

## ERYTHROPOIETIN RESISTANCE INDEX AND MORTALITY OF HEMODIALYSIS PATIENTS-MULTICENTER STUDY

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**INTRODUCTION AND AIMS:** A considerable proportion of hemodialysis (HD) patients exhibit a suboptimal hematologic response to erythropoiesis-stimulating agents (ESA) and evidence suggests those patients with resistance (ERI) to have a higher mortality rate. This multi-center study was aimed to investigate the risk factors for ERI and association between ESA responsiveness and mortality in HD population.

**METHODS:** A prospective multicenter cohort study included seven hemodialysis centers with overall 603 patients on chronic HD for at least one year during 2015. We collected data on patient demographic factors, comorbidities, dialysis vintage and type, vascular access, body weight, ESA dose and hemoglobin concentration, as well as data on known risk factors for ESA hyporesponsiveness (mean values from multiple measurements). ERI was calculated by dividing the weekly body-weight-adjusted epoetin dose by the hemoglobin concentration. The association between ESA hyporesponsiveness and mortality was investigated by using the Cox proportional hazards model with adjustments for demographic factors, comorbidities, dialysis adequacy and serum biochemical data.

## RESULTS

## Table 1. Data on patients with different categories of ERI

|                   | ERI<10 (86%)       | ERI>10 (14%)      | р     |
|-------------------|--------------------|-------------------|-------|
| Age               | 63.3±12.41         | 61.8±13.1         | 0.301 |
| Males             | 63.7%              | 45.9%             | 0.003 |
| HD vintage, m     | 61 ±43             | 60±52             | 0.009 |
| DM, yes           | 20.3%              | 21.2%             | 0.855 |
| HT, yes           | 69.5%              | 81.2%             | 0.028 |
| Hepatitis, yes    | 13.5%              | 10.6%             | 0.602 |
| nonHDF/HDF        | 83.8% vs.16.2%     | 89.4% vs. 10.6%   | 0.255 |
| AVF/AVG/cath.     | 90%/2.3%/7.7%      | 87.1%/1.2%/11.8%  | 0.378 |
| ACE/ARB/both      | 43.7%/ 55.9% /0.4% | 29.6%/66.7% /3.7% | 0.002 |
| Promena doze      | 4.4±2.5            | 3.7±2.7           | 0.028 |
| Feritin           | 562±394            | 724±525           | 0.002 |
| Tsat              | 32±13              | 28±12             | 0.005 |
| Kt/V              | 1.32±0.32          | 1.32±0.53         | 0.827 |
| CRP               | 10.5±15.0          | 16.5±18.7         | 0.002 |
| PTH               | 239±310            | 237±317           | 0.951 |
| Albumin           | 36.0±3.7           | 36.3±2.9          | 0.490 |
| CCI               | 2.6±1.5            | 3.3±1.9           | 0.001 |
| Iron availability | 8.0±5.4            | 6.1±5.1           | 0.005 |

Table 2. Risk factor for ERI>10 (multivariat analysis)

| varijabla         | HR     | CI (95%)      | р     |
|-------------------|--------|---------------|-------|
| CRP >10           | 2.735  | 1.592-4.698   | 0.000 |
| Ž.pol             | 2.581  | 1.495-4.457   | 0.001 |
| HD staž           | 0.993  | 0.988-0.998   | 0.010 |
| ACE ili ARB       | 1.810  | 1.003-3.267   | 0.049 |
| ACE + ARB         | 11.641 | 1.073-126.273 | 0.044 |
| Iron availability | 0.918  | 0.848-0.993   | 0.032 |

During the one-year period, 92 (15.3%) patients died, 51% from CV reasons. Significant risk factors for Mt marked by Cox multivariate analysis



| <b>PREDICTOR</b> (univariate) | HR    | CI           | р     |
|-------------------------------|-------|--------------|-------|
| age                           | 1.038 | 1.019-1.057  | 0.000 |
| non-HDF treatment             | 3.346 | 1.358-8.241  | 0.009 |
| Hickman catheter              | 2.404 | 1.381-4-183  | 0.002 |
| Kt/V                          | 0.277 | 0.148-0.518  | 0.000 |
| albumin                       | 0.925 | 0.895-0.956  | 0.000 |
| ACE + ARB                     | 4.864 | 1.481-15.975 | 0.009 |

**CONCLUSION:** Although ERI above 10 caries 1.22 HR for Mt, its significance was not confirmed by multivariate analysis. Factors that cause an increase in ESA dose is likely to be of greater importance for mortality of HD patients than ERI.

