

IMPACT OF INTRAVENOUS HEPARIN ADMINISTERED PRIOR TO PRIMARY PERCUTANEOUS CORONARY INTERVENTION: ANALYSIS FROM THE HORIZONS-AMI TRIAL

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Background There is no consensus whether early administration of unfractionated heparin (UFH) is beneficial in pts prior to mechanical reperfusion for STEMI. We assessed outcomes of pts in HORIZONS-AMI trial in relation to UFH administration prior to randomization to bivalirudin (Biv) or UFH+a glycoprotein IIb/IIIa inhibitor (GPI).

Methods and Results In HORIZONS-AMI, randomization was stratified by the administration of UFH in the ER pre enrollment. Among a total of 3595 pts, 2357 pts received UFH pre randomization to Biv (n=1178) or to UFH+GPI (n=1179). A total of 1238 pts were not administered UFH pre randomization (619 pts in each arm). Administration of UFH pre randomization was associated with older age, > frequent use of clopidogrel loading dose of 600 mg, and > treated lesions. Pts who were administered UFH pre randomization had higher incidence of TIMI flow grade 2/3 on baseline both in Biv arm (37% vs 26%, p<0.0001) and UFH+GPI arm (39% vs 29%, p<0.0001). There was a significant interaction (Table) between UFH administration pre randomization and pharmacologic assignment; 30-day rates of MACE and NACE were significantly improved in Biv pts (but not UFH+GPI pts) who received UFH in the ER, without increase in major bleeding.

Conclusion In the HORIZONS-AMI trial, administration of UFH pre randomization to Biv or UFH+GPI was associated with reduced ischemic complications without an increase in bleeding events, benefits that were especially evident in pts treated with a bivalirudin monotherapy strategy.

| 30-day outcomes, % | Patients with STEMI n=3595 | | | | | | |
|---|--|---------------------------------------|-------------------------|--|---|--------------------|---------------|
| | Bivalirudin monotherapy n=1797 | | | UFH + GPI n=1798 | | | |
| | Pre-randomization UFH (+) n=1178 | Pre-randomization UFH (-) n=619 | HR [95%CI] | Pre- randomization UFH (+) N=1179 | Pre- randomization UFH (-) N=619 | HR [95%CI] | P interaction |
| Death | 2.0% | 2.3% | -0.31 [-1.72,1.11] | 3.2% | 2.9% | 0.32 [-1.35,1.99] | 0.57 |
| Reinfarction | 1.5% | 2.6% | -1.07 [-2.52,0.38] | 2.0% | 1.5% | 0.51 [-0.74,1.76] | 0.09 |
| Ischemic TVR | 1.8% | 3.9% | -2.11 [-3.82,- 0.40] | 2.0% | 2.0% | 0.01 [-1.34,1.37] | 0.09 |
| Major bleeding | 5.1% | 5.2% | -0.08 [-2.25,2.09] | 9.2% | 8.2% | 0.94 [-1.79,3.68] | 0.62 |
| Definite/probable stent thrombosis* | 2.5% | 5.7% | -3.22 [-5.39,- 1.04] | 2.1% | 2.2% | -0.08 [-1.60,1.44] | 0.12 |
| MACE† | 4.5% | 7.3% | -2.78 [-5.14,- 0.41] | 5.8% | 5.2% | 0.60 [-1.60,2.80] | 0.03 |
| Net adverse clinical events‡ | 8.6% | 10.8% | -2.25 [-5.18,0.68] | 13.6% | 11.0% | 2.59 [-0.56,5.74] | 0.03 |
| *By ARC definition; †Death, reinfarctio | n, ischemic TVR or str | oke; ‡MACE or major | bleeding | | | | |