The Utilization and Outcomes of Hypoxemic Donors in Combined Heart-Lung Transplantation

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Combined heart-lung transplantation (HLTx) is a complex yet life-saving procedure for patients with concomitant end-stage heart and lung failure.

While the ISHLT has found that HLTx recipients who survive the first year have favorable long-term survival with a conditional median survival of 12.8y¹, the impact of donor hypoxemia remains unknown in HLTx.

Prior studies have found that hypoxemic donor lungs, measured as PaO2/FiO2 (PF) ≤300mmHg, do not confer adverse outcomes in lung transplant.^{2,3}

However, this has not been investigated in HLTx or since UNOS modified the lung allocation score (LAS) system in 2017.

We aim to analyze the utilization and outcomes of hypoxemic donor lungs in HLTx before and after the modified LAS system.

Methods

We identified all adults (≥ 18yo) with data on donor P/F ratio undergoing HLTx in the US from 1/1/2010 – 4/30/2024.

Donor PF Ratio was used to group recipients into two categories: ≤300mmHg and >300mmHg.

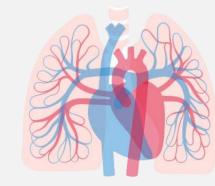
Recipients and matched donors were also stratified by before and after LAS modification: **Era 1** (1/1/10 – 11/24/17) vs. **Era 2** (11/25/17 – 4/30/2024)

Baseline recipient & donor characteristics were compared.

30-day, 1-, and 3-year survival were compared using the Kaplan-Meier method, stratified by both Era and PF Ratio.

Multivariate Cox proportional hazard regression models were used to assess the impact of PF Ratio on mortality.



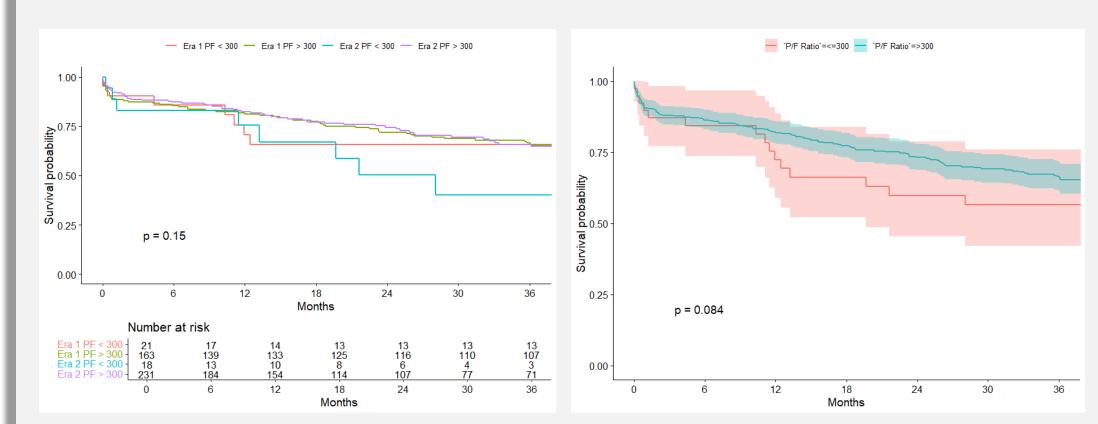


Results Characteristic 163 (41%) 184 (42%) 21 (53%) 255 (58%) 19 (48%) 236 (59%) 195.71 (187.11) 167.40 (173.42) 198.55 (188.39 45.25 (13.12) 44.73 (13.13) 0.007 50.43 (12.00) 0.10 219 (50%) 204 (51%) 15 (38%) 220 (50%) 25 (63%) 195 (49%) 66 (17%) 76 (17%) 10 (26%) 144 (33%) 15 (38%) 129 (33%) Cigarette Use 348 (87%) 382 (87%) 34 (85%) 51 (13%) 57 (13%) 6 (15%) 1.78 (0.24) 1.83 (0.28) 1.77 (0.24) 0.2 **LVAD** explant 1 (2.5%) 2 (0.5%) 3 (0.7%) 11 (2.5%) 2 (5.1%) 9 (2.3%) 97 (22%) 89 (22%) 4 (1.0%) **ECMO** explant 4 (0.9%) 0 (0%) 21 (4.8%) 1 (2.6%) 20 (5.1%) 32.45 (11.93) 32.45 (11.78) 32.45 (13.53) **Donor Age** 61.38 (6.82) 62.23 (7.74) 61.30 (6.73) 0.5 3.78 (1.02) 3.77 (1.09) 3.78 (1.01) 254 (58%) 33 (83%) 221 (56%) 94 (22%) 14 (38%) 80 (21%) 0.015 **Abnormal Bronchoscopy**

