Pre-hospital determinants of successful resuscitation after traumatic and nontraumatic out-of-hospital cardiac arrest

List of Authors

Ed B G Barnard (1,2) Daniel D Sandbach (1) Tracy L Nicholls (3) Alastair W Wilson (1) Ari Ercole (1,4)

1. East Anglian Air Ambulance, Hangar E, Gambling Close, Norwich Airport, Norwich, UK

2. Academic Department of Military Emergency Medicine, Royal Centre for Defence Medicine (Research & Clinical Innovation), Birmingham, UK

3. East of England Ambulance Service NHS Trust, Whiting Way, Melbourn, UK

4. University of Cambridge Division of Anaesthesia, Addenbrooke's Hospital, Cambridge, UK

Corresponding Author:

Dr. Ed Barnard PhD East Anglian Air Ambulance Hangar E Gambling Close Norwich Airport Norwich, NR6 6EG UK

Tel: 07946 021207 (not for dissemination) Email: ukbarnard@gmail.com

Keywords: cardiac arrest; traumatic cardiac arrest; out-of-hospital cardiac arrest; emergency medical services; pre-hospital care

Word Count (excluding title, abstract, tables, and references): 3141

ABSTRACT

Background

Out-of-hospital cardiac arrest (OHCA) is prevalent in the UK. Reported survival is lower than in countries with comparable healthcare systems; a better understanding of outcome determinants may identify areas for improvement.

Methods

An analysis of 9,109 OHCA attended in East of England between 1st January 2015 and 31st July 2017. Univariate descriptives and multivariable analysis were used to understand determinants of survival for non-traumatic cardiac arrest (NTCA) and traumatic cardiac arrest (TCA). Two Utstein outcome variables were used: survival to hospital admission, and hospital discharge.

Results

The incidence of OHCA was 55.1 per 100,000 population/year. The overall survival to hospital admission was 27.6% (95%CI 26.7-28.6) and the overall survival to discharge was 7.9% (95%CI 7.3-8.5). Survival to hospital admission and survival to hospital discharge were both greater in the NTCA group compared with the TCA group: 27.9% versus 19.3% p=0.001, and 8.0% versus 3.8% p=0.012 respectively.

Determinants of NTCA and TCA survival were different, and varied according to the outcome examined. In NTCA, bystander-CPR was associated with survival at discharge but not at admission, and the likelihood of bystander CPR was dependent on geographical socioeconomic status. An air ambulance was associated with increased survival to both hospital admission and discharge in NTCA, but only with survival to admission in TCA.

Conclusion

NTCA and TCA are clinically distinct entities with different predictors for outcome – future OHCA reports should aim to separate arrest aetiologies. Determinants of survival to hospital admission and discharge differ in a way that likely reflects the determinants of neurological injury. Bystander-CPR public engagement may be best focused in more deprived areas.

INTRODUCTION

Every year the ambulance services in England attempt resuscitation on approximately 30,000 cases of out of hospital cardiac arrest (OHCA).[1] Survival to hospital discharge in England is less than 10% -lower than that of several other countries with developed emergency healthcare systems,[2] indicating the need for improvement. Key to addressing this disparity is a better understanding of determinants of outcome.

Demographic data on OHCA in England is routinely collected as part of the Out-of-Hospital Cardiac Arrest Outcome project,[3] providing a national, linkable source of epidemiological data. This project has reported a 25% survival to hospital admission, and a survival to discharge of 8%. However, it has not modeled the associations between pre-hospital variables and survival outcomes,[2] a process which could highlight opportunities for system improvements. Achieving a return of spontaneous circulation (ROSC) cannot be of benefit if the individual sustains a neurological injury from which they subsequently die; the pre-hospital phase plays an important role in this and deserves particular attention. Neurological outcome is critically determined by ambulance response time,[4] and there are many prehospital interventions (initially, and during post-ROSC care) that may be important in determining outcome. These interventions potentially require specialist medical teams that are both expensive and limited in number. It is therefore important that they are properly evaluated and appropriately used.[5]

Whilst research into management of OHCA is constrained by its very nature (unpredictable, resource-limited, in an uncontrolled environment),[6] the wide inclusion of all cause OHCA in epidemiological studies potentially leads to a lack of clarity when reporting determinants of outcome. For example, traumatic cardiac arrest (TCA), predominantly caused by traumatic brain injury and haemorrhage,[7] is a fundamentally different disease process to non-traumatic cardiac arrest (NTCA) that is largely of primary cardiac aetiology.[8] Therefore, it would be expected that determinants of outcome are different for TCA and NTCA. However, most previous studies, including the Out-of-Hospital Cardiac Arrest Outcome project,[3] do not differentiate these conditions. There has been a substantial increase in reported survival from TCA since 2005.[9] It is likely that a better understanding of this disease process, distinct from NTCA, has played a part in this improvement, but there remains a paucity of data comparing determinants of outcome from NTCA and TCA in the same patient cohort.

