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Intensive care-treated cardiac arrest: A retrospective study on the impact of extended age on mortality, neurological outcome, received treatments and healthcare-associated costs

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Bakgrund: Hjärtattacker är en av de ledande dödsorsakerna internationellt. Detta i kombination med en åldrande befolkning leder till ett ökat behov för forskning kring ålderns effekt på utfall och vård i hjärtattacker. I denna studie delade vi in vårt patientmaterial i två grupper: patienter 75 år och äldre samt patienter yngre än 75 år. Efter detta jämförde vi grupperna i kategorierna vårdintensitet, neurologiskt utfall efter 12 månader, dödlighet samt hälsovårdskostnader.

Metoder: Vårt material inkluderade alla vuxna patienter som vårdades på Mejlans sjukhus intensivvårdsavdelning på grund av hjärtattack åren 2005-2013. Vårdintensitet bedömdes med hjälp av de totala och de dagliga median TISS-76-poängen (Therapeutic intervention scoring system-76). Kostnadseffektiviteten räknades ut genom att dela de totala hälsovårdskostnaderna av alla patienter med mängden patienter som hade ett gott neurologiskt utfall 12 månader efter hjärtattacken. Vi definierade ett gott neurologiskt utfall med hjälp av CPC-skalan (cerebral performance category), där CPC 1–2 bedömdes som ett gott utfall. För att illustrera skillnader i fördelningen av kostnader användes stapeldiagram. Efter detta bedömde vi också med hjälp av logistisk regression olika faktors samband med ett dåligt neurologiskt utfall och hög mortalitet. Kaplan-Meier kurvor användes för att illustrera skillnaden i mortalitet mellan åldersgrupperna.

Resultat och slutsats: Totalt 1285 patienter inkluderades i studien, 212 (16%) av dessa var 75 år eller äldre. Det var en klar skillnad i behandlingsintensitet mellan åldersgrupperna, vi konstaterade färre vårdinterventioner hos de äldre både i medeltal totalt och dagligen. Här funderade vi i artikeln om detta kunde vara på grund av en aktiv daglig bedömning av situationen och att man snabbare bedömer att äldre inte tjänar på att vårdas aggressivt. Trots detta var den effektiva kostnaden högre för de äldre, vilket implicerar att vården för äldre ännu kan effektiviseras och att vi behöver fortsätta utveckla bra verktyg för bedömning



av vilka åldringar som tjänar på en aktiv och intensiv vård. Det neurologiska utfallet var också markant sämre för den äldre åldersgruppen, dock mycket på grund av den höjda mortaliteten. I våra multivariabel-modeller var faktorer som påverkade det neurologiska utfallet samt mortaliteten: ålder, gerasteni, hjärtrytmen, komorbiditeten och tid till spontan blodcirkulation vid återupplivningen. Därmed kan man konstatera att ålder föga överraskande påverkar utfallet efter en hjärtattack, men att det samtidigt också är viktigt att ta i beaktande övriga faktorer som påverkar utfallet.

(376 ord)

1 **Intensive care-treated cardiac arrest: A retrospective study on the impact**
2 **of extended age on mortality, neurological outcome, received treatments**
3 **and healthcare-associated costs**

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12 Abstract

13 **Background:** Cardiac arrest (CA) is a leading cause of death worldwide. As population ages, the
14 need for research focusing on CA in elderly increases. This study investigated treatment intensity,
15 12-month neurological outcome, mortality and healthcare-associated costs for patients aged over 75
16 years treated for CA in an intensive care unit (ICU) of a tertiary hospital.

17 **Methods:** This single-centre retrospective study included adult CA patients treated in a Finnish
18 tertiary hospital's ICU between 2005 and 2013. We stratified the study population into two age
19 groups: <75 and ≥ 75 years. We compared interventions defined by the median daily therapeutic
20 intervention scoring system (TISS-76) between the age groups to find differences in treatment
21 intensity. We calculated cost-effectiveness by dividing the total one-year healthcare-associated costs
22 of all patients by the number of survivors with a favourable neurological outcome. Favourable
23 outcome was defined as a cerebral performance category (CPC) of 1–2 at 12 months after cardiac
24 arrest. Logistic regression analysis was used to identify independent association between age group,
25 mortality and neurological outcome.