A total of 439 patients underwent HLTx (40 hypoxemic vs. 399 normal donors). Hypoxemic donor utilization has remained low between eras (21 cases in era 1 vs. 19 cases in era 2; p=0.2).

² Welch Two Sample t-test; Pearson's Chi-squared test

Recipients of hypoxemic donors were older (p=0.007). Hypoxemic donors had increased rates of abnormal chest x-ray (p=0.001) and bronchoscopy results (p=0.015) compared to normal donors.



There was no difference in survival between recipients of hypoxemic vs. normal donor lungs (p=0.084), regardless of era (p=0.15). Hypoxemic donors had 90.5, 70.6, and 65.6% survival in era 1, compared to 88.9, 75.4, and 40.2% survival in era 2 at 30-days, 1-, and 3-year intervals, respectively.



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| Post-HLTx Complications | Era 1 PF < 300, N = 21 ¹ | Era 1 PF > 300, N = 163 ¹ | Era 2 PF < 300, N = 19 ¹ | Era 2 PF > 300, N = 236 ¹ | p- value ² | Post-HLTx Complications | <=300, N = 40¹ | >300, N = 399 ¹ | p-value ² |
|-------------------------|-------------------------------------|--------------------------------------|---|--------------------------------------|--------------------------|-------------------------|-------------------|----------------------------|----------------------|
| Acute Rejection | 3 (14%) | 25 (15%) | 0 (0%) | 28 (12%) | 0.3 | Acute Rejection | 3 (7.7%) | 53 (13%) | 0.3 |
| Length of Stay | 34.95 (29.29) | 44.13 (56.64) | 44.24 (41.24) | 49.62 (53.58) | | Length of Stay | 39.22 (35.08) | 47.36 (54.86) | 0.2 |
| Pacemaker | 0 (0%) | 3 (1.9%) | 0 (0%) | 4 (1.7%) | >0.9 | Pacemaker | 0 (0%) | 7 (1.8%) | >0.9 |
| Stroke | 1 (5.0%) | 9 (5.6%) | 1 (5.6%) | 11 (4.8%) | >0.9 | Stroke | 2 (5.3%) | 20 (5.1%) | >0.9 |
| Dialysis | 4 (20%) | 48 (30%) | 6 (33%) | 74 (32%) | 0.7 | Dialysis | 10 (26%) | 122 (31%) | 0.5 |
| Airway Dehiscence | 0 (0%) | 1 (0.6%) | 0 (0%) | 9 (3.9%) | 0.2 | Airway Dehiscence | 0 (0%) | 10 (2.6%) | 0.6 |

In both eras, rates of acute rejection, length of stay, pacemaker implantation, stroke, dialysis, and airway dehiscence were similar between HLTx recipients with hypoxemic vs. normal donor lungs.

| Covariates | Hazard Ratios | 95 % Confidence Intervals | p-value |
|-----------------------|---------------|------------------------------|---------|
| PF>300 | 0.64 | 0.40 - 1.04 | 0.073 |
| Era 2 | 1.20 | 0.85 - 1.70 | 0.300 |
| Abnormal Chest X Ray | 0.62 | 0.45 – 0.84 | 0.002 |
| Abnormal Bronchoscopy | 0.86 | 0.59 – 1.26 | 0.444 |
| Recipient Age | 1.00 | 0.99 - 1.02 | 0.465 |

After adjusting for covariates, neither PF>300mmHg (HR 0.64, p=0.073) nor Era 2 (HR 1.20, p=0.300) had an impact on HLTx recipient survival.

Conclusions

Since 2017, hypoxemic donors continue to be underutilized in HLTx despite comparable long-term outcomes to normal donors. However, due to the small number of patients receiving hypoxemic donor lungs in HLTx, further studies are warranted to better interpret these results.

References

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