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Title: Acute respiratory failure and severe sepsis may identify different clinical phenotypes in community-acquired pneumonia

Dr. Stefano 24211 Aliberti stefano.aliberti@unimib.it MD ¹, Dr. Anna Maria 24212 Brambilla brambi.brambi@gmail.com MD ², Dr. James D. 24213 Chalmers jamesdchalmers@googlemail.com MD ³, Dr. Catia 24214 Cilloniz catiacilloniz@yahoo.com MD ⁴, Prof. Angelo 24215 Bignamini angelo.bignamini@fastwebnet.it ⁵, Dr. Elena 24216 Prina eleprina@gmail.com MD ², Dr. Alice 24221 D'Adda alice.dadda@gmail.com MD ⁶, Dr. Marco 24222 Mantero mantero.marco@gmail.com MD ⁶, Prof. Dr Alberto 24224 Pesci alberto.pesci@unimib.it MD ¹, Prof. Dr Julio 24225 Ramirez j.ramirez@louisville.edu MD ⁷, Prof. Dr Antoni 24232 Torres ATORRES@clinic.ub.es MD ⁴, Prof. Dr Francesco 24239 Blasi francesco.blasi@unimi.it MD ⁶ and Dr. Roberto 24240 Cosentini r.cosentini@gmail.com MD ². ¹ Health Science Department, University of Milan Bicocca, AO San Gerardo, Milan, Italy ; ² Emergency Medicine Department, IRCCS Fondazione Cà Granda Ospedale Maggiore Policlinico, Milan, Italy ; ³ Tayside Respiratory Research Group, University of Dundee, Ninewells Hospital and Medical School, Dundee, United Kingdom ; ⁴ Department of Pneumology, Institut Clinic Del Tórax, Hospital Clinic of Barcelona - Institut D'Investigacions Biomèdiques August Pi I Sunyer (IDIBAPS), University of Barcelona, Barcelona, Spain ; ⁵ School of Specialization in Hospital Pharmacy, University Of Milan, Milan, Italy ; ⁶ Department of Pathophysiology and Transplantation, University of Milan, IRCCS Fondazione Ca' Granda Ospedale Maggiore Policlinico, Milan, Italy and ⁷ Division of Infectious Diseases, Department of Internal Medicine, University of Louisville, Louisville, KY, United States .

Body: The aim of the study was to evaluate the impact of acute respiratory failure (ARF) and severe sepsis (SS) on outcomes of patients with community-acquired pneumonia (CAP). This was a multicenter, observational, prospective study of immunocompetent CAP patients admitted to three hospitals in Italy, Spain and Scotland between 2008 and 2010. ARF was defined as the presence of at least one among: PaO₂<60 mmHg; PaO₂/FiO₂ ratio<250; oxygen saturation<90%; respiratory acidosis. SS was defined according to the 2008 Surviving Sepsis Campaign. Three groups were identified: patients with neither ARF nor SS (Group 1), those with only ARF (Group 2) and those with both ARF and SS (Group 3) on admission. A total of 2,271 consecutive CAP patients were enrolled (48% male, mean±SD age 68±19 yrs). ARF and SS were diagnosed in 969 (49%) and 473 (21%) patients, respectively. 242 (11%) patients died during hospitalization. Prevalence and mortality according to the study groups are depicted in Figure. At the multivariable logistic regression model, after adjustment for centers and confounders, the presence of only ARF had an OR for mortality of 2.5 (95%CI: 1.6-4.0, p<0.001) and the presence of both ARF and SS an OR of 5.9 (95%CI: 3.6-9.8, p<0.001). The significant difference in mortality in patients with or without ARF and

SS on admission may indicate different clinical phenotypes.