26 **Results:** This study included a total of 1,285 patients, of which 212 (16%) were ≥ 75 years of age.
27 Treatment intensity was lower for the elderly compared to the younger group, with median TISS
28 scores of 116 and 147, respectively ($p < 0.001$). The effective cost in euros for patients with a good
29 one-year neurological outcome was €168,000 for the elderly and €120,000 for the younger group.
30 At 12 months after CA 24% of the patients in the elderly group and 47% of the patients in the younger
31 group had a CPC of 1-2 ($p < 0.001$). Age was an independent predictor of mortality (multivariate OR
32 = 3.36, 95% CI:2.21-5.11, $p < 0.001$) and neurological outcome (multivariate OR = 3.27, 95% CI:
33 2.12-5.03, $p < 0.001$).

34 **Conclusions:** The elderly ICU-treated CA patients in this study had worse neurological outcomes,
35 higher mortality and lower cost-effectiveness than younger patients. Further efforts are needed to
36 recognize the tools for assessing which elderly patients benefit from a more aggressive treatment
37 approach in order to improve the cost-effectiveness of post-CA management.

38
39 **Keywords:** Elderly, Cardiac arrest, intensive care unit, critical care, cardiopulmonary resuscitation,
40 OHCA, IHCA, ICUCA

41

42 **Background**

43 CA is one of the leading causes of death in the developed world [1], with over three million patients
44 affected each year worldwide [2]. This, in addition to a clear increase in patient longevity globally,
45 mandates more research efforts towards care of the elderly post CA [3, 4]. Not unexpectedly,
46 cardiopulmonary resuscitation (CPR) is more commonly initiated in younger patients, and younger
47 patients receive more aggressive treatment by mobile medical teams [4, 5]. Although post-CA
48 mortality increases with age, it has been disputed whether this is due to age in itself or other CA
49 characteristics [6-8]. Pre-arrest comorbidity and CA factors still need more research in order to
50 explain the variability of outcome in CA among elderly [9].

51 Only a few studies have focused on the post CA treatment of the elderly in the intensive care unit
52 (ICU) while including long-term outcome [3, 10]. The aim of this study is to explore treatment
53 intensity, outcome and healthcare-associated costs of the ICU-treated elderly CA patients treated in
54 a single centre over a 9-year period. We hypothesise that compared with younger patients, the elderly
55 have higher mortality and worse neurological outcome despite high treatment costs and intensity.

56 **Methods**

57 **Study design and setting**

58 This retrospective cohort study was conducted at Meilahti Hospital, Helsinki, Finland, which serves
59 as the primary referral centre for CA patients in the Helsinki and Uusimaa region. This region has a
60 population of approximately 1.7 million people (30% of the total Finnish population). Data were
61 extracted from the Finnish Intensive Care Consortium (FICC) database [11] and include adult CA
62 patients (≥ 18 years of age) treated in the hospital's ICU between January 1, 2005 and December 31,
63 2013. We reviewed Electronic health records (EHR) of individual patients for relevant data. Patients
64 with incomplete or missing data and patients where return of spontaneous circulation (ROSC) was
65 not achieved were excluded from the analyses. The patients were divided into two age groups for
66 descriptive purposes: <75 (young) and ≥ 75 years (elderly).

67 The study was approved by the ethics committee of the Operative Division of Helsinki University
68 Hospital (June 2014: 194/13/03/02/2014 TMK02 §97), the Finnish National Institute for Health and
69 Welfare (May 2014: THL/713/5.05.01/2014), Statistics Finland (May 2014: TK-53-1047-14), the

70 Social Insurance Institution (September 2015: Kela 55/522/2015) and the Office of the Data
71 Protection Ombudsman (February 2016: 2794/204/2015).

