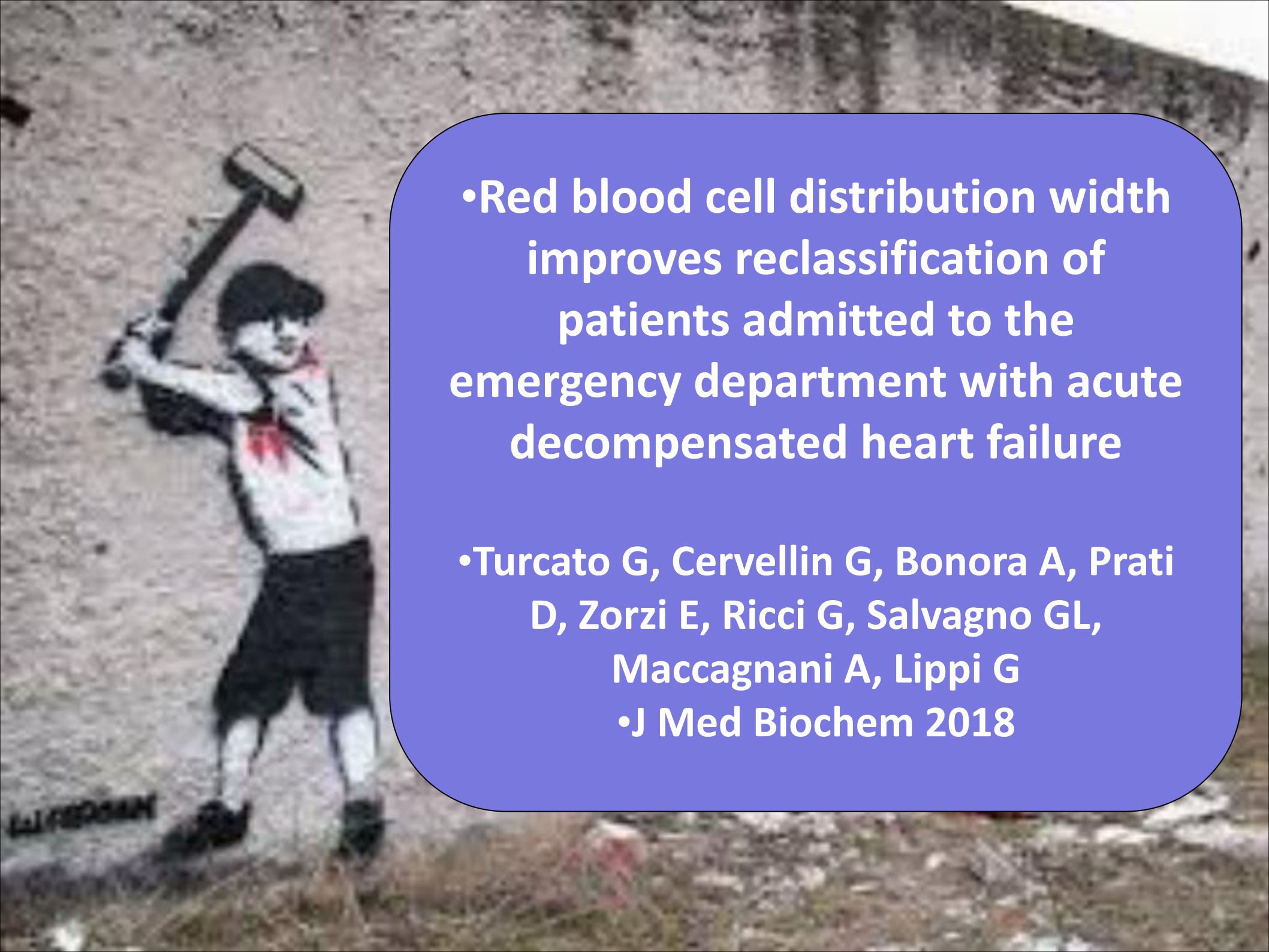
- 588 pazienti
- 319 maschi
- 269 femmine
- età media 83.5 anni
- (range 74 – 91)