Our primary aim was to compare the differential determinants of survival to hospital admission and survival to hospital discharge for non-traumatic and traumatic cardiac arrest in a large regional cohort of OHCA.

METHODS

Emergency Medical Service

East of England is a geographic area of 20,000 km², containing a population of 6,395,000 (June 2016).[10] The East of England Ambulance Service NHS Trust (EEAST) is the statutory emergency medical service for this area and receives over one million emergency calls per year to three Emergency Operations Centres. OHCA cases are assigned an immediate dispatch of a minimum of two double-staffed ambulances, including at least one paramedic. In addition, EEAST utilises layperson community first responders, police and fire service co-responders, and British Association for Immediate Care Scheme responders. All carry automated external defibrillators (AED) and are trained in basic life support as a minimum standard. These assets, as well as ambulance service rapid response vehicles, are used to meet the statutory eight-minute response standard.[11] Dispatch (by helicopter or car) of one of five physician-paramedic pre-hospital critical care teams in the East of England is at the discretion of the paramedic-led critical care desk at one of the EEAST Emergency Operations Centres.

Data collection and variable definitions

Data were obtained from EEAST for all cases of OHCA in which a resuscitation attempt was made between 1st January 2015 and 31st July 2017; EEAST only capture data in cases in which the ambulance service attempt resuscitation.[12] These data are routinely collected, in an Utstein 2004 template,[13] for submission to UK Government and the Out-of-Hospital Cardiac Arrest Outcome Project. Neurological outcome at discharge from hospital is not routinely recorded. The presence of an 'air ambulance' (pre-hospital critical care team, which may have been deployed by helicopter or rapid response vehicle) is also recorded.

Complete postcode data of the emergency call were only available for 2015 (from 2016 onwards data governance procedures precluded the storage of a complete postcode, reducing the accuracy of these data). From these, we obtained indices of social deprivation from Ministry of Housing, Communities and Local Government data.[14]

The data were dichotomised into NTCA and TCA. NTCA, is dominated by all-cause medical cardiac arrests, but also includes asphyxiation, drowning and electrocution. TCA is defined as cardiac arrest resulting from an external application of kinetic energy.

Outcomes

In keeping with the international consensus reporting guidelines for resuscitation (Utstein), we have reported two patient outcomes: 'survival to hospital admission'

and 'survival to hospital discharge'.[15] Survival to hospital in the absence of longer-term survival is clearly not of benefit to the patient. However, in addition to the adherence to international data standards, the authors perceive substantial benefit to pre-hospital providers (who are not routinely informed of final patient outcome) in including the 'survival to hospital admission' outcome in this report.

Statistical analyses

Utstein 2004 variables (excluding neurological outcome), presence of an air ambulance medical team, postcode, and indices of social deprivation were refactored into putative explanatory variables of interest after which <<10% of the total was missing, except for social deprivation status where approximately 66% (i.e. 2016 and 2017 patients) was not available. The response time was defined as emergency call origin to ambulance arrival at scene. Patients who had a cardiac arrest with the ambulance crew present (i.e. those for which the original call was not a cardiac arrest but where the patient subsequently required CPR) had the response time set to zero, rather than the response time of the initial call. Missing data were imputed under the assumption of missingness-at-random; predictive mean matching was used to generate 100 imputed datasets for each of the NTCA and TCA subsets.[16]

Multivariable logistic regression models were then constructed for the NTCA subgroup and pooled across imputations. Plausible covariates and interactions were introduced and the model successively simplified by eliminating predictors in such a way as to retain those that were significant for at least one outcome. To assess differences between the non-trauma (NTCA) and trauma (TCA) groups, this final model was then applied to the TCA group to assess its performance before further simplification, again to the point that only predictors with statistical significance for at least one outcome measure were retained. We assumed a significance level of 5%.