72 **Data collection and extracted variables**

73 The FICC database provided data on hospital survival, preadmission physical status (a modified
74 World Health Organization/Eastern Cooperative Oncology Group (WHO/ECOG) classification
75 implemented by FICC), mean TISS-76 score and its components for the complete ICU stay, and
76 Acute Physiology and Chronic Health Evaluation II (APACHE II) components and scores [12-15].
77 In this study APACHE II scores were used excluding the points for age. We obtained the confirmed
78 date of death by linking the patients' unique personal identification numbers with the Finnish
79 Population Register Centre database, which registers all deaths of Finnish residents. Detailed
80 information regarding preadmission physical status, time of CA, time to ROSC, initial CA rhythm
81 and location as well as CPC for survivors at one year after CA was collected from the hospital's EHR
82 [16-19]. We determined preadmission functional status by using a simplified WHO/ECOG
83 classification, where "independent" was defined as the patient being independent in self-care and
84 "dependent" was defined as the patient being partly or fully dependent on help in self-care prior to
85 hospital admission. [20]. A favourable neurological outcome was defined as CPC scores of 1–2 and
86 an unfavourable neurological outcome as CPC scores of 3–4 [19].

87 **Healthcare-associated costs**

88 Healthcare-associated costs included three parameters: index hospital costs, rehabilitation costs and
89 social security costs. We obtained hospital costs from the hospital's billing records. These included
90 costs incurred during the entire treatment period, such as costs of personnel, surgery, diagnostics as
91 well as ICU and ward stay. Rehabilitation costs were calculated by multiplying the length of stay
92 (LOS) in the rehabilitation unit with the average cost per day for the respective level of care unit [21].
93 Social security costs were retrieved from the national Social Insurance Institution. This is a
94 government-based social security and healthcare system. All reimbursements made by the Social
95 Insurance Institution, up to one year after the admission, were obtained and summed. These included
96 disability allowances, sickness allowances, private physician and physiotherapist expenses,
97 prescription drug expenses and medical transport expenses. All costs were converted to euros based
98 on the 2021 currency rate. Cost data analysis included the calculation of median healthcare costs for
99 each age group and separately for the survivors with a favourable neurological outcome in the studied
100 age groups.

101 Effective cost per survivor with favourable neurological outcome (ECPSFNO) was calculated by
102 dividing the sum of the total cost for all patients within each age group by the number of patients
103 within that group with a favourable neurological outcome (CPC of 1–2) after 12 months [22]. We
104 further stratified costs according to the location of CA (out-of-hospital CA (OHCA), in-hospital CA
105 (IHCA) and in-ICU CA (ICUCA)). A mean of hospital, rehabilitation and social insurance institution
106 costs was separately calculated for each age group.

107 **Statistical analysis**

108 For statistical analyses we used SPSS statistics for MAC, version 25.0, released in 2017 (IBM Corp,
109 Armonk, NY, USA). The baseline characteristics of the study cohort are described using proportions
110 with percentiles for categorical values and medians with interquartile range for continuous variables.
111 We tested group differences with Mann-Whitney U-test or Chi-square test, as appropriate. Logistic
112 regression was used with original data to calculate univariable odds ratios with corresponding 95%
113 confidence intervals regarding impact on mortality and neurological outcome. Significant factors
114 were included in a multivariate regression model to identify independent predictors of unfavourable
115 neurological outcome and mortality. We illustrated the difference in mortality between the two age
116 groups by using Kaplan Meyer survival curves and a clustered bar of cumulative percentages.
117 Chi-square tests were used on TISS-point distribution to determine if there were significant
118 differences in treatment intensities between the two age groups and if the location of CA (OHCA or
119 IHCA) affected/influenced treatment intensity.

120 **Results**

121 **Study population and factors at resuscitation**

122 The study included 1,285 patients, of which 1,073 (84%) were younger than 75 years and 212 (16%)
123 75 years or older (Table 1). OHCA were less common among the elderly with an occurrence of 43%
124 compared to 64% in the young group, $p < 0.001$ (Table 1). A number of other differences between
125 the younger and the elderly population were noted: less elderly patients had an independent
126 preadmission functional status (75% vs. 90%, $p < 0.001$), a non-shockable initial CA rhythm was
127 more common (49% vs. 35%, $p < 0.001$), and ROSC was achieved faster among the elderly patients
128 (median of 10 min vs. 16 min, $p < 0.001$).