Relationship between 30-day mortality and variation of red blood cell distribution width (RDW) between measurement obtained at ED admission and after 48 or 96 hours of hospital stay

30-day mortality		Univariate analysis	p	Multivariate analysis	p
		Odds Ratio (95% CI)		Adjusted odds ratio (95% CI)	
30-day mortality	ED admission and after 48 hours of hospital stay				
	As continuous variable				
	DeltaRWD	1.711 (1.25-2.35)	<0.001	1.46 (1.05-2.03)	0.024
	Delta%RDW	1.09 (1.03-1.14)	<0.001	1.07 (1.01-1.19)	0.036
	As categorical variable				
	DeltaRWD >0.4%	3.54 (2.01-6.25)	<0.001	3.04 (1.56-5.94)	0.001
	Delta%RDW >1%	1.85 (1.17-2.94)	0.009	1.59 (1.01-2.68)	0.048
	ED admission and after 96 hours of hospital stay				
60-day mortality	As continuous variable				
	DeltaRWD	2.29 (1.71-3.07)	<0.001	1.96 (1.45-2.65)	<0.001
	Delta%RDW	1.15 (1.09-1.20)	<0.001	1.12 (1.07-1.18)	<0.001
	As categorical variable				
	DeltaRWD >0.4%	4.34 (2.66-7.09)	<0.001	3.65 (2.02-6.15)	<0.001
	Delta%RDW >1%	3.67 (2.25-5.99)	<0.001	2.86 (1.67-4.91)	<0.001



acute heart failure. Int J Cardiol 2016



- Red blood cell distribution width improves reclassification of patients admitted to the emergency department with acute decompensated heart failure
- Turcato G, Cervellin G, Bonora A, Prati D, Zorzi E, Ricci G, Salvagno GL, Maccagnani A, Lippi G
- J Med Biochem 2018

•popolazione dello studio

- 2278 pazienti osservati in DEA per scompenso cardiaco acuto
- gen 2013 – dic 2016

- 554 esclusi

- 1704 pazienti
 - 824 maschi
 - 880 femmine
- età media 84 anni
- (range 76 – 92)

•Turcato G, Cervellin G, Bonora A et al. Red blood cell distribution width improves reclassification of patients admitted to emergency department with acute decompensated heart failure. J Med Biochem 2016

Table II Multivariable logistic regression analysis for prediction of 30-day mortality in patients with acute decompensated heart failure (BNP, creatinine, serum sodium and chloride are considered as continuous variables).

Coefficient	Standard error	Odds ratio	95% CI	p	
Final model					
BNP	0.511	0.088	1.666	1.403–1.979	<0.001
Creatinine	0.331	0.071	1.392	1.212–1.599	<0.001
Hypernatremia	0.323	0.110	1.381	1.112–1.715	0.003
Hypochloremia	0.509	0.111	1.665	1.340–2.068	<0.001
Final model + RDW					
BNP	0.460	0.088	1.584	1.333–1.882	<0.001
Creatinine	0.278	0.072	1.321	1.148–1.520	<0.001
Hypernatremia	0.306	0.110	1.358	1.094–1.686	0.006
Hypochloremia	0.496	0.111	1.642	1.322–2.039	<0.001
RDW	0.379	0.074	1.461	1.263–1.690	<0.001