Data manipulation and statistical analyses were performed using the R statistical programming language (R Core Team (2018); R: A language and environment for statistical computing (R Foundation for Statistical Computing, Vienna, Austria). Statistical significance (*p*-values) are presented without correction for multiple comparison.

Data have been reported as number (percentage), number (percentage (95% confidence interval)), and median [interquartile range] as appropriate. Continuous data have been analysed with a Mann-Whitney U test, and categorical data have been analysed with a Chi-squared test.

Ethical review

Institutional approval was granted by EEAST as a service evaluation (EEAST/1819/0019) in order to access, analyse and publish these anonymised data, and therefore specific ethical review was waived.

RESULTS

Total population

There were 9,109 OHCA cases during the period studied. Overall, the median age was 73 years [60-83], and 5,721 (62.8%) were male. The incidence of OHCA was 55.1 per 100,000 population/year. The survival to hospital admission was 27.6% (95%CI 26.7-28.6), and the survival to hospital discharge was 7.9% (95%CI 7.3-8.5). The median response time (defined as emergency call origin to ambulance arrival at scene) was 6.3 [4.0-10.0] minutes; 67.1% were within the statutory eight-minute standard.[11]

Comparison of NTCA and TCA cohorts

The majority (*n*=8,805) of patients had suffered NTCA, Table 1. The incidence of NTCA per 100,000 population/year was 53.3, Fig. 1 (left panel). 304 patients were reported as TCA, Table 1. The incidence of TCA per 100,000 population/year was 1.8, Fig. 1 (right panel).

Table 1 – Descriptive characteristics of non-traumatic and traumatic cardiac arrest cohorts, with outcomes.

Variable	NTCA	ТСА	<i>p</i> -value
Total, <i>n</i> (%)	8,805 (96.7)	304 (3.3)	
Median age, years [IQR]	74 [61-83]	43 [30-61]	< 0.001 ***
Gender, male (%)	62.5	77.0	< 0.001 ***
Witnessed arrest, n (%)	4,950 (56.4)	103 (34.0)	< 0.001 ***
- not recorded, <i>n</i>	24	1	
Bystander-CPR, n (%)	4,719 (54.6)	156 (52.2)	0.40 NS
- not recorded, <i>n</i>	166	5	
First monitored rhythm			
- Asystole, n (%)	4,689 (56.0)	204 (70.6)	< 0.001 ***
- PEA, n (%)	1,862 (22.3)	69 (23.9)	0.52 NS
- VF, n (%)	1,697 (20.3)	14 (4.8)	< 0.001 ***
- VT, n (%)	87 (1.0)	1 (0.3)	0.25 NS

- other, <i>n</i> (%)	32 (0.4)	1 (0.3)	0.92 NS
- not recorded, <i>n</i>	438	15	
Survival			
- admission, <i>n</i> (%,	2,423 (27.9,	58 (19.3,	< 0.001 ***
95%CI)	95%CI 27.0-28.9)	95%CI 15.3-24.2)	
- discharge, <i>n</i> (%,	623 (8.0,	10 (3.8,	0.012 *
95%CI)	95%CI 7.4-8.7)	95%CI 2.1-6.8)	

NTCA – non-traumatic cardiac arrest, TCA – traumatic cardiac arrest, IQR – interquartile range, PEA – pulseless electrical activity, VF – ventricular fibrillation, VT – ventricular tachycardia.

The NTCA cohort were significantly older, contained a higher proportion of female patients, and were more likely to have had a witnessed arrest compared to the TCA cohort; rates of bystander-CPR were comparable, Table 1. The most prevalent initial cardiac rhythm in both groups was asystole. However, the NTCA cohort contained a significantly smaller proportion of asystole and a significantly larger proportion of VF compared to the TCA cohort, Table 1. Survival to hospital admission and survival to hospital discharge were both significantly higher in the NTCA group compared to the TCA group, Table 1 & Fig. 2.

Multivariable analysis

Non-traumatic cardiac arrest

The final logistic regression models for NTCA survival are summarised in Table 2.

Table 2 - Results of multivariable logistic regression on imputed dataset for NTCAsurvival to hospital admission or survival to receiving hospital discharge outcomes.

