



University of Groningen

Long-term outcome of elderly out-of-hospital cardiac arrest survivors as compared with their younger counterparts and the general population

Hiemstra, Bart; Bergman, Remco; Absalom, Anthony R; van der Naalt, Joukje; van der Harst, Pim; de Vos, Ronald; Nieuwland, Wybe; Nijsten, Maarten W; van der Horst, Iwan C C

Published in: Therapeutic advances in cardiovascular disease

DOI: 10.1177/1753944718792420

IMPORTANT NOTE: You are advised to consult the publisher's version (publisher's PDF) if you wish to cite from it. Please check the document version below.

Document Version Publisher's PDF, also known as Version of record

Publication date: 2018

Link to publication in University of Groningen/UMCG research database

Citation for published version (APA): Hiemstra, B., Bergman, R., Absalom, A. R., van der Naalt, J., van der Harst, P., de Vos, R., ... van der Horst, I. C. C. (2018). Long-term outcome of elderly out-of-hospital cardiac arrest survivors as compared with their younger counterparts and the general population. *Therapeutic advances in cardiovascular disease*, *12*(12), 309-320. https://doi.org/10.1177/1753944718792420

Copyright

Other than for strictly personal use, it is not permitted to download or to forward/distribute the text or part of it without the consent of the author(s) and/or copyright holder(s), unless the work is under an open content license (like Creative Commons).

Take-down policy If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

Downloaded from the University of Groningen/UMCG research database (Pure): http://www.rug.nl/research/portal. For technical reasons the number of authors shown on this cover page is limited to 10 maximum.

Original Research

Ther Adv Cardiovasc Dis

1–12

DOI: 10.1177/ 1753944718792420



Article reuse guidelines: sagepub.com/journalspermissions

Correspondence to: Bart Hiemstra Department of Critical Care, University of Groningen, University Medical Center Groningen, Hanzeplein 1, PO Box 30.001, Groningen, 9700 RB, The Netherlands

b.hiemstra01@umcg.nl Remco Bergman Anthony R. Absalom Joukje van der Naalt Pim van der Harst Ronald de Vos Wybe Nieuwland Maarten W. Nijsten Iwan C. C. van der Horst University of Groningen, University Medical Center Groningen, Groningen, The Netherlands

Long-term outcome of elderly out-of-hospital cardiac arrest survivors as compared with their younger counterparts and the general population

Bart Hiemstra^(b), Remco Bergman, Anthony R. Absalom^(b), Joukje van der Naalt, Pim van der Harst, Ronald de Vos, Wybe Nieuwland, Maarten W. Nijsten and Iwan C. C. van der Horst

Abstract

Background: Over the past decade, prehospital and in-hospital treatment for out-of-hospital cardiac arrest (OHCA) has improved considerably. There are sparse data on the long-term outcome, especially in elderly patients. We studied whether elderly patients benefit to the same extent compared with younger patients and at long-term follow up as compared with the general population.

Methods: Between 2001 and 2010, data from all patients presented to our hospital after OHCA were recorded. Elderly patients (\geq 75 years) were compared with younger patients. Neurological outcome was classified as cerebral performance category (CPC) at hospital discharge and long-term survival was compared with younger patients and predicted survival rates of the general population.

Results: Of the 810 patients admitted after OHCA, a total of 551 patients (68%) achieved return of spontaneous circulation, including 125 (23%) elderly patients with a mean age of 81 \pm 5 years. In-hospital survival was lower in elderly patients compared with younger patients with rates of 33% *versus* 57% (p < 0.001). A CPC of 1 was present in 73% of the elderly patients *versus* 86% of the younger patients (p = 0.031). In 7.3% of the elderly patients, a CPC >2 was observed *versus* 2.5% of their younger counterparts (p = 0.103). Elderly patients had a median survival of 6.5 [95% confidence interval (CI) 2.0–7.9] years compared with 7.7 (95% CI 7.5–7.9) years of the general population (p = 0.019).

Conclusions: The survival rate after OHCA in elderly patients is approximately half that of younger patients. Elderly patients who survive to discharge frequently have favorable neurological outcomes and a long-term survival that approximates that of the general population.

Keywords: cardiopulmonary resuscitation, cerebral performance, elderly, long-term mortality, out-of-hospital cardiac arrest, prognosis

Received: 2 January 2018; revised manuscript accepted: 20 June 2018.

Introduction

Outcome after out-of-hospital cardiac arrest (OHCA) has markedly improved in the past years.^{1–3} While the overall mortality rate is still considerable, the patients who do survive tend to have a good neurological outcome.^{4–6} Many studies have focused on short-term outcome after OHCA,^{7–11}

and studies on the elderly have increased during the past years.^{12–18} Since older studies reported dismal outcomes in elderly patients after cardiopulmonary resuscitation (CPR),¹⁹ there still may be reluctance to institute maximal treatment in all patients for fear of only generating a greater number of survivors with poor functional outcomes.^{20,21}

Several recent studies show that with increasing age, short-term survival decreases but survivors display a good neurologic outcome.^{13–15} Long-term survival after OHCA seems to approach the general population,^{22–25} but little is known on the long-term survival of elderly patients who survive until hospital discharge.^{16–18} To what extent elderly OHCA patients benefit from early treatment as compared with younger patients and the general population was investigated in the current study.