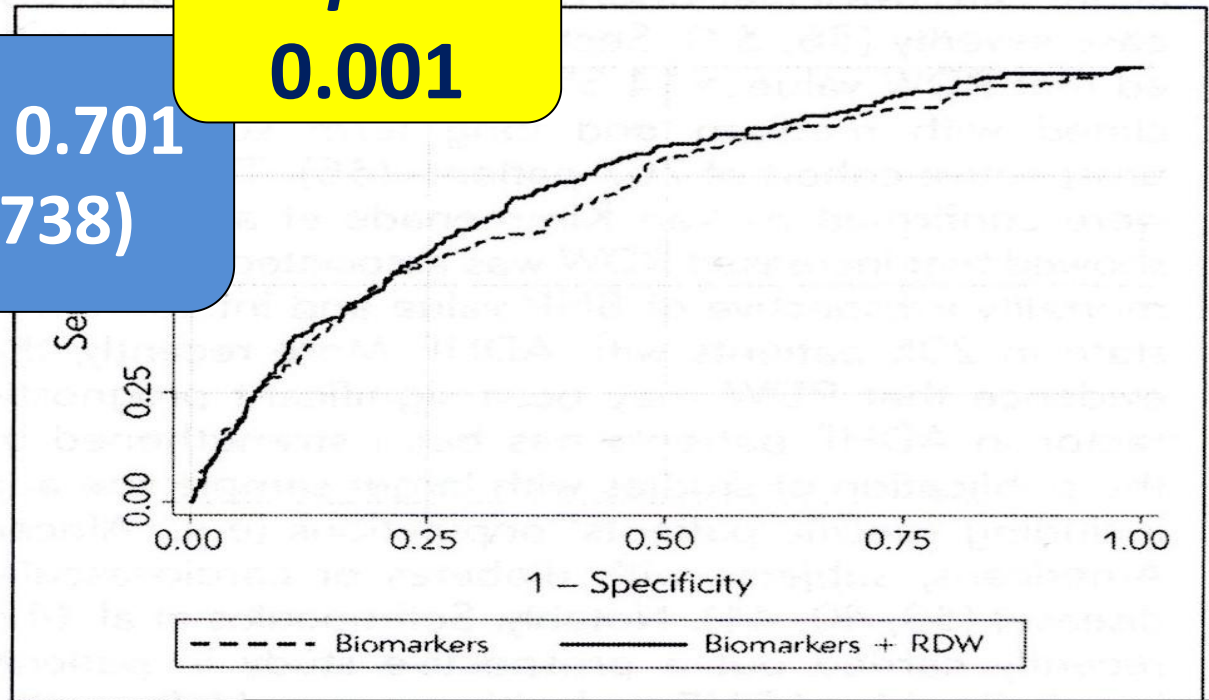
BNP, B-type natriuretic peptide; RDW, red blood cell distribution width.

Turcato G, Cervellini G, Bonora A et al. Red blood cell distribution width improves reclassification of patients admitted to emergency department with acute decompensated heart failure. J Med Biochem 2016

•modello predittivo

• $p =$
0.001

•AUC modello finale 0.701
•(95% CI 0.662 – 0.738)



• $p <$
0.001

•AUC modello finale + RDW
0.723
•(95% CI 0.662 – 0.738)

