



# Impact of Regional Organ Sharing and Allocation in the UK Northern Liver Alliance (NLA) on Waiting Time to Liver Transplantation and Waitlist survival



Malik AK, Masson S, Allen E, Akyol M, Bathgate A, Davies M, Hidalgo E, Hudson M, Powell J, Zarankaite A, Manas DM











# Background

- Deceased-donor liver grafts (DDL) have been allocated in a centrebased system until recently
- A supra-regional organ allocation system (NLA) was established in 2006 to improve access to DDLs for the sickest patients awaiting liver transplantation
  - 3 Northern centres (Edinburgh, Newcastle and Leeds)
    - Patients with UKELD ≥62\* listed on 'top-band'
    - Prioritisation by UKELD score
    - Organs shared between centres
    - Organ 'payback' scheme
- Scheme mirrors "Share-35" system in US
  - Implemented from 2013
- We aimed to investigate the impact on waiting list outcomes

## Methods

- Data retrospectively extracted from UK transplant registry (NHSBT)
  - Apr 2013 to Dec 2016
  - NLA centres compared with two non-NLA centres (King's and Cambridge liver transplant units)
  - Changes in UKELD captured by sequential data
- Adult patients registered for first DDL transplant included in analysis
- Once patients are registered into top-band, they are not removed
- Periods of suspension from WL not included in WT

## Methods, cont.

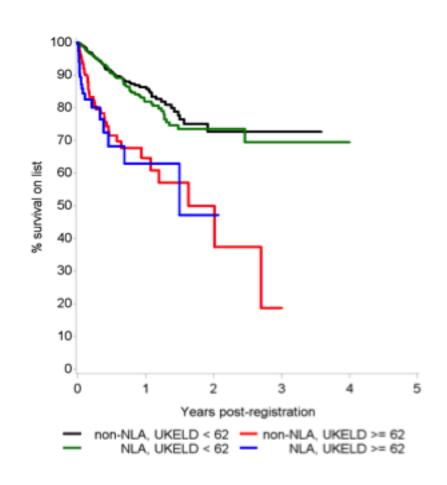
- Kaplan-Meier method used to estimate WL survival and WT to transplantation
- WL survival patients who were suspended, removed from WL (non-transplant reason), or transplanted were censored
- WT patients removed/suspended or died on WL were censored
- Log-rank test used for comparisons
  - Bonferroni correction for multiple testing
- Cox proportional hazards model used to ascertain impact of WT on post-transplant survival
  - All 7 liver transplant units included
  - Adjusted with risk factors for post-transplant mortality

## Results

Transplant centre	Non-top-band	Top-band	Total
NLA centres	880	159	1039
Newcastle	129	32	161
Edinburgh	319	62	381
Leeds	432	65	497
Non-NLA centres	923	200	1123
Cambridge	322	65	387
King's	601	135	736
Total	1803	359	2162

#### WL survival

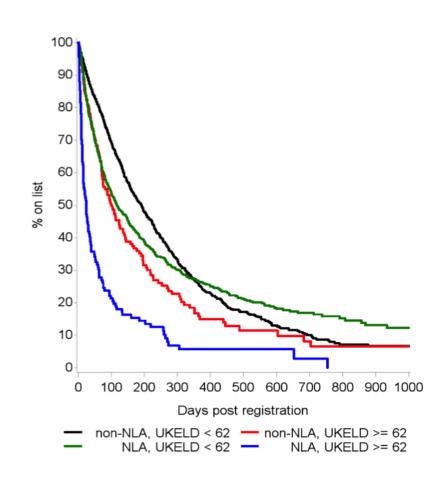
- NLA top-band vs non-NLA top-band p=0.9999
  - 62.9 vs. 64.6%
- NLA top-band vs NLA non-top-band p<0.0001</li>
  - 62.9 vs. 81.9%
- No difference between NLA non-top-band and non-NLA non-top-band
  - 81.9 vs. 86.4%



Significantly different amongst 4 groups (p<0.0001)

# WT to transplantation

- Median WT NLA top-band shorter compared to non-NLA top-band
  - 23 vs. 99 days
  - p<0.0001
- Median WT non-top-band no different between NLA and non-NLA
  - 117 vs. 192 days
  - p=0.2288



Significantly different among 4 groups (p<0.0001)