Materials and methods

Study design and setting

The University Medical Center Groningen (UMCG) is the main tertiary referral hospital for the north-eastern part of the Netherlands, which provides 24/7 emergency care in a region with 750,000 inhabitants.²⁶ It operates in combination with multiple emergency medical services (EMS) in this area. A driver with a paramedical background and a nurse staff each ambulance, and both are trained and qualified to perform Advanced Cardiac Life Support (ACLS) according to the European Resuscitation Council guidelines, such as advanced airway interventions, administration of inotropic drugs and use of a defibrillator.²⁷ In case of resuscitation, a backup ambulance is always sent. The nurse staff always commences CPR but has the discretion to discontinue if the patient has a 'do not attempt CPR' decision or a nonshockable rhythm present for more than 20 minutes. Except for these patients, all other OHCAs including EMS-witnessed arrests are transported to our center where resuscitation is continued or postresuscitation care is given following ACLS guidelines.²⁷ When patients present with an ST-segment elevation myocardial infarction (STEMI) after OHCA we perform percutaneous coronary intervention (PCI) in patients with return of spontaneous circulation (ROSC), including elderly patients.²⁸ Patients requiring continuous mechanical ventilation were transferred to the intensive care unit (ICU) for postresuscitation care. Treatment was left at the discretion of the attending physicians. If brainstem reflexes (i.e. absent pupillary light response and corneal reflexes) and motor scores were absent, no further life-sustaining treatment was delivered. In all other cases, additional somatosensory-evoked-potential (SSEP) monitoring was done and in the case of absent bilateral negative 20 (N20) SSEPs, active treatment was also

withdrawn. Electroencephalography (EEG) analysis was performed when seizures or myoclonus was present. Seizures were treated with antiepileptic drugs; in the case of *myoclonus status epilepticus*, treatment was also stopped.^{29,30} In these patients, active therapies were discontinued, and treatment was altered to comfort care. The institutional review board [Medisch Ethische Toetsingscommissie (METc) of the UMCG] approved the study and waived the need for informed consent due to the observational nature of the study (METc 2011.374).

Patients

We studied all adult (\geq 18 years) patients who were transferred to our hospital after OHCA during the years 2001–2010. Data were collected retrospectively from the medical records of the Ambulance System. During the study period, mild therapeutic hypothermia (MTH) was initiated if patients had a Glasgow Coma Score < 8. These patients were cooled to 32–34°C for 24 h and thereafter, passively rewarmed to 36°C. In this cohort, we defined patients with an age of \geq 75 years as elderly and used the younger patients for comparison of outcomes in both categories.

Data on prehospital characteristics, medical history, electrocardiography (ECG), SSEP, EEG and laboratory values were gathered from the hospital information system. Neurological status was classified using the cerebral performance category (CPC) from the clinical patient notes made during outpatient follow-up assessments by cardiologists, rehabilitation specialists or, when indicated, neurologists. Survival status of patients discharged alive was obtained from the municipal personal records database on 20 December 2017, with a maximum follow up of 16.4 years.

Statistical analysis

Continuous data are presented as either means \pm standard deviation for normally distributed variables or medians and interquartile ranges for skewed data, or as group percentages with 95% confidence intervals (CI) for categorical variables. We determined outcome in subgroups dependent on initial rhythm [ventricular fibrillation (VF) versus no-VF] and implemented treatment (PCI versus no PCI, MTH versus no MTH). We recorded age, sex and known factors associated with outcome: initial rhythm and in-hospital treatment (PCI and MTH) and left ventricular function.

B Hiemstra, R Bergman et al.

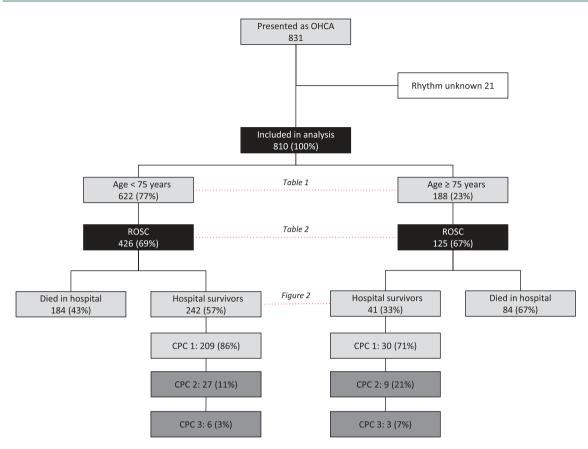


Figure 1. Flow diagram of study population.

CPC, cerebral performance category; OHCA, out-of-hospital cardiac arrest; ROSC, return of spontaneous circulation.

Statistical analysis was performed using the chisquare test for categorical and the Mann-Whitney U test or Student's t test for skewed continuous and normally distributed variables, respectively. Kaplan-Meier survival analysis was used to assess long-term survival and the logrank test was used to compare survival across subgroups. For each OHCA patient, the actual survival was compared with the predicted survival rates of the general population standardized for age, sex and calendar year. These predicted survival rates were obtained from the Dutch Census Bureau, which predicts general population survival annually.³¹ We identified independent predictors of in-hospital mortality, coronary angiography (CAG), PCI and MTH using multivariable logistic regression analyses. Univariable analyses were conducted on all known predictors for mortality, which were added without any transformation. A p < 0.25was used for inclusion in the multivariable model. Calibration of the multivariable models was checked with Hosmer-Lemeshow tests and by plotting observed proportions against the predicted risks of 10 equally sized groups. We used complete-case analyses because all variables in our multivariable models had fewer than 5% missing values.³² Statistical significance was defined as a two-sided *p* value of < 0.05. Statistical analyses were performed with Stata version 15.1 (StataCorp, 2017, College Station, Texas: StataCorp LLC).