129 **Table 1.** Patients characteristics

	Age <75 (n=1073)	Age ≥75 (n=212)	<i>p</i>
Women, % (no.)	281 (26)	70 (33)	0.041
Location of arrest, % (no.)			<0.001
OHCA	64 (691)	43 (92)	
IHCA	27 (286)	46 (97)	
ICUCA	9 (96)	11 (23)	
Women, % (no.)	281 (26)	70 (33)	0.041
Witnessed arrest, % (no.)¹	935 (87)	193 (91)	0.130
Initial cardiac-arrest rhythm, % (no.)			<0.001
Shockable (VT or VF)	641 (60)	94 (44)	
Non-Shockable (all other rhythms)	378 (35)	104 (49)	
Unknown	54 (5)	14 (6.6)	
Time to ROSC in minutes, median (IQR)²	16 (10-23)	10 (5-18)	<0.001
Independent preadmission functional status % (no.)³	90 (960)	75 (158)	<0.001

130 ¹2% of patients are missing this information

131 ²9,5% of patients are missing this information

132 ³4.7% of patients are missing this information

133

134 **Treatment intensity & ICU factors**

135 No difference was observed in the APACHE II scores between the elderly and younger patients when
 136 points for age were excluded (Table 2). Treatment intensity was lower in the elderly than in the
 137 younger age group, with median daily average TISS scores of 34 and 37 for the elderly and younger
 138 patients, respectively, $p < 0.001$. The total amount of TISS points was also lower for the elderly (116
 139 vs. 147, $p < 0.001$) (Table 2). In-hospital as well as in-ICU mortality was higher for the elderly group
 140 (ICU mortality 33% vs. 18%, $p < 0.001$; hospital mortality 49% vs. 33%, $p < 0.001$). The ICU LOS
 141 was shorter for the elderly than for the younger patients (Table 2). The ICU LOS among the survivors
 142 was however not different. Table 2 details the ICU factors, in-hospital mortality, TISS-point
 143 distribution and the difference in the selected treatments received at the hospital. TISS-point
 144 distribution can be viewed in more detail in the supplementary material (Additional file 1–3).

145 **Table 2.** Intensive care unit-factors

	Age <75 (n=1073)	Age ≥75 (n=212)	<i>p</i>
APACHE II-score excluding age points, median (IQR)	20 (15-27)	22 (15-27)	0.181
TISS-Score, median (IQR)			
Daily average	37 (31-43)	34 (28-41)	<0.001
Total TISS-score	147 (93-227)	116 (65-192)	<0.001
Treatments received, no. (%)			
Controlled ventilation with or without PEEP	1055 (98)	197 (93)	<0.001
Induced hypothermia	450 (42)	34 (16)	<0.001
Vasoactive drug infusion (> 1 drug)	503 (47)	80 (38)	0.015
Continuous antiarrhythmia infusions	212 (20)	35 (17)	0.273
Rx of seizures	140 (13)	13 (6)	0.004
Hemodialysis in unstable patient	18 (2)	8 (4)	0.048
Arterial line	1071 (100)	211 (100)	0.432
In-hospital mortality %(no.)			
Dead in ICU	18 (194)	33 (69)	<0.001
Dead in hospital	33 (357)	49(104)	<0.001
Length of stay in days, median (IQR)			
ICU	3 (2-5)	2 (1-4)	<0.001
Hospital	10 (4-20)	8 (3-16)	0.003
Length of stay in days among patients discharged alive, median (IQR)			
ICU	3 (2-6)	3 (2-6)	0.085
Hospital	14 (8-24)	14 (8-23)	0.654

146

147 **Healthcare-associated costs**

148 The ECPSFNO was €168,000 and €120,000 for the elderly and young group, respectively. The
 149 effective cost for the elderly patient group was higher than that for the younger patient group in all
 150 locations of CA except for ICUCA, where it was €173,000 and €308,000, respectively. The elderly
 151 patient group received less median funding from the Social Insurance Institution, €714 compared to
 152 €1,670 in the younger age group (Table 3, Figure 1, Additional file 4). Median rehabilitation costs
 153 were higher for the elderly patient group when we only included those with a favourable 12-month
 154 neurological outcome, 5,700 compared to 2,000 (*p* = 0.012) (Table 3).