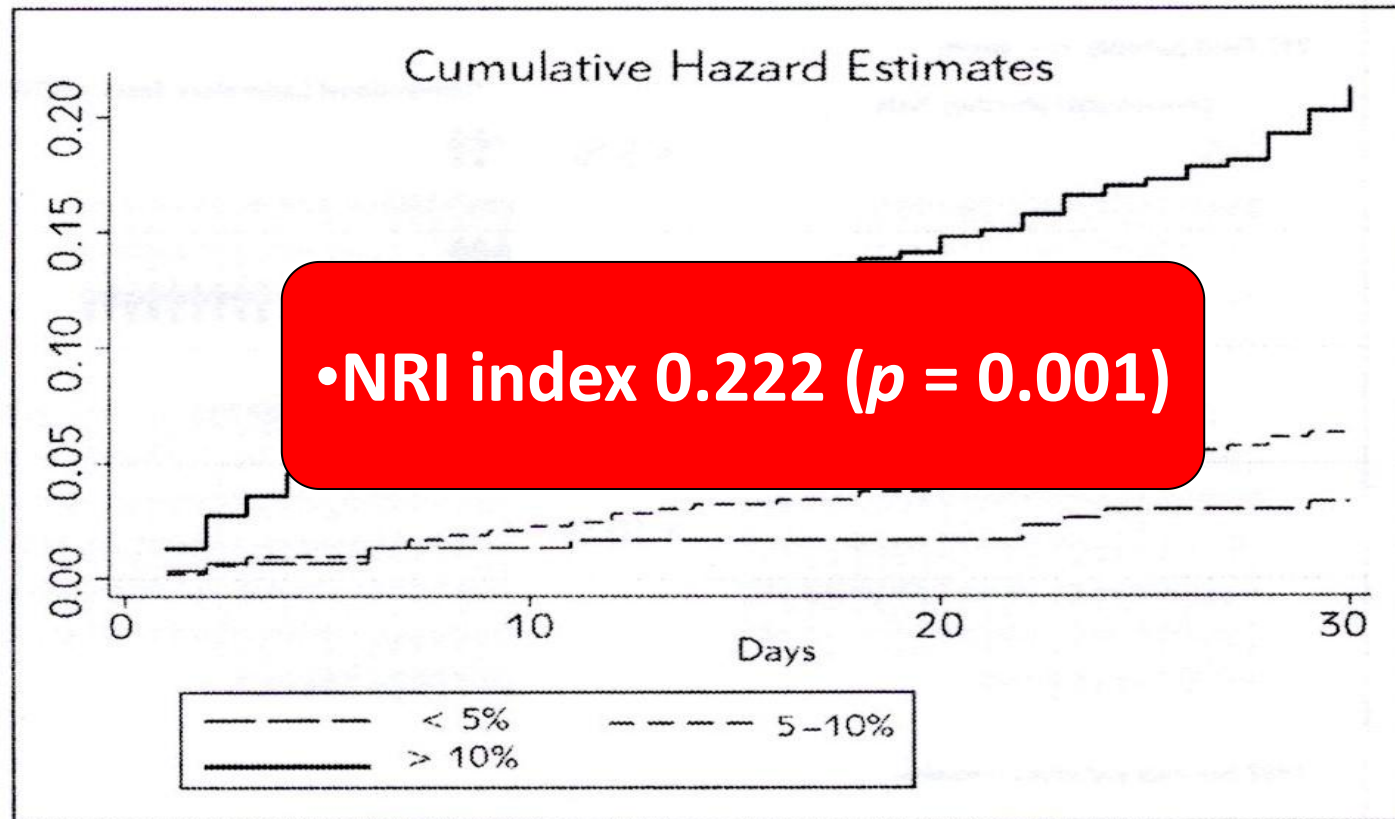
•Turcato G, Cervellin G, Bonora A et al. Red blood cell distribution width improves reclassification of patients admitted to emergency department with acute decompensated heart failure. J Med Biochem 2016

•net reclassification improvement

(N)

Table III Results of net reclassification improvement (NRI) obtained by combining red blood cell distribution width (RDW) with conventional laboratory tests (B-type natriuretic peptide, creatinine, sodium and chloride).