Results

Prehospital characteristics of all admitted patients

From 2001 to 2010, a total of 810 patients were admitted after OHCA (Figure 1). Of these, 188 (23%) patients were \geq 75 years with a mean age of 81 \pm 5 years. Baseline and clinical characteristics of patients aged < 75 years and \geq 75 years are presented in Table 1. The location of arrest differed between both groups and elderly patients less frequently had an initial rhythm of VF. There was no difference between the two age groups in the fraction of patients who

	Total n = 810	< 75 years n = 622	≥ 75 years n = 188	<i>p</i> value
Age*	63 ± 15	57 ± 12	81 ± 5	<0.001
Male sex (%)	572 (71%)	448 (72%)	124 (66%)	0.110
Prehospital characteristics				
Location (%)				< 0.001
Home	374 (46%)	280 (45%)	94 (50%)	
Public place	190 (24%)	164 (26%)	26 (14%)	
Other	116 (14%)	91 (15%)	25 (13%)	
Ambulance	60 (7%)	47 (8%)	13 (7%)	
Unknown	70 (9%)	40 (6%)	30 (16%)	
Witnessed arrest (%)	642 (79%)	504 (81%)	138 (73%)	0.964
Bystander CPR (%)	460 (57%)	366 (59%)	94 (50%)	0.160
Initial rhythm (%)				< 0.001
VF	518 (64%)	421 (68%)	97 (52%)	
PEA	149 (18%)	97 (16%)	52 (28%)	
Asystole	116 (14%)	84 (14%)	32 (17%)	
Pulseless VT	27 (3%)	20 (3%)	7 (4%)	
ROSC before ED arrival (%)	473 (58%)	374 (60%)	99 (53%)	0.013
ROSC (%)	551 (68%)	426 (69%)	125 (67%)	0.606
Cardiovascular history and risk	factors			
History of CVD (%)	257 (32%)	169 (27%)	88 (48%)	< 0.001
Myocardial infarction (%)	148 (18%)	98 (16%)	50 (27%)	< 0.001
STEMI (%)	131 (16%)	86 (14%)	45 (24%)	< 0.001
PCI (%)	61 (8%)	46 (7%)	15 (8%)	0.680
CABG (%)	57 (7%)	31 (5%)	26 (14%)	< 0.001
Angina (%)	79 (10%)	49 (8%)	30 (16%)	<0.001
Cerebrovascular accident (%)	51 (6%)	33 (5%)	18 (10%)	0.024
History of VF (%)	16 (2%)	11 (2%)	5 (3%)	0.400
Malignancy (%)	35 (4%)	16 (4%)	19 (10%)	< 0.001
Hypertension (%)	196 (24%)	141 (23%)	55 (29%)	0.031
Hypercholesterolemia (%)	78 (10%)	65 (11%)	13 (7%)	0.180
Diabetes (%)	108 (13%)	72 (12%)	36 (19%)	0.004

Table 1 .(Continued)

· ·				
	Total n = 810	< 75 years n = 622	≥ 75 years n = 188	<i>p</i> value
Medication history				
Beta blocker (%)	185 (23%)	124 (20%)	61 (32%)	< 0.001
ACE inhibitor (%)	125 (15%)	80 (13%)	45 (24%)	< 0.001
AT-II receptor antagonist (%)	56 (7%)	39 (6%)	17 (9%)	0.043
Diuretics (%)	142 (18%)	87 (14%)	55 (29%)	< 0.001
Antiplatelet therapy (%)	126 (16%)	83 (13%)	43 (23%)	< 0.001
Coumarin derivatives (%)	110 (14%)	72 (12%)	38 (20%)	< 0.001
Statins (%)	150 (19%)	107 (17%)	43 (23%)	0.003
Antiarrhythmics (%)	33 (4%)	18 (3%)	15 (8%)	< 0.001
Oral antidiabetic therapy (%)	57 (7%)	37 (6%)	20 (11%)	0.003
Insulin (%)	35 (4%)	30 (5%)	5 (3%)	0.390

*Mean \pm standard deviation.

ACE, angiotensin-converting enzyme; AT, angiotensin; CABG, coronary artery bypass grafting; CPR, cardiopulmonary resuscitation; CVD, cardiovascular disease; ED, emergency department; PCI, percutaneous coronary intervention; PEA, pulseless electrical activity; ROSC, return of spontaneous circulation; STEMI, ST-segment elevation myocardial infarction; VF, ventricular fibrillation; VT, ventricular tachycardia.

achieved ROSC, although upon arrival at the emergency department, younger patients more often had achieved ROSC. Elderly patients also had an increased incidence of prior cardiovascular disease and risk factors. More specifically, elderly patients were more frequently known with a previous myocardial infarction, cerebrovascular accident, hypertension, diabetes and concurrent malignancy.

In-hospital characteristics of patients who achieved return of spontaneous circulation

Table 2 summarizes the clinical characteristics, angiography data and outcome of the patients who achieved ROSC. After ROSC, the initial Glasgow Coma Scores were equally low in the elderly and younger cohort, with median scores of 3 (3–5) and 3 (3–6), p = 0.358, respectively. Elderly patients underwent fewer primary CAGs, were less often diagnosed with a STEMI, had smaller creatine kinase-myocardial band (CK-MB) and troponin T levels and presented with higher N-terminal prohormone of brain natriuretic peptide (NT-proBNP) levels (p < 0.001 for all). There were no

differences in glucose, lactate, pH, and base excess between these groups. Among the elderly, CAG was associated with an initial rhythm of VF, an ST-segment elevation on the ECG and higher arterial pH levels at admission (Supplementary Table 1).