#### Impact of WT on post-transplant survival in topband patients

- 315 top-band transplants from Apr 2013 to Dec 2016
  - 273 included in analysis
  - 42 excluded from analysis due to missing survival/risk factor data, or auxiliary transplant
- WT has no significant impact on 3-year risk-adjusted posttransplant survival (p=0.712)
  - 1-month increase in WT is associated with 4.6% increase in risk of death
  - HR 1.046 (95% CI 0.825-1.327)

## Discussion

- NLA significantly shortened WT for top-band patients
- No improvement in WL survival
  - No adverse impact upon non-top-band patients either
- WT did not impact long-term survival in top-band patients
- Results can be seen as an ethically positive outcome
  - Sicker patients warrant special priority, irrespective of potentially lower 'benefit'/utility
  - Mirrors US organ allocation policy and 'final rule'

## Discussion, cont.

- Survival benefit may not have been detected as each centre ensures timely transplantation of sickest patients
  - Patients censored at transplantation type 2 error
- Initial analysis of Share-35 reported 30% reduction in WL mortality in patients with MELD ≥35
  - Patients with MELD <35 not adversely affected</li>
  - Analyses limited through historical controls
- Under Share-35, WL patients reprioritised were not disadvantaged by losing allocation
  - Similar analysis into NLA not possible retrospectively

## Conclusions

- NLA achieved its aim of improving DDL transplantation access to those most in need
- Prioritisation of sickest patients did not improve WL survival
  - Did not disadvantage less sick patients
- Similar experience to US
- NLA will be absorbed into UK national allocation scheme based on transplant benefit score
  - Similar analysis into outcomes of patients UKELD ≥62 after national allocation warranted once long-term followup data sufficiently available

## References

- 1. Massie AB, Chow EK, Wickliffe CE, Luo X, Gentry SE, Mulligan DC, et al. Early changes in liver distribution following implementation of Share 35. Am J Transplant. 2015;15(3):659-67.
- 2. Edwards EB, Harper AM, Hirose R, Mulligan DC. The impact of broader regional sharing of livers: 2-year results of "Share 35". Liver Transpl. 2016;22(4):399-409.
- 3. Murken DR, Peng AW, Aufhauser DD, Jr., Abt PL, Goldberg DS, Levine MH. Same policy, different impact: Center-level effects of share 35 liver allocation. Liver Transpl. 2017;23(6):741-50.
- 4. Schaubel DE, Guidinger MK, Biggins SW, Kalbfleisch JD, Pomfret EA, Sharma P, et al. Survival benefit-based deceased-donor liver allocation. Am J Transplant. 2009;9(4 Pt 2):970-81.
- 5. Barber K, Madden S, Allen J, Collett D, Neuberger J, Gimson A, et al. Elective liver transplant list mortality: development of a United Kingdom end-stage liver disease score. Transplantation. 2011;92(4):469-76.
- 6. Rauchfuss F, Zidan A, Scheuerlein H, Dittmar Y, Bauschke A, Settmacher U. Waiting time, not donor-risk-index, is a major determinant for beneficial outcome after liver transplantation in high-MELD patients. Ann Transplant. 2013;18:243-7.
- 7. NHS Blood and Transplant. Annual Report on Liver Transplantation 2015/2016 2016 [cited 2017 28/12/17]. Available from: <a href="https://nhsbtdbe.blob.core.windows.net/umbraco-assets-corp/1314/organ\_specific\_report\_liver\_2016.pdf">https://nhsbtdbe.blob.core.windows.net/umbraco-assets-corp/1314/organ\_specific\_report\_liver\_2016.pdf</a>.
- 8. Howard D. The impact of waiting time on liver transplant outcomes. Health Serv Res. 2000;35(5 Pt 2):1117-34.
- 9. Chow EK, Massie AB, Luo X, Wickliffe CE, Gentry SE, Cameron AM, et al. Waitlist Outcomes of Liver Transplant Candidates Who Were Reprioritized Under Share 35. Am J Transplant. 2017;17(2):512-8.
- 10. Nicolas CT, Nyberg SL, Heimbach JK, Watt K, Chen HS, Hathcock MA, et al. Liver transplantation after share 35: Impact on pretransplant and posttransplant costs and mortality. Liver Transpl. 2017;23(1):11-8.

# Acknowledgments

- Professor Derek M Manas
- Dr Steven Masson
- Dr Elisa Allen
- Agne Zarankaite
- The NLA
- The HPB and Transplant unit at the Freeman Hospital, Newcastle-upon-Tyne

# Acknowledgments

 The organ donors, who with their passing, provide the ultimate gift