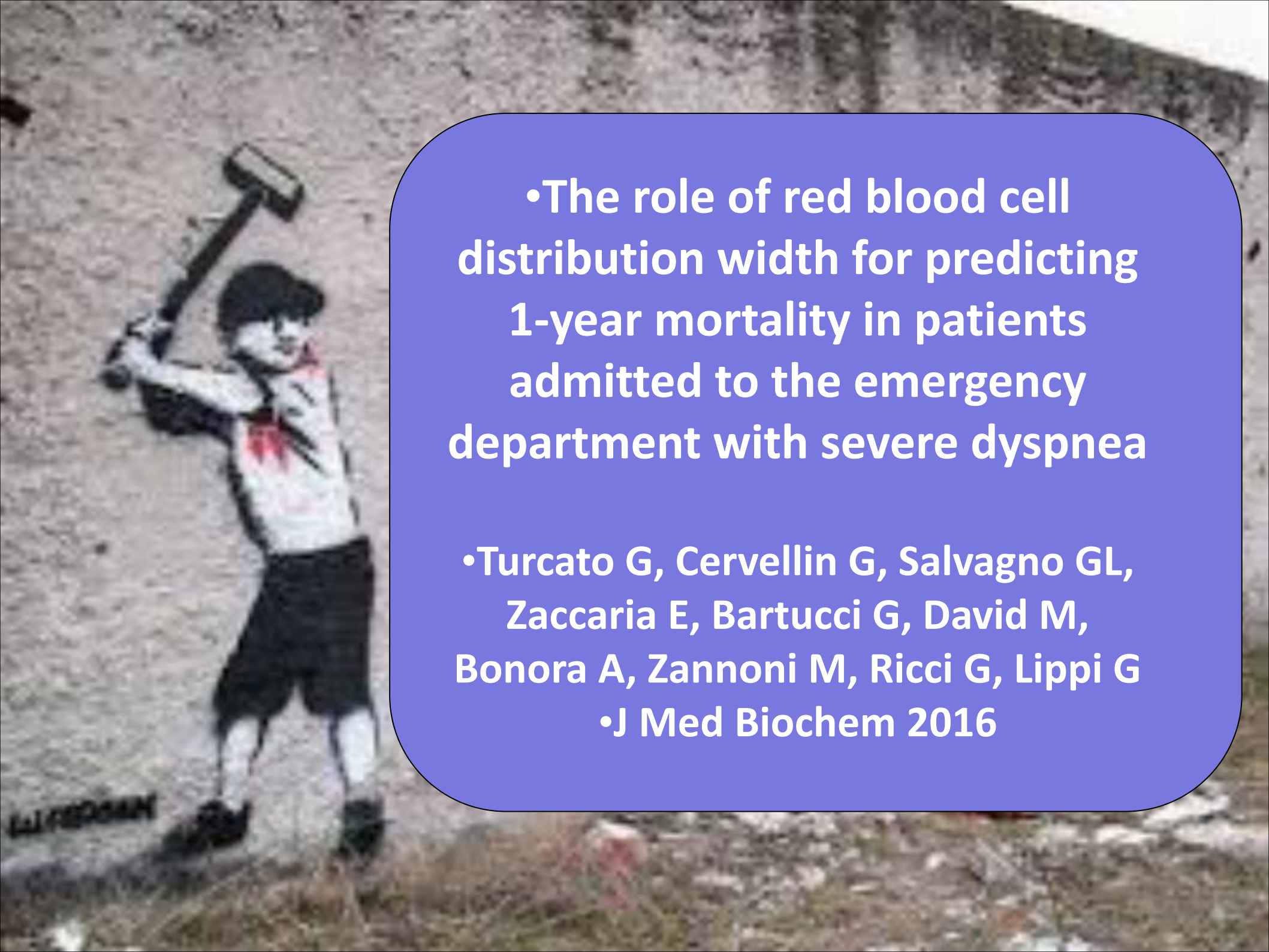
Patients d
< 5%
5-10%
> 10%
Total
Patients s
< 5%
5-10%
> 10%
Total



Bold font, r

Figure 3 Survival curve analysis after including red blood cell distribution width (RDW) in a predictive model based on conventional laboratory tests (i.e., B-type natriuretic peptide, creatinine, sodium and chloride).

•Turcato C et al. Red blood cell distribution width improves reclassification of patients admitted to emergency department with acute decompensated heart failure. J Med Biochem 2016

A young boy with dark hair, wearing a white t-shirt and dark shorts, is captured in the middle of a powerful swing with a large mallet. He is standing on a dirt field, and the background is a blurred natural setting. The mallet is dark and has a large, rectangular head. The boy's expression is one of concentration and effort.

- The role of red blood cell distribution width for predicting 1-year mortality in patients admitted to the emergency department with severe dyspnea

- Turcato G, Cervellin G, Salvagno GL, Zaccaria E, Bartucci G, David M, Bonora A, Zannoni M, Ricci G, Lippi G
•J Med Biochem 2016

Table III Multivariate analysis including the significant predictors of 1-year mortality after emergency department (ED) admission for severe dyspnoea from the univariate analysis.