Treatment

Only 31 patients (25%) from the elderly group underwent PCI versus 184 patients (43%) in the younger group (p < 0.001). In both younger and elderly patients, independent predictors of PCI were the presence of ST-segment elevation on the ECG and higher serum CK-MB levels [Supplementary Table 2(a), 2(b)]. MTH was similarly used in the elderly (32%, 95% CI 24-41%) and younger (38%, 95% CI 33-43%) patients (p = 0.277). In the elderly, an initial rhythm of VF and higher serum CK-MB levels were independently associated with institution of MTH (Supplementary Table 3). There were no statistically significant differences in the use of an intra-aortic balloon pump and mechanical ventilation between both groups (Table 2).

Table 2. Clinical characteristics, treatment and outcome after return of spontaneous circulation.

	Total n = 551 (68%)	< 75 years n = 426 (68%)	≥ 75 years n = 125 (66%)	<i>p</i> value
Clinical characteristics				
Glasgow Coma Score at admission*	4 (3, 13)	4 (3, 14)	3 (3, 9)	0.450
STEMI (%)	183 (33%)	157 (39%)	26 (22%)	< 0.001
NSTEMI (%)	72 (13%)	51 (12%)	21 (17%)	0.160
Wall motion abnormalities (%)	82 (15%)	73 (32%)	9 (21%)	0.150
LVEF (%)				0.830
Poor-moderate	62 (24%)	50 (31%)	12 (33%)	
Reasonable-normal	133 (65%)	109 (69%)	24 (67%)	
Laboratory values				
Glucose*	12.7 (9.9, 16.9)	12.3 (9.6, 16.6)	13.2 (10.6, 17.1)	0.060
Lactate*	6.2 (2.95, 9.75)	6.3 (2.9, 9.9)	5.4 (3.1, 8.7)	0.470
pH*	7.25 (7.10, 7.33)	7.25 (7.1, 7.32)	7.27 (7.11, 7.34)	0.510
Base excess*	-8.1 (-14.5, -3.9)	-8.1 (-14.7, -3.8)	-7.9 (-12.9, -3.95)	0.640
CK max*	1943 (436, 5585)	2390 (607, 6376)	628 (286, 3104)	<0.001
CK-MB max*	169 (64, 554)	214 (77, 587)	110 (44, 241)	<0.001
Troponin T max*	2.22 (0.29, 11.33)	2.53 (0.39, 12.30)	0.99 (0.20, 6.30)	0.016
NT-pro BNP*	313 (75, 1607)	229 (69, 898)	1731 (312, 3619)	<0.001
Treatment				
Coronary angiography (%)	295 (54%)	255 (60%)	40 (32%)	<0.001
Primary	217 (39%)	185 (43%)	32 (26%)	<0.001
Secondary	78 (14%)	70 (16%)	8 (6%)	0.005
Door to balloon time (min)*	88 (56, 128)	88 (55, 125)	83 (61, 137)	0.270
Culprit lesion known (%)	295 (54%)	255 (60%)	40 (32%)	0.089
Right coronary artery	67 (12%)	58 (14%)	9 (7%)	
Left anterior descending	111 (20%)	97 (23%)	14 (11%)	
Circumflex	44 (8%)	37 (9%)	7 (6%)	
Left main	12 (2%)	7 (2%)	5 (4%)	
Graft	4 (1%)	4 (1%)	0 (0%)	
Unknown or multiple	32 (6%)	28 (7%)	4 (3%)	
No culprit	25 (5%)	24 (6%)	1 (1%)	

Table 2 .(Continued)

	Total n = 551 (68%)	< 75 years n = 426 (68%)	≥ 75 years n = 125 (66%)	<i>p</i> value
PCI (%)	215 (39%)	184 (43%)	31 (25%)	<0.001
Primary	191 (35%)	164 (39%)	27 (22%)	< 0.001
Secondary	24 (4%)	20 (5%)	4 (2%)	0.470
Intra-aortic balloon pump (%)	73 (13%)	60 (14%)	13 (10%)	0.290
Mild therapeutic hypothermia (%)	201 (37%)	161 (38%)	40 (32%)	0.280
Mechanical ventilation (%)	145 (26%)	118 (28%)	27 (22%)	0.200
Outcome				
Length of hospital stay (days)*	6 (3, 18)	7 (3, 20)	4 (2,12.5)	< 0.001
In-hospital death (%)	268 (49%)	184 (43%)	84 (67%)	< 0.001
Neurological bad outcome	184 (33%)	132 (31%)	52 (42%)	
Cardiogenic shock	43 (8%)	25 (6%)	18 (14%)	
Multiorgan failure	17 (3%)	12 (3%)	5 (4%)	
Other	24 (4%)	15 (4%)	9 (7%)	
Hospital survival (%)	283 (51%)	242 (57%)	41 (33%)	< 0.001
CPC at discharge (%)				0.041
1	239 (84%)	209 (86%)	30 (71%)	
2	36 (13%)	27 (11%)	9 (21%)	
3	9 (3%)	6 (3%)	3 (7%)	
Survival at end of follow up (%)	185 (33%)	178 (42%)	7 (6%)	< 0.001

*Median (interquartile range).