Covariate	Survival to admis	-	Survival to hospital discharge	
	effect	<i>p</i> -value	effect	<i>p</i> -value
(Intercept)	-0.860	<0.001	-1.20	< 0.001
Age	0.0105	0.090	0.0204	0.068
Age ²	-0.000144	0.0061	-0.000483	<0.001
Male sex	-0.167	0.0025	0.297	0.021
Arrest witnessed	0.752	< 0.001	0.79	<0.001
Response time (minutes)	-0.0356	<0.001	-0.108	< 0.001
Shockable initial rhythm	1.27	<0.001	2.512	< 0.001
Air ambulance	0.676	<0.001	0.527	0.0023
Response time:Bystander CPR	-0.00445	0.57	0.04	0.049
Successfully intubated	0.122	0.039	-0.77	<0.001
Adrenaline administered	-0.81	<0.001	-2.15	< 0.001

Models were simplified by sequential elimination of terms and interactions that were not significant for at least one of the two outcomes. Significant covariate estimates are denoted in **bold** assuming a significance level of 5% and estimates are log-odds. CPR – cardiopulmonary resuscitation.

Terms up to quadratic for age were included as age was found to have a strong nonlinear relationship with outcome. Index of multiple deprivation and bystander-CPR were found to be strongly collinear, and therefore only the latter was retained on mechanistic grounds. Whether the patient had been defibrillated prior to ambulance arrival (n=141) was not statistically significant and eliminated. We found no evidence of statistically significant interactions between response time and either a shockable initial rhythm, or whether the arrest was witnessed. Furthermore, we could find no evidence that cardiac arrests occurring at weekends carried a statistically different outcome, and this variable was eliminated.

Age, male sex, response time, and the documented administration of adrenaline (epinephrine) were all associated with adverse hospital arrival outcome, whereas the presence of a shockable initial rhythm, the attendance of an air ambulance, and successful intubation were positively associated. The pattern is different for survival to hospital discharge: in this case, male sex was now found to carry a survival advantage and successful intubation to be associated with mortality. However, the presence of an air ambulance remained significantly associated with survival to hospital discharge. The documented presence of bystander-CPR only entered into the model via its interaction with response time, but this was only significantly associated with survival to discharge, not admission.

The results presented, using multiple imputation, were consistent with a total case analysis, but for clarity only the final, most robust, model has been included; for completeness, the summary data (before multiple imputation) for the NTCA cohort is presented in Table 3.

	Survived to hospital admission	Did not survive to hospital admission	Survived to hospital discharge	Survived to admission but not discharge
Median age, years [IQR]	71 [58-81]	75 [62-84]	63 [50-73]	74 [62-83]
 Male, n (%)	1,551 (17.6)	3,939 (44.7)	411 (4.7)	663 (7.5)
Female, n (%)	868 (9.9)	2,422 (27.5)	119 (1.4)	418 (4.7)
Witnessed arrest, n (%)	1,752 (19.9)	3,198 (36.3)	440 (5.0)	763 (8.7)
Unwitnessed arrest, n (%)	666 (7.6)	3,165 (35.9)	90 (1.0)	319 (3.6)
Median response time, minutes [IQR]	5.0 [3.0-8.0]	6.0 [3.9-9.0]	4.4 [1.0-7.0]	6.0 [3.1-8.0]

Table 3 – Summary statistics (before multiple imputation) for significant predictors in the NTCA model. Percentages are of the total *n* of NTCA patients.

Shockable rhythm, n (%)	911 (10.3)	775 (8.8)	373 (4.2)	282 (3.2)
Non-shockable rhythm, n (%)	1,319 (15.0)	5,362 (60.9)	98 (1.1)	737 (8.4)
Successfully intubated, n (%)	916 (10.4)	2,228 (25.3)	121 (1.4)	489 (5.6)
Not intubated, n (%)	1,326 (15.0)	3,683 (41.8)	369 (4.2)	558 (6.3)
Adrenaline, n (%)	1,689 (19.2)	5,462 (62.0)	187 (2.1)	895 (10.2)
No adrenaline, n (%)	735 (8.3)	915 (10.4)	343 (3.9)	188 (2.1)

Traumatic cardiac arrest

The results of applying the final NTCA multivariable model to the TCA cohort is shown in Table 4.

Table 4 - Results of multivariable logistic regression on imputed dataset for TCA survival to hospital admission or survival to receiving hospital discharge outcomes.