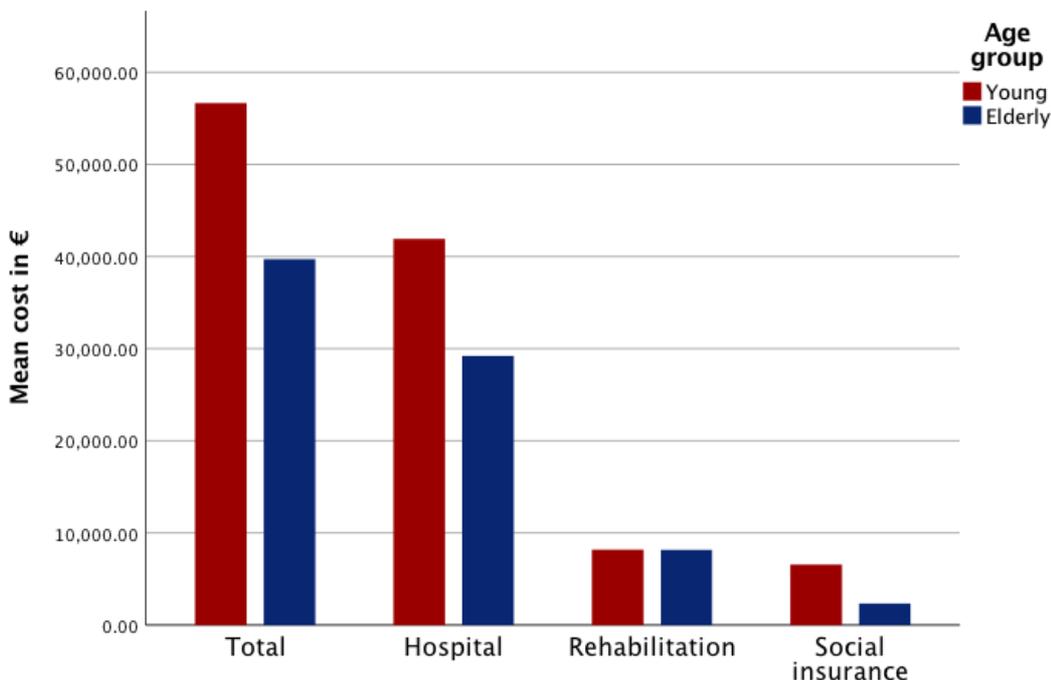
155 **Table 3.** Resource use

	Age <75 (n=1073)	Age ≥75 (n=212)	<i>p</i>
Cost of treatments in €, median (IQR)			
Hospital costs	29 971 (13 381-50 212)	18 356 (9 740-37 102)	<0.001

Rehabilitation	0 (0-6 543)	0 (0-7 574)	0.928
Social Insurance Institution	1 669 (579-6 686)	714 (392-1 760)	<0.001
Cost of treatment of those with CPC 1-2 after 12 months in €, median (IQR) (57%)			
Hospital	41 194 (27 031-63 709)	34 888 (19 083-60 029)	0.071
Rehabilitation	2113 (0-9 603)	6073 (417-13 319)	0.012
Social Insurance Institution	4 561 (1 173-15 043)	2 049 (1 196-3 509)	<0.001
Total cost in €, median (IQR)			
Everyone	38 195 (16 505-71680)	22 641 (12 488-47 006)	<0.001
Those with CPC 1-2 after 12 months (57)	54 510 (36 148-86 461)	39 482 (24 101-93 020)	0.040
Effective cost¹ in €			
Of those with CPC 1-2 after 12 months	119 941	168 416	-
Effective cost in € among those with CPC 1-2 after 12 months			
OHCA	90 499	133 134	-
IHCA	161 670	199 540	-
ICUCA	308 000	172 595	-

156 ¹Effective cost: The total healthcare-associated costs of all patients within their respective age group
157 divided by the number of survivors with a favourable neurological outcome