CK-MB, creatine kinase-myocardial band; CPC, cerebral performance category; LVEF, left ventricular ejection fraction; (N)STEMI, (non)-ST-segment elevation myocardial infarction; NT-pro BNP, N-terminal prohormone of brain natriuretic peptide; PCI, percutaneous coronary intervention; ROSC, return of spontaneous circulation.

Outcome

Survival to hospital discharge was lower in elderly patients with only 33% (95% CI 25–42%) surviving compared with 57% (95% CI 52–62%) in the younger group (p < 0.001; Table 2). Being elderly was independently associated with in-hospital mortality when adjusted for known prognostic factors [i.e. sex, initial rhythm, bystander CPR, witnessed arrest, arterial glucose and pH levels, and prior CVD (Supplementary Table 4)]. Of the 41 elderly patients who survived to hospital discharge, 24 (59%, 95% CI 42–74%) had VF *versus* 17 (41%, 95% CI 26–58%) patients with non-VF (p = 0.600). In younger patients, survival in the

VF group was 207 (86%, 95% CI 81–90%) versus 35 (15%, 95% CI 10–20%) in the non-VF patients (p < 0.001). Only Glasgow Coma Score at admission was independently associated with mortality in the elderly (Supplementary Table 5). Length of hospital stay was shorter in the elderly patients (Table 2), but length of hospital stay was similar for hospital survivors in both groups.

After hospital discharge, survival in elderly patients was worse in comparison with younger patients (Figure 2). The 1-year survival in the overall OHCA group after hospital discharge was 94% (95% CI 91–97%) and 5-year survival was

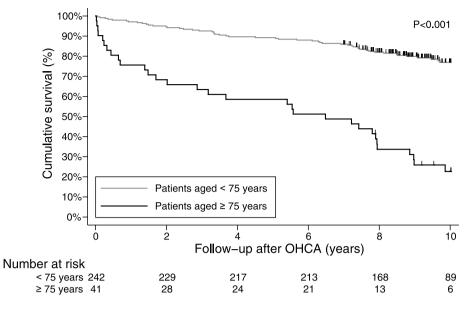


Figure 2. Survival curves of younger (< 75 years) and elderly (≥ 75 years) patients. OHCA, out-of-hospital cardiac arrest.

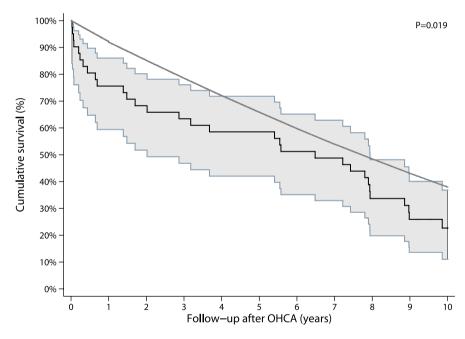


Figure 3. Survival curve elderly versus matched control population with 95% confidence intervals.

still 85% (95% CI 80–89%). In contrast, in the patients aged \geq 75 years, 1-year survival was only 76% (95% CI 60–88%) and 5-year survival dropped to 59% (95% CI 42–74%). However, when standardized for sex, age and calendar year, long-term survival approached that of the general population (Figure 3). The median survival of the elderly patients was 6.5 (95% CI 2.0–7.9) years compared with 7.7 (95% CI 7.5–7.9) years for a matched general population (p = 0.019).

Neurological outcome at hospital discharge

The younger cohort had a better neurological outcome, as scored by the CPC (Table 2). In the elderly, 73% (95% CI 57–86%) had a CPC of 1 compared with 86% (95% CI 81–90%) in the younger patients (p = 0.031). The incidence of poor outcome (CPC > 2) was 7.3% (95% CI 1.5–20%) in the elderly *versus* 2.5% (95% CI 0.9–5.3%) in the younger cohort (p = 0.103).

Cause of death

Causes of in-hospital death did not vary substantially between age groups (Table 2). The major cause of death was neurological damage that was deemed not compatible with survival. Other important causes of death were secondary circulatory failure and multiorgan failure (Table 2). Of the 35 patients with a known malignancy, only one survived until end of follow up (p = 0.003).

Discussion

From the long-term follow up of a cohort of 810 patients after OHCA, two main observations emerged. First, despite similar percentages of patients regaining spontaneous circulation, inhospital mortality was considerably higher in elderly compared with younger patients. Second, we add to the current literature, that elderly patients who were discharged alive had a median survival of 6.5 years compared with 7.7 years in matched controls in the general population. Moreover, in these elderly patients, the neurological outcome was good in a high percentage of patients, with less than 10% with a poor neurological outcome.