Covariate	Survival to hospital admission effect <i>p</i> -value		Survival to hospital discharge		
			effect	<i>p</i> -value	
(Intercept)	-0.970	0.0014	-2.89	<0.001	
Arrest witnessed	-0.76	0.035	-0.784	0.26	
Response time (minutes)	-0.072	0.057	-0.00944	0.89	
Shockable initial rhythm	0.777	0.22	2.49	0.0030	
Air ambulance	0.952	0.012	0.180	0.83	

Models were simplified by sequential elimination of terms and interactions that were not significant for at least one of the two outcomes. Significant covariate estimates are denoted in **bold** assuming a significance level of 5% and estimates are log-odds.

For the hospital admission outcome, only the presence of an air ambulance was associated with survival, but this did not translate to improved survival to hospital discharge, and there was a negative association with survival to hospital for witnessed TCA, Table 4. For the survival to hospital discharge end-point, only an initial shockable rhythm was significant. The median [interquartile range] age of TCA patients with and without a shockable rhythm was 63 [30-82] years and 43 [30-60] years respectively, p=0.025. Summary data (before multiple imputation) is presented in Table 5.

	Survived to hospital admission	Did not survive to hospital admission	Survived to hospital discharge	Survived to admission but not discharge
Witnessed arrest, n (%)	13 (4.3)	90 (30.0)	2 (0.7)	5 (1.6)
Unwitnessed arrest, n (%)	44 (14.5)	156 (51.3)	6 (2.0)	19 (6.3)
Median response time, minutes [IQR]	5.0 [3.0-7.8]	6.0 [4.0-9.7]	4.0 [3.5-7.3]	5.0 [2.5-6.6]
Shockable rhythm, n (%)	4 (1.3)	11 (3.6)	3 (1.0)	0 (0)
Non-shockable rhythm, n (%)	50 (16.4)	224 (73.7)	4 (1.3)	25 (8.2)
Air ambulance, n (%)	14 (4.6)	29 (9.5)	1 (0.3)	8 (2.6)
No air ambulance, n (%)	44 (14.5)	217 (71.4)	7 (2.3)	17 (5.6)

Table 5 – Summary statistics (before multiple imputation) for significant predictors in the TCA model. Percentages are of the total *n* of TCA patients.

DISCUSSION

Our study has demonstrated significant differences in the epidemiology, presentation, and predictors of outcome between non-traumatic and traumatic cardiac arrest. Furthermore, we have identified differences in the pre-hospital determinants of survival to hospital admission and survival to hospital discharge.

The overall incidence of OHCA in the East of England during the study period is comparable to England in 2014: 55.1 and 53.2 per 100,000 population/year respectively.[2] The overall survival to hospital arrival was significantly higher than national data (27.6% versus 25.8%, *p*<0.001), but the survival to discharge was the same (both 7.9%).[2] This suggests that even though a higher proportion of patients are initially 'successfully' resuscitated in the East of England, this does not translate to greater survival to hospital discharge. Our figure of 3.8% survival to discharge following TCA is lower than the 7.5% recently reported in England and Wales (which excluded patients pronounced life extinct pre-hospital);[7] our figure is therefore likely to be a more accurate total population estimate.[17]

Comparison of NTCA and TCA

Comparison of NTCA and TCA cohorts demonstrated that TCA patients were younger and more likely to be male. This is not a surprising result, and similar findings have recently been reported.[18] However, we are unable to explain why TCA was less likely to be witnessed than NTCA; it might be that a proportion of NTCA patients were unwell and therefore sought assistance prior to arrest, whereas TCA had a more sudden onset. The only pre-hospital variable associated with increased survival to hospital discharge in both NTCA and TCA cohorts was a 'shockable initial rhythm'. An initial shockable rhythm is well known to increase the chance of survival in NTCA, but this finding is puzzling following TCA, in which shockable rhythms are rare.[19] It is possible that this group comprises patients in whom the 'trauma' was preceded by a primary cardiac OHCA, which would most likely be a ventricular dysrhythmia (VF/VT), and as such would have a 'more-survivable' underlying aetiology. The TCA group with a shockable rhythm were significantly older than those without, and emphasizes the need for caution when applying bespoke TCA management protocols to older patients. Therefore, the only common pre-hospital determinant of survival to discharge between NTCA and TCA is potentially due to misdiagnosis, and together with the differences observed in epidemiology and presenting cardiac rhythm, provides evidence that these are clinically distinct entities that should be analysed separately and managed differently.

