Prospective Changes of Cardiac and Hepatic Iron and Cardiac Function in Low and Intermediate-1 Risk MDS Patients

Antonella Meloni¹, Michele Rizzo², Giovanni Carulli³, Esther Natalie Oliva⁴, Francesco Arcioni⁵, Sergio Storti⁶, Maria Giovanna Neri¹, Stefania Renne⁷, Emanuele Grassedonio⁸, Gennaro Restaino⁹, Vincenzo Positano¹, and Alessia Pepe¹

¹CMR Unit, Fondazione G. Monasterio CNR-Regione Toscana, Pisa, Italy, ²Reparto di Ematologia, Azienda Sanitaria Provinciale Caltanissetta - Ospedale "Sant'Elia, Caltanisetta, Italy, ³Dip. di Oncologia, dei Trapianti e delle Nuove Tecnologie in Medicina – Divisione di Ematologia, Facoltà di Medicina e chirurgia – Università degli Studi di Pisa, Pisa, Italy, ⁴Hematology Unit, A.O. Bianchi-Melacrino-Morelli, Reggio Calabria, Italy, ⁵Dipartimento di Medicina Clinica e Sperimentale, Università degli Studi di Perugia, Perugia, Italy, ⁶UOC di Onco-Ematologia, Università Cattolica del Sacro Cuore, Campobasso, Italy, ⁷Struttura Complessa di Cardioradiologia-UTIC, P.O. "Giovanni Paolo II", Lamezia Terme, Italy, ⁸Dipartimento di Radiologia, Policlinico "Paolo Giaccone", Palermo, Italy, ⁹Dipartimento di Radiologia, Università Cattolica del Sacro Cuore, Campobasso, Italy

<u>Introduction</u>. In patients with myelodysplastic syndrome (MDS) no longitudinal studies on myocardial and hepatic iron and on cardiac function and fibrosis are available in literature. So, the aim of our study was to assess the changes in cardiac and hepatic iron overload and the morpho-functional cardiac parameters by Magnetic Resonance Imaging (MRI) in MDS patients enrolled in the MIOMED (Myocardial Iron Overload in MyElodysplastic Diseases) study who performed the follow-up (FU) MRI at 12 months.

Methods. MIOMED is an observational, MRI multicentre study in low and intermediate-1 risk MDS patients who have not received regular iron chelation therapy. Out of the 51 MDS patients enrolled, 48 underwent the baseline MRI exam and 28 performed the MRI FU. This analysis was limited to patients who performed both the MRIs. Mean age was 72.8±7.6 years and 8 patients were females. MIO was assessed using a multislice multiecho T2* approach. Hepatic T2* values were assessed in a homogeneous tissue area and converted into liver iron concentration (LIC). Biventricular function parameters were quantified by cine sequences. Myocardial fibrosis was evaluated by late gadolinium enhancement (LGE) acquisitions.

Results. The FU MRI was not performed for the following reasons: 4 deaths and 16 patient refusal.

At baseline only one patient showed cardiac iron overload (global heart T2*=14.8 ms) but he recovered at the FU (T2*=28.8 ms). He was not transfused. Out of the 27 patients without significant cardiac iron at the baseline, 26 maintained the same status at the FU while one showed cardiac iron (T2*=12.3 ms).

Thirteen patients (8 transfusion-dependent - TD) had hepatic iron overload (MRI LIC≥3mg/g/dw) at the baseline. For this subgroup, the baseline and the FU LIC values were, respectively, 14.9±12.0 mg/g/dw and 20.1±16.1 mg/g/dw. The increase in MRI LIC values was not significant (P=0.196). Out of the 11 patients with a baseline MRI LIC<3 mg/g/dw, two (13.3%) showed hepatic iron at the FU. Only one patient was TD but both patients had not received any chelation therapy.

Due mainly to technical reasons, biventricular function

60 MRI LIC (mg/g/dw) 40 Global heart T2* 50 40 30 20 10 0 0 FU 80 80 8 Ē § 70 70 ≥ 60 出 60 50 30 30 FU Baseline Baseline

was assesses at both baseline and FU MRIs in 22 patients. At baseline 6 patients showed a reduced left ventricular ejection fraction (LV EF) and 4 of them recovered at the FU. All patients had a baseline global heart T2*>20 ms (one with 2 segmental T2* values<20 ms). At baseline 5 patients showed a reduced right ventricular (RV) EF and all recovered at the FU. One patient with normal LV EF at baseline showed pathological LV EF at the and 2 patients with normal RV EF at baseline showed reduced RV EF at the FU. At the FU we detected a significant increase in the LV end-diastolic volume index (EDVI) (mean difference: 6.5±11.3 ml/m2; P=0.015) as well as in the RV EDVI (mean difference: 7.8±9.3 ml/m2; P=0.002). The change in the LV mass index was not significant.

For 18 patients the presence of myocardial fibrosis was investigated at both baseline and FU MRIs, and this subgroup was considered. Eight patients had myocardial fibrosis at the baseline. One patient showed a subendocardial ischemic pattern and seven patients showed a non-ischemic pattern and myocardial fibrosis was detected for all of them also at the FU. At the FU one new occurrence of non-ischemic myocardial fibrosis was detected.

<u>Conclusions.</u> The new occurrences of cardiac iron, reduced cardiac function, increased LV and RV EDVI and myocardial fibrosis and the worsening in MRI LIC values suggest the need of performing periodic MRI scans, in order to better manage these patients.

<u>References.</u> [1] Meloni A et al. Magn Reson Med 2010;64:211-9. [2] Meloni A et al. J Magn Reson Imaging 2011;33:348-55. [3] Marsella M et al. Haematologica 2011;96:515-20. [4] Pepe A et al. Heart 2009;95:1688-93.