Our findings confirm that elderly patients have worse outcomes than younger patients: Funada and colleagues studied 334,730 elderly (≥ 75 vears) and 23,520 very elderly (\geq 95 years) patients,12,33 and in 2017, five more studies described worse outcomes of elderly patients after 1 month: Aissaoui and colleagues ($n = 464, \ge 65$ years),³⁴ Andrew and colleagues (n = 14,678, ≥ 65 years),¹⁸ Segal and colleagues (n = 18,249, > 65 years),¹⁴ Sulzgruber and colleagues (n =1315, ≥ 65 years),¹⁵ and Wiel and colleagues (n = 4347, > 65 years).¹³ In a study with 34,291 patients Wong and colleagues showed in 2014 that over time, the survival of the elderly at 1 year has increased.³⁶ The favorable outcome trend for functional outcome reflected by good CPC scores at 1 month was recently shown in large Japanese cohort studies.^{12,37,38} Beesems and colleagues showed worse survival and equally favorable functional outcome in survivors compared with our study.³⁹ In 1332 patients aged \geq 70 years, 1-year survival in patients discharged alive was 88% and neurologic outcome, as defined by a CPC of 1 or 2, was good in 90% of patients. Winther-Jensen and colleagues showed that 70% of the very elderly patients (i.e. ≥ 80 years) had CPC scores of 1 or 2 at 30-day follow up, which is comparable with the observed 70% elderly with

In our study, several factors were associated with both mortality and unfavorable neurological outcome. In line with observations of others, the elderly have a higher cardiovascular risk profile and more comorbidities.^{39–41} We also observed a higher incidence of an initial rhythm other than VF, which is associated with a poor outcome, in the elderly patients.⁴² Similar to Sulzgruber and colleagues, the initial rhythm and other known prognostic factors such as witnessed arrest, bystander CPR, arterial glucose, base excess and lactate levels were not associated with outcome in the elderly.¹⁵ More data are needed to identify early prognostic variables, as other cohort studies did find associations between these variables and outcome.^{6,42–44}

Compared with other studies, the incidence of ROSC in the elderly in our population was high and almost equal to younger patients.47,48 A selection bias could be present as referral to our hospital may have been considered futile in patients with very poor early prognostic variables or a nonshockable rhythm present for more than 20 minutes. Like in the study of Winther-Jensen and colleagues, CAG and PCI were employed less frequently in the elderly patients.¹⁷ Our data show that in elderly patients, OHCA less frequently has an acute cardiac origin. Elderly patients less often suffer from a myocardial infarction, initial rhythm of VF and have lower cardiac biomarkers. Also, the higher incidence of previous myocardial infarction in the elderly might underlie secondary rhythm disturbances that cause OHCA. The different underlying pathology makes them less suitable for treatment such as PCI.6,49,50 Another reason could be that elderly patients tend to receive a less aggressive diagnostic work up compared with younger patients.13,51-53 MTH was employed equally in both groups in our cohort.

Treatment decisions can be particularly difficult for clinicians caring for elderly patients. CPR for older patients is a controversial issue.^{16,54,55} Initiating CPR in the elderly is a decision that is affected by sometimes conflicting opinions for what is best for the patient, and these opinions may vary from country to country. It involves not only medical, but also ethical decisions, especially when there are important associated comorbidities that directly affect the patient's prognosis.²¹ As shown in our study, the incidence of malignancies is significantly higher in the elderly. Since long-term survival of elderly hospital survivors is reasonable with a median survival of 6.5 years, our data thus do not support the withholding of resuscitation based on age only.¹⁶

Limitations

This study has several limitations. First, it was performed retrospectively on data of patients admitted to our hospital. Detailed data are lacking on the person who witnessed the arrest (EMS or bystander) and time frames of initial treatment, such as time to CPR, ACLS and ROSC for instance. Second, not all treatment was standardized; various potential sources of unidentified bias such as confounding by indication in the care of older and younger patients may be present. However, highlighting this potential bias is also an important finding in this study. Finally, these results only relate to patients presenting to the emergency unit of a single center. This limits the external validity of our findings. Some patients who were resuscitated during the study period in our region may not have been referred to our hospital, since PCI was not deemed necessary for some patients. This may explain differences in our patient population in comparison with other studies.^{10,56} As such, the outcome data we present do not pertain to the total population of patients suffering OHCA in our region.

Conclusion

Despite similar rates of ROSC after OHCA in elderly and younger patients, hospital survival for elderly patients was approximately half that of younger patients. However, the majority of elderly patients who survived initial treatment had a CPC score of 1 and a lifespan that approximates that of the general population.

Acknowledgements

Bart Hiemstra and Remco Bergman contributed equally to this work.

Funding

This research received no specific grant from any funding agency in the public, commercial, or notfor-profit sectors.

Conflict of interest statement

The authors declare that there is no conflict of interest.

ORCID iDs

Bart Hiemstra (D) https://orcid.org/0000-0001-6547-2138 Anthony R Absalom (D) https://orcid.org/0000-0001-7563-9157

References

- 1. Nolan JP. Optimizing outcome after cardiac arrest. *Curr Opin Crit Care* 2011; 17: 520–526.
- Savastano S, Klersy C, Raimondi M, et al. Positive trend in survival to hospital discharge after out-of-hospital cardiac arrest: a quantitative review of the literature. J Cardiovasc Med (Hagerstown) 2014; 15: 609–615.
- 3. Arrich J, Holzer M, Havel C, *et al.* Hypothermia for neuroprotection in adults after cardiopulmonary resuscitation. *Cochrane Database Syst Rev* 2016; 2: CD004128.
- 4. Bro-Jeppesen J, Kjaergaard J, Horsted TI, *et al.* The impact of therapeutic hypothermia on neurological function and quality of life after cardiac arrest. *Resuscitation* 2009; 80: 171–176.
- Kim WY, Giberson TA, Uber A, *et al.* Neurologic outcome in comatose patients resuscitated from out-of-hospital cardiac arrest with prolonged downtime and treated with therapeutic hypothermia. *Resuscitation* 2014; 85: 1042–1046.
- Bergman R, Hiemstra B, Nieuwland W, et al. Long-term outcome of patients after out-ofhospital cardiac arrest in relation to treatment: a single-centre study. Eur Heart J Acute Cardiovasc Care 2016; 5: 328–338.
- Hosmane VR, Mustafa NG, Reddy VK, et al. Survival and neurologic recovery in patients with ST-segment elevation myocardial infarction resuscitated from cardiac arrest. J Am Coll Cardiol 2009; 53: 409–415.
- 8. Lang ES. ACP Journal Club. Review: therapeutic hypothermia improves neurologic outcome and survival to discharge after cardiac arrest. *Ann Intern Med* 2010; 152: 22.
- Hinchey PR, Myers JB, Lewis R, et al. Improved out-of-hospital cardiac arrest survival after the sequential implementation of 2005 AHA guidelines for compressions, ventilations, and induced hypothermia: the Wake County experience. Ann Emerg Med 2010; 56: 348–357.
- 10. Berdowski J, Berg RA, Tijssen JG, *et al.* Global incidences of out-of-hospital cardiac arrest and survival rates: systematic review of 67 prospective studies. *Resuscitation* 2010; 81: 1479–1487.