Comparison of survival outcomes

It is not possible for a patient to survive to hospital discharge without a return of spontaneous circulation. Conversely, survival to hospital admission cannot be of benefit, and is potentially harmful, if the patient does not survive to hospital discharge. Determinants of survival at the two time points were found to be different in both NTCA and TCA cohorts. In NTCA, male sex conferred a survival disadvantage at hospital admission but a survival advantage at discharge. This is presumably owing to a difference in the aetiologies between male and female OHCA. Documented bystander-CPR reduced the negative effect of longer response times; this is consistent with previous findings that bystander-CPR does not improve neurological outcome in short response times.[20] That this conferred a survival benefit at hospital discharge but not at admission has not to our knowledge been previously described. This is likely to reflect a greater degree of neurological injury (probably the dominant cause of ultimate death) in patients with a period without CPR, whereas ROSC is still obtained in such patients due to the relative tolerance of the heart to ischaemia; this matters less when response times are short.

Successful pre-hospital intubation was positively associated with survival to hospital admission, but negatively associated with survival to discharge. This is likely to reflect the effectiveness of resuscitation to achieve a ROSC even after a period of arrest that is neurologically devastating, and is in keeping the findings of the recent AIRWAYS-2 trial.[21] This trial demonstrated no difference in neurologically-intact survival between supraglottic airway (SGA) use and intubation, but also included n=1,707 (18.4%) patients who did not receive an advanced airway (owing to a short-duration arrest), 21.1% of these patients had a good outcome compared to 3.3% of those that received an advanced airway.[21] In addition, there was a higher proportion of patients in the intubation arm that did not receive an advanced airway compared to those randomised to the SGA arm (985/4410, 22.3% and 722/4886, 14.8% respectively), indicating that OHCA

patients who are intubated are likely to have had a longer duration arrest, and by inference a worse outcome.

A negative association between the administration of pre-hospital adrenaline and outcome has been previously reported.[22] The recent randomised trial PARAMEDIC-2 demonstrated a 30-day survival advantage for adrenaline versus placebo, but no difference in neurological-intact survival.[23] This suggests that whilst adrenaline administration may increase survival this does not translate to meaningful survival in OHCA. This trial excluded patients who had a ROSC during initial resuscitation (before adrenaline was indicated), and are therefore a cohort of longer duration arrests that would be expected to have a worse outcome. Our data also included patients who had an early ROSC, in whom adrenaline was not administered, which might explain our findings that adrenaline administration was associated with worse outcomes (compounded by the length of arrest).

The presence of an air ambulance was strongly associated with improved survival at both time points in NTCA and at admission in TCA. It would be tempting to attribute this to the critical care skills of air ambulance providers, and previous literature has suggested this.[24] However, in the East of England this is also likely to represent effective triage of patients who are more likely to survive. In particular, we cannot determine how many of the patients attended by an air ambulance had already obtained ROSC, with the critical care team instead being tasked to assist with post-ROSC care.

It is interesting that we were not able to demonstrate a statistically significant effect of defibrillation prior to arrival of EEAST, which we take to be a measure of the effectiveness of public access defibrillators corrected for the other factors that we considered. However, only a small number of patients (n=141) were defibrillated prior to EEAST arrival, introducing the chance of a type 2 error.

The Index of Multiple Deprivation was removed from the multivariable model as it was found to strongly covary with documented bystander-CPR. Univariable analysis found a strong association between receiving bystander-CPR and Index of Multiple Deprivation with patients more likely to receive bystander-CPR in less deprived postcodes (p=0.002). We cannot draw conclusions on whether this is a result of social, cultural or educational factors, or whether this is the result of geographic population sparsity, but our results do suggest a potentially high impact target area for policy efforts to improve public engagement with bystander-CPR.

For patients suffering from TCA, the multivariable analysis results were very different to NTCA. There was only weak evidence (p=0.057) that shorter response times were associated with increased survival at hospital arrival, and no evidence for increased survival at discharge. For survival to hospital discharge, an initial shockable rhythm was the only significant covariate.

Our approach, using regional data has the advantage of limiting variations in care provision and geographical influences. This approach may offer the possibility of exploiting regional differences in a comparative effectiveness research model to further understand determinants of outcome. Our work motivates a national, or indeed international, effort towards provider profiling regions to look for important covariates.

Limitations

Like any retrospective observational analysis, it is difficult to assign causality and our results are limited by any inherent errors in the data. However, it is reassuring that the missingness was low. We were unable to include in-hospital management variables in our model, these may have potentially confounded the survival to discharge outcome.