158 **Figure 1.** Mean cost distribution in 2021 euros



159

160 **Neurological outcome and mortality**

161 Neurological outcome was worse for the elderly group, with only 24% (50/212 patients) having CPC
 162 scores of 1–2 after 12 months, compared with 47% (507/1073 patients) of the younger age group, p
 163 < 0.001 . Long-term mortality was higher for the elderly group compared to the younger group; 70%
 164 of the elderly (vs. 44%) had died within two years, $p < 0.001$. Mortality in the elderly versus the
 165 younger age group during the first year is shown in Figure 2. Separate Kaplan Meier curves
 166 illustrating mortality during the whole follow-up period for all patients, patients based on location of
 167 arrest (OHCA, IHCA and ICUCA) as well as based on initial rhythm (shockable and non-shockable)
 168 can be found in the supplementary material (Additional file 5-6). The median follow-up time was 1.6
 169 years per patient.

170 Factors independently associated with unfavourable neurological outcome was age above 75 years
 171 (OR = 3.27, 95% CI: 2.12-5.03, $p < 0.001$), dependent pre-admission functional status (OR = 3.13,
 172 95% CI: 1.69-5.79, $p < 0.001$), non-shockable initial CA rhythm (OR shockable rhythm = 0.43, 95%
 173 CI: 0.30-0.60, $p < 0.001$), time to ROSC in 10 minutes (OR = 1.61, 95% CI: 1.34-1.93, $p < 0.001$)
 174 and APACHE II score excluding points for age (OR = 1.96, 95% CI: 1.61–2.39, $p < 0.001$).

175 Factors independently associated with mortality were age above 75 years (OR = 3.36, 95% CI: 2.21-
 176 5.11, $p < 0.001$), dependent pre-admission functional status (OR = 2.96, 95% CI: 1.60-5.50, $p <$
 177 0.001), initial CA rhythm (OR shockable rhythm= 0.56, 95% CI: 0.40-0.77, $p < 0.001$), location of
 178 CA (ICUCA OR = 2.27, 95% CI: 1.26-4.08, $p = 0.006$), time to ROSC (OR = 1.36, 95% CI: 1.16-
 179 1.60, $p < 0.001$) and APACHE II score excluding age (OR = 2.06, 95% CI: 1.70-2.50, $p < 0.001$).
 180 Table 4 details the independent predictors of an unfavourable neurological outcome and Table 5 the
 181 independent predictors of mortality.

182 **Table 4.** Univariate models and multivariate models for risk factors predicting 12-month
 183 unfavourable cerebral performance status

Variable	Univariate model		Multivariate model	
	OR (95% CI)	p	OR (95% CI)	p
Age				
Young (<75y)	1		1	
Elderly (>=75)	3.09 (2.19-4.36)	<0.001	3.27 (2.12-5.03)	<0.001
Pre-admission functional status				
Independent	1		1	
Dependent	4.07 (2.48-6.67)	<0.001	3.13 (1.69-5.79)	<0.001
Initial CA-rhythm				
Non-shockable	1		1	