- Van der Wal G, Brinkman S, Bisschops LL, et al. Influence of mild therapeutic hypothermia after cardiac arrest on hospital mortality. *Crit Care Med* 2011; 39: 84–88.
- Funada A, Goto Y, Maeda T, *et al.* Improved survival with favorable neurological outcome in elderly individuals with out-of-hospital cardiac arrest in Japan – a nationwide observational cohort study. *Circ* 7 2016; 80(5): 1153–1162.
- Wiel E, Di Pompeo C, Segal N, et al. Age discrimination in out-of-hospital cardiac arrest care: a case-control study. Eur J Cardiovasc Nurs. Epub ahead of print 1 December 2017. DOI: 10.1177/1474515117746329.
- 14. Segal N, di Pompeo C, Escutnaire J, et al. Evolution of survival in cardiac arrest with age in elderly patients: is resuscitation a dead end? *f Emerg Med* 2017; 54(3): 295–301.
- Sulzgruber P, Sterz F, Poppe M, et al. Agespecific prognostication after out-of-hospital cardiac arrest - the ethical dilemma between 'lifesustaining treatment' and 'the right to die' in the elderly. Eur Heart J Acute Cardiovasc Care 2017; 6(2): 112–120.
- Van de Glind EM, van Munster BC, van de Wetering FT, *et al.* Pre-arrest predictors of survival after resuscitation from out-of-hospital cardiac arrest in the elderly a systematic review. *BMC Geriatr* 2013; 13: 68.
- Winther-Jensen M, Pellis T, Kuiper M, et al. Mortality and neurological outcome in the elderly after target temperature management for out-ofhospital cardiac arrest. *Resuscitation* 2015; 91: 92–98.
- Andrew E, Mercier E, Nehme Z, *et al.* Long-term functional recovery and health-related quality of life of elderly out-of-hospital cardiac arrest survivors. *Resuscitation* 2018; 126: 118–124.
- Applebaum GE, King JE and Finucane TE. The outcome of CPR initiated in nursing homes. J Am Geriatr Soc 1990; 38: 197–200.
- 20. Nagappan R and Parkin G. Geriatric critical care. *Crit Care Clin* 2003; 19: 253–270.
- 21. Edin MG. Cardiopulmonary resuscitation in the frail elderly: clinical, ethical and halakhic issues. *Isr Med Assoc J* 2007; 9: 177–179.
- Andrew E, Nehme Z, Wolfe R, et al. Long-term survival following out-of-hospital cardiac arrest. *Heart* 2017; 103(14): 1104–1110.
- Smith K, Andrew E, Lijovic M, et al. Quality of life and functional outcomes 12 months after out-of-hospital cardiac arrest. *Circulation* 2015; 131(2): 174–181.

- Kragholm K, Wissenberg M, Mortensen RN, et al. Bystander efforts and 1-year outcomes in out-of-hospital cardiac arrest. N Engl J Med 2017; 376: 1737–1747.
- 25. Nichol G, Guffey D, Stiell IG, *et al.* Postdischarge outcomes after resuscitation from out-of-hospital cardiac arrest: a ROC PRIMED substudy. *Resuscitation* 2015; 93: 74–81.
- 26. Mahmoud KD, Gu YL, Nijsten MW, et al. Interhospital transfer due to failed prehospital diagnosis for primary percutaneous coronary intervention: an observational study on incidence, predictors, and clinical impact. Eur Heart J Acute Cardiovasc Care 2013; 2: 166–175.
- Soar J, Nolan JP, Bottiger BW, et al. European Resuscitation Council guidelines for resuscitation 2015: section 3. Adult advanced life support. *Resuscitation* 2015; 95: 100–147.
- 28. Peels HO, Jessurun GA, Van der Horst IC, et al. Outcome in transferred and nontransferred patients after primary percutaneous coronary intervention for ischaemic out-of-hospital cardiac arrest. *Catheter Cardiovasc Interv* 2008; 71(2): 147–151.
- Wijdicks EF, Hijdra A, Young GB, et al. Practice parameter: prediction of outcome in comatose survivors after cardiopulmonary resuscitation (an evidence-based review): report of the Quality Standards Subcommittee of the American Academy of Neurology. *Neurology* 2006; 67: 203–210.
- Bouwes A, Binnekade JM, Kuiper MA, et al. Prognosis of coma after therapeutic hypothermia: a prospective cohort study. *Ann Neurol* 2012; 71: 206–212.
- Central Bureau for Statistics. Statistics Netherlands: life expectancy; gender and age from 1950 on (per year), http://statline.cbs.nl/StatWeb/ publication/?PA=37360ned (2015, accessed 13 September 2017).
- 32. Jakobsen JC, Gluud C, Wetterslev J, et al. When and how should multiple imputation be used for handling missing data in randomised clinical trials a practical guide with flowcharts. BMC Med Res Methodol 2017; 17(1): 162.
- 33. Funada A, Goto Y, Maeda T, et al. Prehospital predictors of neurological outcomes in out-ofhospital cardiac arrest patients aged 95 years and older: a nationwide population-based observational study. J Cardiol 2017; 69(1): 340–344.
- 34. Aissaoui N, Bougouin W, Dumas F, et al. Postresuscitation care in elderly patients: insights from the sudden death expertise center registry. *Eur Heart J* 2017; 38(1): ehx504.P3022.