`Survival to discharge' from the receiving hospital may include patients who leave the receiving hospital not because they ultimately survive but because they are transferred to specialist centres for prognostication and perhaps subsequently go on to die. Unfortunately, we have no way of knowing this from our data. Furthermore, survival to hospital discharge data was obtained by follow-up and therefore may be missing not-at-random.

Conclusions

NTCA and TCA are clinically distinct entities with different predictors for outcome – future OHCA reports should aim to separate arrest aetiologies. Determinants of survival to hospital admission and discharge differ in a way that likely reflects the determinants of neurological injury. Bystander-CPR public engagement may be best focused in more deprived areas.

Contribution statement

The study was conceived by EB, AE, and AW. Data were supplied by TN. Data analyses were undertaken by AE and DS, with input from EB, AW, and TN. The manuscript was drafted by EB, DS, and AE, with critical review by TN and AW. All authors have reviewed and agreed the final manuscript.

Acknowledgements

We would like to acknowledge the assistance of the Audit & Research team at East of England Ambulance Service NHS Trust in compiling the original data, and the assistance of Mr. Andrew Downes, Dr. Tobias Gauss, and Dr. James Price (all East Anglian Air Ambulance) in the development of the AngliaOHCA project.

Competing interests

None declared by any author; EB is a serving member of UK Defence.

Funding

No external funding was obtained for this project.

KEY MESSAGES

What is already known on this subject

- The ambulance services in England attempt resuscitation on 30,000 out-ofhospital cardiac arrest (OHCA) patients per year.

- OHCA survival to discharge in England is lower than that of countries with comparable healthcare systems.

- A better understanding of outcome determinants may identify specific targets for improvement of OHCA survival.

What this study adds

- Pre-hospital determinants of non-traumatic and traumatic cardiac arrest outcome are different; these conditions should be analysed and managed independently.

- The determinants of survival identified are likely to reflect the magnitude of neurological injury incurred during resuscitation.

- Social deprivation and bystander CPR rates are strongly collinear –presenting a target for public engagement in OHCA.

REFERENCES

- 1. Resuscitation Council (UK). NHS England, British Heart Foundation. Consensus paper on out-of-hospital cardiac arrest in England. Resuscitation Council; 2014.
- Hawkes C, Booth S, Li C, Brace-McDonnell SJ, Whittington A, Mapstone J, et al. Epidemiology and outcomes from out-of-hospital cardiac arrests in England. Resuscitation 2016;110:133–40.
- Rajagopal S, Booth SJ, Brown TP, Ji C, Hawkes C, Siriwardena AN, et al. Data quality and 30-day survival for out-of-hospital cardiac arrest in the UK out-of-hospital cardiac arrest registry: a data linkage study. BMJ Open 2017;7(11):e017784. doi: 10.1136/bmjopen-2017-017784.
- 4. Leis CC, Hernández CC, Blanco MJG-O, Paterna PCR, Hernández R de E, Torres EC. Traumatic cardiac arrest: should advanced life support be initiated? J Trauma Acute Care Surg 2013;74(2):634–8.
- 5. Vopelius-Feldt von J, Powell J, Morris R, Benger J. Prehospital critical care for out-of-hospital cardiac arrest: An observational study examining survival and a stakeholder-focused cost analysis. BMC Emerg Med 2016;16(1):47.
- 6. Vopelius-Feldt von J, Brandling J, Benger J. Systematic review of the effectiveness of prehospital critical care following out-of-hospital cardiac arrest. Resuscitation 2017;114:40–6.
- 7. Barnard E, Yates D, Edwards A, Fragoso-Iñiguez M, Jenks T, Smith JE. Epidemiology and aetiology of traumatic cardiac arrest in England and Wales - A retrospective database analysis. Resuscitation 2017;110:90–4.
- 8. H Hayashi M, Shimizu W, Albert CM. The spectrum of epidemiology underlying sudden cardiac death. Circ Res 2015;116(12):1887–906.
- 9. Lockey DJ, Lyon RM, Davies GE. Development of a simple algorithm to guide the effective management of traumatic cardiac arrest. Resuscitation 2013;84(6):738–42.
- 10. Office for National Statistics. Revised population estimates for England and Wales: mid-2012 to mid-2016, 2018. (Accessed 01 August 2018, at https://www.ons.gov.uk/releases/revisedannualmidyearpopulationestimat es2012to2016).
- 11. Department of Health and Social Care. NHS Constitution for England: resources, 2015. (Accessed 01 August 2018, at

https://www.gov.uk/government/collections/nhs-constitution-for-england-resources).

