

# Long-term prognostic value of growth-differentiation factor 15 in acute myocardial infarction complicated by cardiogenic shock

## *Purpose*

Growth-differentiation factor 15 (GDF-15), a stress-responsive member of the transforming growth factor beta cytokine superfamily, has proven prognostic impact in cardiovascular disease. In acute myocardial infarction (AMI) complicated by cardiogenic shock (CS) impact on short-term the impact of GDF-15 was shown recently, but a possible long-term prognostic impact beyond the acute phase has not been investigated yet.

## *Methods*

In 190 patients with CS complicating AMI blood samples were collected during primary percutaneous coronary intervention (PCI). The blood was centrifuged immediately and the serum was frozen at -87° C. GDF-15 was measured with a standard ELISA-Kit. All-cause mortality at 1 year was used for long-term outcome assessment.

## *Results*

Patients with positive 1 year survival had in median significant lower levels of GDF-15 (5002 [IQR 2297;9134] vs. 10618 [IQR 6406;14458] pg/ml;  $p<0.001$ ). GDF-15 levels above 7452 pg/ml (best cut off by Youden-index) showed higher rates of death at 1 year (71.4 vs. 34.8 %,  $\text{Chi}^2$   $p<0.001$ ; log-rank-testing [HR 2.61 {95%CI 1.77-3.85};  $p<0.001$ ]). A landmark analysis in 30 day survivors showed a persistent discriminating effect of GDF-15 (log-rank-test day 30 to 1 year: HR 4.92 [95%CI 2.15-11.21];  $p<0.001$ ). In a multivariable stepwise Cox-regression model including all baseline variables with an univariable association to 1 year mortality ( $p<0.1$ : GDF-15, age, serum creatinine and lactate, ejection fraction, sex, prior stroke, NT-ProBNP, presence of coronary 3-vessel disease, patent culprit vessel after PCI and mechanical ventilation at admission) GDF-15, age, ejection fraction, serum lactate and a patent culprit vessel after PCI remained significant predictors of time to death (HR per 10 µg/L GDF-15 1.77 [95%CI 1.13-2.81],  $p=0.01$ ). Adding GDF-15 to a model including all multivariable significant predictors resulted in a significant increase of the area under the curve for prediction of 1 year mortality (0.767 without vs. 0.817 with GDF-15,  $p=0.046$ ).

## *Conclusions*

GDF-15 levels at baseline are an independent predictor of long-term mortality in acute myocardial infarction complicated by CS.