- 35. Wong MK, Morrison LJ, Qiu F, *et al.* Trends in short- and long-term survival among out-ofhospital cardiac arrest patients alive at hospital arrival. *Circulation* 2014; 130: 1883–1890.
- SOS-KANTO 2012 Study Group. Changes in treatments and outcomes among elderly patients with out-of-hospital cardiac arrest between 2002 and 2012: a post hoc analysis of the SOS-KANTO 2002 and 2012. *Resuscitation* 2015; 97: 76–82.
- Kitamura T, Iwami T, Kawamura T, et al. Nationwide improvements in survival from outof-hospital cardiac arrest in Japan. *Circulation* 2012; 126(24): 2834–2843.
- Beesems SG, Blom MT, Van der Pas MH, et al. Comorbidity and favorable neurologic outcome after out-of-hospital cardiac arrest in patients of 70 years and older. *Resuscitation* 2015; 94: 33–39.
- Bagnall AJ, Goodman SG, Fox KA, *et al.* Influence of age on use of cardiac catheterization and associated outcomes in patients with non-ST-elevation acute coronary syndromes. *Am J Cardiol* 2009; 103: 1530–1536.
- Mosier J, Itty A, Sanders A, et al. Cardiocerebral resuscitation is associated with improved survival and neurologic outcome from out-of-hospital cardiac arrest in elders. *Acad Emerg Med* 2010; 17: 269–275.
- Wissenberg M, Lippert FK, Folke F, et al. Association of national initiatives to improve cardiac arrest management with rates of bystander intervention and patient survival after out-of-hospital cardiac arrest. *JAMA* 2013; 310: 1377–1384.
- 42. McGill JW and Ruiz E. Central venous pH as a predictor of arterial pH in prolonged cardiac arrest. *Ann Emerg Med* 1984; 13: 684–687.
- 43. Mullner M, Sterz F, Domanovits H, *et al.* The association between blood lactate concentration on admission, duration of cardiac arrest, and functional neurological recovery in patients resuscitated from ventricular fibrillation. *Intensive Care Med* 1997; 23: 1138–1143.

44. Takasu A, Sakamoto T and Okada Y. Arterial

base excess after CPR: the relationship to CPR

Visit SAGE journals online http://tac.sagepub.com

SAGE journals

duration and the characteristics related to outcome. *Resuscitation* 2007; 73: 394–399.

- 45. Fukuda T, Ohashi-Fukuda N, Matsubara T, et al. Trends in outcomes for out-of-hospital cardiac arrest by age in Japan: an observational study. *Medicine (Baltimore)* 2015; 94: e2049.
- 46. Andersen LW, Bivens MJ, Giberson T, *et al.* The relationship between age and outcome in outof-hospital cardiac arrest patients. *Resuscitation* 2015; 94: 49–54.
- 47. Hollenbeck RD, McPherson JA, Mooney MR, et al. Early cardiac catheterization is associated with improved survival in comatose survivors of cardiac arrest without STEMI. *Resuscitation* 2014; 85: 88–95.
- Camuglia AC, Randhawa VK, Lavi S, et al. Cardiac catheterization is associated with superior outcomes for survivors of out of hospital cardiac arrest: review and meta-analysis. *Resuscitation* 2014; 85: 1533–1540.
- 49. Scott IA and Guyatt GH. Cautionary tales in the interpretation of clinical studies involving older persons. *Arch Intern Med* 2010; 170(7): 587–595.
- 50. Lindner TW, Langorgen J, Sunde K, *et al.* Factors predicting the use of therapeutic hypothermia and survival in unconscious out-ofhospital cardiac arrest patients admitted to the ICU. *Crit Care* 2013; 17(4): R147.
- 51. Hirlekar G, Karlsson T, Aune S, *et al.* Survival and neurological outcome in the elderly after in-hospital cardiac arrest. *Resuscitation* 2017; 118: 101–106.
- 52. Gordon M. Ethical challenges in end-of-life therapies in the elderly. *Drugs Aging* 2002; 19: 321–329.
- Deasy C, Bray JE, Smith K, et al. Out-ofhospital cardiac arrests in the older age groups in Melbourne, Australia. *Resuscitation* 2011; 82: 398–403.
- 54. Van Alem AP, Chapman FW, Lank P, *et al.* A prospective, randomised and blinded comparison of first shock success of monophasic and biphasic waveforms in out-of-hospital cardiac arrest. *Resuscitation* 2003; 58: 17–24.