Variables	% CI			p
RDW	-2.162			0.001
Lactate	0.411	1.508	1.174-1.937	0.001
Heart rate >100	0.855	2.352	0.884-6.260	0.087
SpO ₂ < 90%RDW	0.992	2.696	1.017-7.144	0.046

•mortalità a 1
anno

•36 paz (12.5%)

•Turcato G, Cervellin G, Salvagno GL et al. The role of red blood cell distribution for predicting 1-year mortality in patients admitted to emergency department with severe dyspnoea. J Med Biochem 2016

•modello predittivo

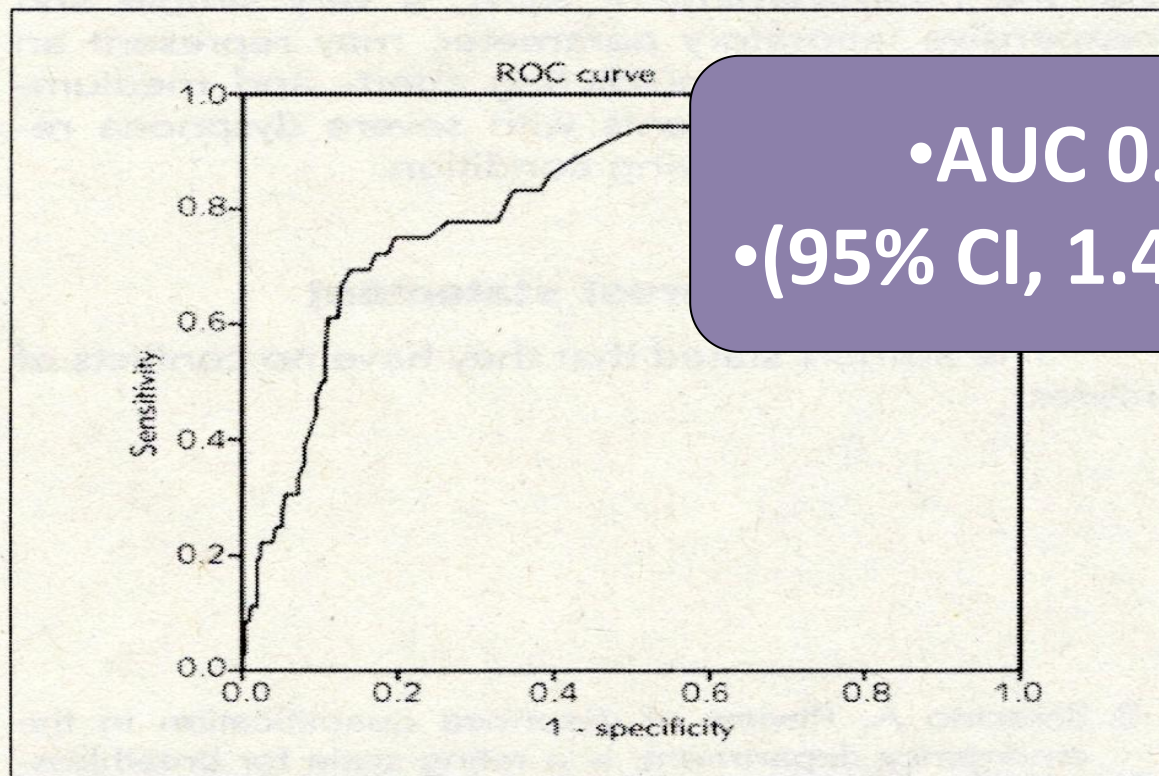


Figure 1 Receiver operating characteristics (ROC) curve analysis of red blood cell distribution width (RDW) for predicting 1-year mortality in patients admitted to the emergency department with severe dyspnoea.

•Turcato G, Cervellin G, Salvagno GL et al. The role of red blood cell distribution for predicting 1-year mortality in patients admitted to emergency department with severe dyspnoea. J Med Biochem 2016

•outcome mortalità

•sensibilità

87%

•specificità

•RDW $\geq 15\%$
•adjusted OR 1.72
•(95% CI 1.37 – 2.16)