- 12. Joint Royal Colleges Ambulance Liaison Committee, Association of Ambulance Chief Executives. UK Ambulance Services Clinical Practice Guidelines 2016. Bridgewater: Class Publishing; 2016.
- 13. Jacobs I, Nadkarni V, Bahr J, Berg RA, Billi JE, Bossaert L, et al. Cardiac arrest and cardiopulmonary resuscitation outcome reports: update and simplification of the Utstein templates for resuscitation registries: a statement for healthcare professionals from a task force of the International Liaison Committee on Resuscitation (American Heart Association, European Resuscitation Council, Australian Resuscitation Council, New Zealand Resuscitation Council, Heart and Stroke Foundation of Canada, InterAmerican Heart Foundation, Resuscitation Councils of Southern Africa). Resuscitation 2004;63(3):233-49.
- 14. Ministry of Housing, Communities & Local Government. English indices of deprivation 2015, 2015. (Accessed 01 August 2018, at https://www.gov.uk/government/statistics/english-indices-of-deprivation-2015).
- 15. Perkins GD, Jacobs IG, Nadkarni VM, Berg RA, Bhanji F, Biarent D, et al. Cardiac arrest and cardiopulmonary resuscitation outcome reports: update of the Utstein Resuscitation Registry Templates for Out-of-Hospital Cardiac Arrest: a statement for healthcare professionals from a task force of the International Liaison Committee on Resuscitation (American Heart Association, European Resuscitation Council, Australian and New Zealand Council on Resuscitation, Heart and Stroke Foundation of Canada, InterAmerican Heart Foundation, Resuscitation Council of Southern Africa, Resuscitation Council of Asia); and the American Heart Association Emergency Cardiovascular Care Committee and the Council on Cardiopulmonary, Critical Care, Perioperative and Resuscitation. Resuscitation 2015;96:328-40.
- 16. Buuren SV, Groothuis-Oudshoorn K. mice: Multivariate Imputation by Chained Equations in R. J Stat Soft 2011;45(3). doi: 10.18637/jss.v045.i03.
- 17. Barnard E, Yates D, Edwards A, Smith JE. Reply to Letter: Mortality in traumatic cardiac arrest. Resuscitation 2017;113:e23. doi: 10.1016/j.resuscitation.2017.01.004.
- Djarv T, Axelsson C, Herlitz J, Stromsoe A, Israelsson J, Claesson A. Traumatic cardiac arrest in Sweden 1990-2016 - a population-based national cohort study. Scand J Trauma Resusc Emerg Med 2018;26(1):30. doi: 10.1186/s13049-018-0500-7.

- 19. Smith JE, Rickard A, Wise D. Traumatic cardiac arrest. J R Soc Med 2015;108(1):11–6.
- Goto Y, Funada A, Goto Y. Relationship Between Emergency Medical Services Response Time and Bystander Intervention in Patients With Outof-Hospital Cardiac Arrest. J Am Heart Assoc 2018;7(9):e007568. doi: 10.1161/JAHA.117.007568.
- Benger JR, Kirby K, Black S, Brett SJ, Clout M, Lazaroo MJ, et al. Effect of a Strategy of a Supraglottic Airway Device vs Tracheal Intubation During Out-of-Hospital Cardiac Arrest on Functional Outcome: The AIRWAYS-2 Randomized Clinical Trial. JAMA 2018;320(8):779–91.
- 22. Dumas F, Bougouin W, Geri G, Lamhaut L, Bougle A, Daviaud F, et al. Is epinephrine during cardiac arrest associated with worse outcomes in resuscitated patients? J Am Coll Cardiol 2014;64(22):2360–7.
- 23. Perkins GD, Ji C, Deakin CD, Quinn T, Nolan JP, Scomparin C, et al. A Randomized Trial of Epinephrine in Out-of-Hospital Cardiac Arrest. N Engl J Med 2018;379(8):711-721.
- 24. Hamilton A, Steinmetz J, Wissenberg M, Torp-Pedersen C, Lippert FK, Hove L, et al. Association between prehospital physician involvement and survival after out-of-hospital cardiac arrest: A Danish nationwide observational study. Resuscitation 2016;108:95–101.

Age Group