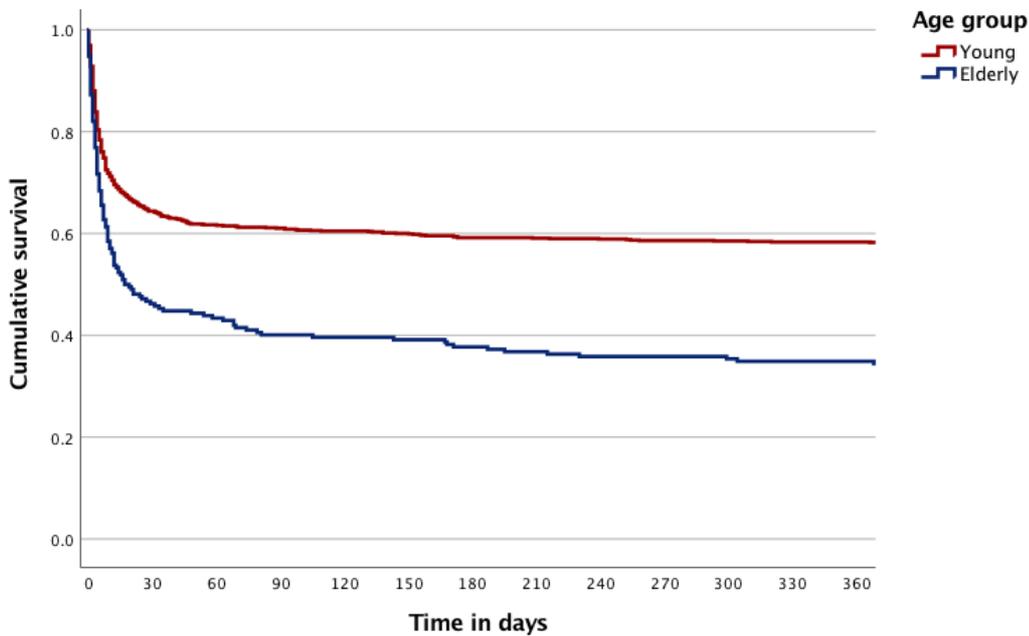
Shockable	0.32 (0.25-0.41)	<0.001	0.43 (0.30-0.60)	<0.001
Location of arrest				
OHCA	1		1	
IHCA	1.78 (1.37-2.30)	<0.001	1.46 (0.98-2.18))	0.065
ICUCA	2.24 (1.47-3.41)	<0.001	2.24(1.21-4.16)	0.010
Witnessed arrest (not witnessed=1)	0.47 (0.31-0.70)	<0.001	0.70(0.43-1.12)	0.134
Time to ROSC in 10 minutes	1.15 (1.02-1.29)	0.020	1.61 (1.34-1.93)	<0.001
APACHE II-score excluding age points¹	2.51 (2.13-2.95)	<0.001	1.96 (1.61-2.39)	<0.001
A total of 980 patients were included. 94 patients had missing CPC, 60 patients had missing functional status, 68 patients had missing initial rhythm, 24 patients had missing if the arrest was witnessed, 118 patients had missing time to ROSC, 1 patient had missing APACHE II-score.				

184 ¹ Each step increases the variable by 10

185 **Table 5.** Univariate models and multivariate models for risk factors predicting 12-month mortality.

Variable	Univariate model		Multivariate model	
	OR (95% CI)	<i>p</i>	OR (95% CI)	<i>p</i>
Age				
Young (<75y)	1		1	
Elderly (>=75)	3.44 (2.44-4.84)	<0.001	3.36 (2.21-5.11)	<0.001
Pre-admission functional status				
Independent	1		1	
Dependent	4.20 (2.55-6.92)	<0.001	2.96 (1.60-5.50)	<0.001
Initial CA-rhythm				
Non-shockable	1		1	
Shockable	0.33 (0.26-0.42)	<0.001	0.56 (0.40-0.77)	<0.001
Location of arrest				
OHCA	1		1	
IHCA	2.07 (1.61-2.67)	<0.001	1.61 (1.11-2.34)	0.12
ICUCA	2.29 (1.52-3.449)	<0.001	2.27 (1.26-4.08)	0.006
Witnessed arrest (not witnessed=1)	0.59 (0.40-0.86)	<0.006	0.80 (0.51-1.27)	0.353
Time to ROSC in 10 minutes	1.04 (0.93-1.16)	0.520	1.36 (1.16-1.60)	<0.001
APACHE II-score excluding age points¹	2.34 (2.01-2.73)	<0.001	2.06 (1.70-2.50)	<0.001
A total of 1055 patients were included. 60 patients had missing functional status, 68 patients had missing initial rhythm, 24 patients had missing if the arrest was witnessed, 118 patients had missing time to ROSC, 1 patient had missing APACHE II-score.				

186 **Figure 2.** Kaplan-Meier survival curve of all cases during the first year, Log Rank $p < 0.001$



187

188 Discussion

189 This current study presents a comprehensive estimation of CA-associated costs including hospital
190 costs, rehabilitation costs and social insurance costs at a tertiary university hospital. Long-term
191 survival and functional outcome was as expected significantly lower among the elderly. In addition
192 they received less intensive ICU treatment and had a shorter ICU LOS overall. Cost effectiveness
193 was lower for the elderly patient group than for the younger patient group in most cases except
194 ICUCAs. Given our ageing population, our study highlights the need to individually assess the cost
195 effectiveness of care post CA.