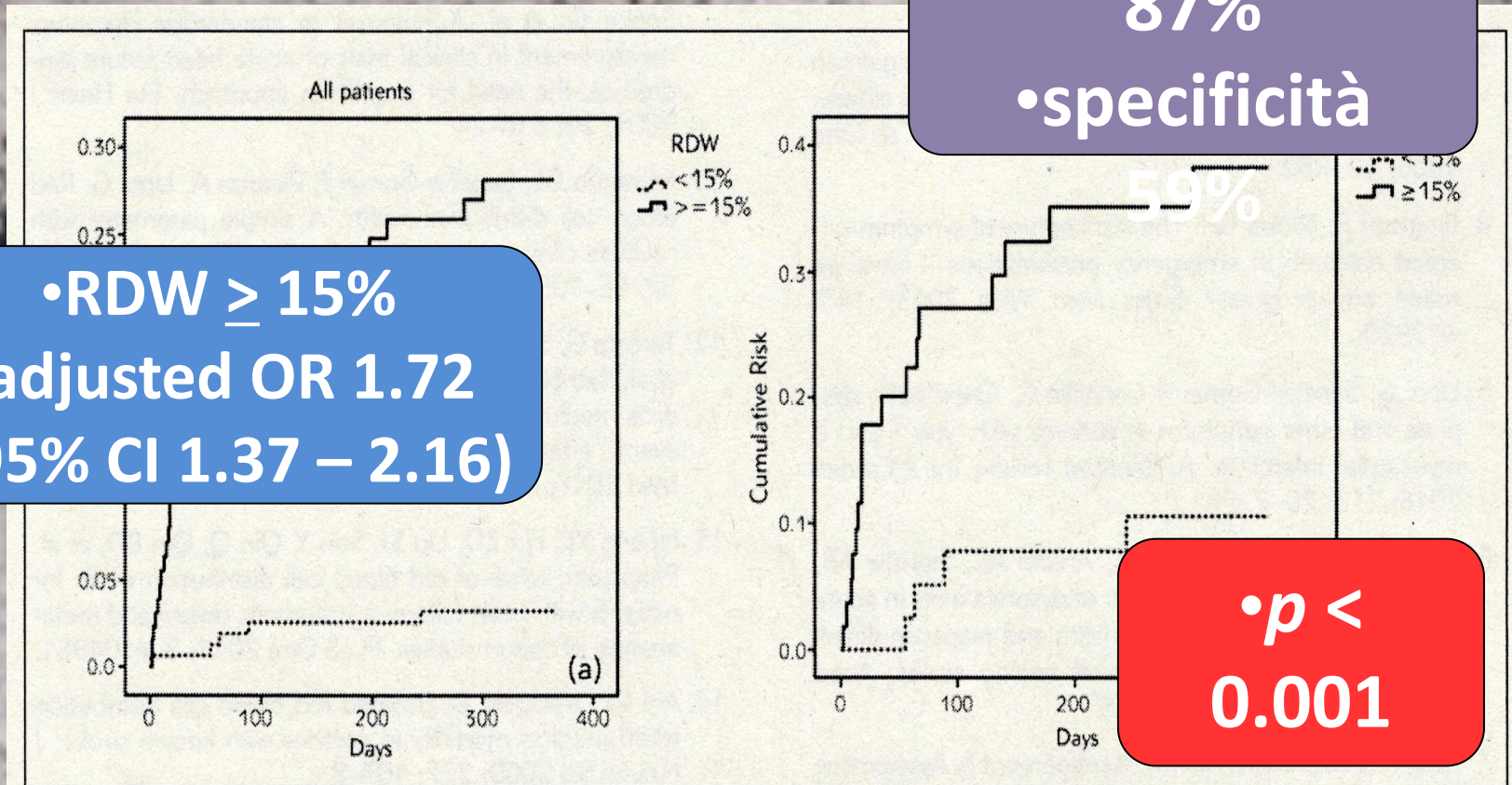


Figure 2 Survival curve analysis of 1-year mortality according to RDW value in patients admitted to the emergency department with severe dyspnoea.

•Turcato G, Cervellin G, Salvagno GL et al. The role of red blood cell distribution for predicting 1-year mortality in patients admitted to emergency department with severe dyspnoea. J Med Biochem 2016

RDW



**A SIMPLE KIND OF MAN
MARKER**