

METFORMIN-INDUCED LACTIC ACIDOSIS: A CASE ANALYSIS



Gioia D.¹, Netti G.S.¹, Cisero R.¹, Fili A.M.¹, Mangiameli I.S.¹, Cavagnero M.¹, Nobile E.¹, Ghignone F.¹, Quaranta S.¹, Bosso S.², Roasio A.², Bianchi A².

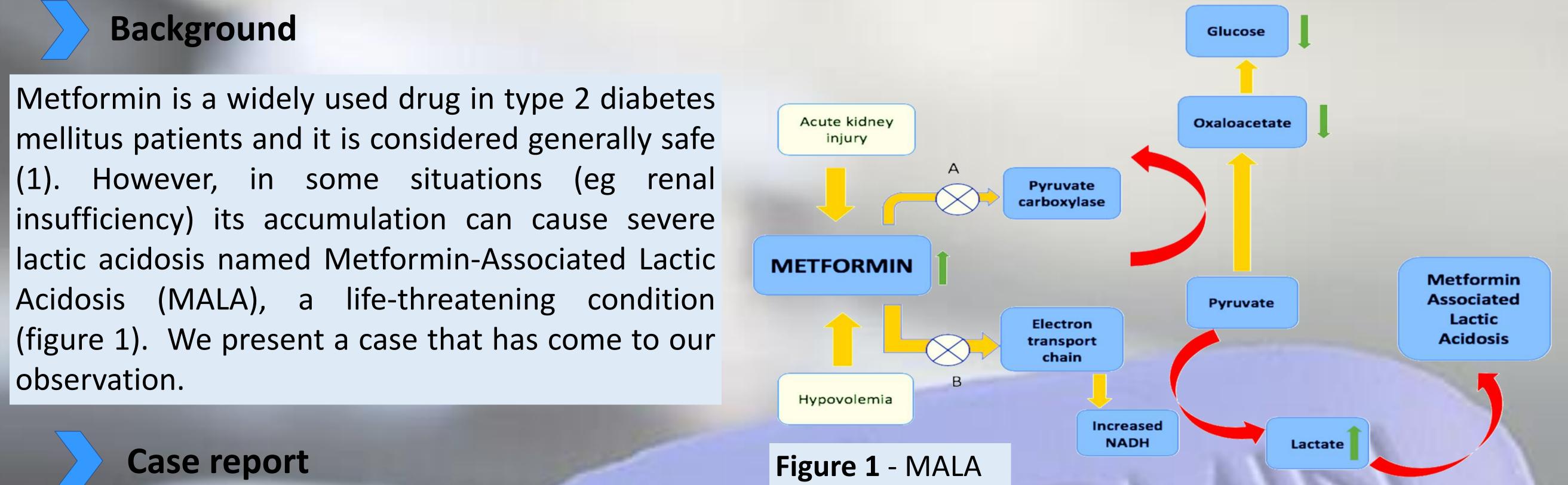
1) Department of Medical Science – University of Turin - Italy

2) Department of Anaesthesia and Intensive Care Medicine – "Cardinal Massaia" Hospital – Asti - Italy

Background

Discussion

renal



A 70-year-old man was admitted in Emergency Department for dyspnea and chest pain. At the entrance his vital signs were: GCS 15, respiratory rate 24 act/min and blood pressure 180/100 mmHg; the first arterial blood sample showed a severe metabolic acidosis (table 1). At first he was treated with NaHCO3 5.4% 500ml. After the start of treatment, we recorded a severe hypotension poorly responsive to fluid challenge with crystalloids; therefore norepinephrine was started (max 0.2mcg/kg/min). The subsequent anamnesis reveals that he had taken metformin to treat type 2 diabetes mellitus. The first blood sample collected in the Emergence Department indicated a severe renal impairment with creatinine 14.8 mg/dl associated with anuria. Hence we decided to admit the patient to ICU and submit him to urgent RRT with CVVHDF (within 3 h from accessing the ED). Our diagnostic hypothesis was metformin induced lactic acidosis (MALA). In the following 48 hours there was a gradual improvement of the metabolic state with normalization of the pH, return to the normal range of lactacidemia and resumption of diuresis with pharmacological stimulation. After 72 hours from ICU admission, RRT was suspended and then the patient was transferred to the medical department.

Approach to the patient with hyperlactatemia on metformin

Arterial blood sample at ED admission

Elevated lactate level plus metformin prescription

Is patient actually taking metformin		ot metformin-related	рН	6.93
Yes Yes Yes Yes Yes Probably m		metformin-induced lactic acidosis (MILA)	paCO2	12.4 mmHg
No No	(especially no al	(especially no alternative explanation i.e. coingestion)	paO2	139 mmHg
	Lactate due	to other acute illness	HCO3 -	2.6 mmol/l
Lactate due to metformin			BE	-28
MILA	MALA	MULA (metformin-unrelated lactic acidosis)	Lat	17 mmol/l
(metformin-induced lactic acidosis)	(metformin-associated lactic acidosis)		Glyc	241 mg/dl
Features favoring MILA end of spectrum				
Higher metformin dose		Lower metformin dose	K+	5.6 mEq/l
Chronic or subacute renal failure Markedly reduced GFR		No renal failure (or very acute renal failure) Fairly preserved GFR		1*
No other obvious causes of elevated lactate		Other causes of elevated lactate (e.g. sepsis)	Anion	20 6
No response to basic supportive/resuscitative n	neasures Lactate	The Internet Book of Critical Care, by @PulmCrit	Gap	38.6
Discussion	Figure 2 – MAL	A: decision-making approach		Table 1

Metformin intoxication should be suspected in a diabetic patient with metabolic acidosis, high anion gap and severe increase in lactacidemia. The history and blood samples allow an easy and early diagnosis and the measurement of the blood metformin level is not necessary. (figure 2). However, early supportive therapies such

as hemodinamic support and hemodialysis are essential to improve the outcome of these patients.	
References: 1) A. Protti et al. Critical Care 2012, 16:R75. 2) R. DeFronzo et al. Metabolism Clinical And Experimental 6	5. 2016 20-29