196 Both TISS-point distribution and the median total cost are lower for the elderly group. This in
197 combination with an even APACHE-score distribution between the age groups (indicating roughly
198 the same comorbidity pre-CA) indicates that age seems to have been a factor affecting treatment
199 intensity. The difference in total TISS-point-distribution can be affected by the LOS, but the LOS
200 does not explain the difference in average daily TISS-points. We speculate that initial treatment
201 intensity was high for both age groups, but that some treatments were stopped earlier in the elderly
202 group due to a perceived poor prognosis, thus decreasing the average daily TISS-score. This indicates
203 that an active daily evaluation of patients was in use in order to allocate resources in a more beneficial
204 way. Recent studies demonstrate that ROSC rates, one-year survival and favourable neurological
205 outcome at one month among elderly CA patients have increased over time with increase in the

206 proportion of advanced in-hospital treatments (i.e. extracorporeal membrane oxygenation,
207 therapeutic hypothermia and/or percutaneous coronary angiogram/intervention) provided [23]. In our
208 study, in-hospital costs of the total provided treatments were lower for the elderly, even when
209 excluding those with a poor one-year neurological outcome. One could argue that the elderly may not
210 benefit from more aggressive treatment, but age in itself should not affect the administered treatments
211 even if it affects mortality, as neurological outcome seems to remain good for survivors [3]. In this
212 study, the ECPSFNO was higher for the elderly group in all locations of CA except ICUCA. Thus,
213 although fewer resources were used by the elderly, the cost per survivor remained higher than the
214 younger age group owing to the high mortality in the elderly group.

215 Due to marked differences in healthcare funding, direct comparisons of our results with other studies
216 are difficult. Our results also indicate a clear inter-patient variation. Costs in the range of €20,000–
217 40,000 for ICU-treated CA survivors have been determined in previous studies [24, 25]. Nonetheless,
218 several studies and meta-analyses have shown that age negatively affects post-CA mortality [26].
219 Long-term survival among elderly CA patients is generally lower than that among younger age groups
220 in the case of OHCA [27, 28].

221 We also looked at the distribution of costs in three separate categories (hospital costs, rehabilitation
222 costs and social insurance costs) among different age groups. Costs were higher for the younger
223 patient group in all categories except rehabilitation. The difference in hospital costs could be
224 attributable to the elderly receiving less aggressive treatment and having to be in a better initial
225 condition in order to survive CA and be taken to the ICU. Less intensive treatment is needed to attain
226 a favourable outcome if the pre-arrest comorbidities are lower, which also decreases hospital costs.
227 The younger age group probably received more funding from the Social Insurance Institution because
228 they got a paid sick-leave from work. Patients over 68 years of age receive pension, which does not
229 alter if the patient is severely ill and therefore isn't included in these calculations. The higher risk of
230 early post-CA mortality in elderly patients might also have decreased social insurance
231 reimbursements as compared to the younger patients. We noticed that median rehabilitation costs
232 were higher among the elderly when only including those with a favourable 12-month neurological
233 outcome. This could be an indicator for the elderly having more long-time problems post-CA. We
234 can probably not see the same difference in rehabilitation costs when taking into account all patients
235 as the early mortality among elderly decreases the median rehabilitation costs. The effect of less
236 aggressive treatment on the need for rehabilitation among elderly is something further research could
237 focus on.

238 Additionally, we demonstrated worse long-term outcome in elderly compared to younger patients
239 following care in the ICU after CA. This difference was the most pronounced in OHCA but was
240 evident in patients with IHCA as well. Interestingly, this study shows that the same percentage of
241 patients in both age groups had a CPC of 3-4 12 months post CA, but our multivariable model still
242 indicates that age affects neurological outcome. It is debatable how much old age correlates with
243 worse neurological outcome as the high mortality probably affects the statistics. Previous studies also
244 indicate that there isn't a difference in neurological outcome among elderly compared to younger CA
245 survivors[3, 28]. It is worth noting that age does not always correlate with outcome and is not in itself
246 an adequate prognostic factor, as two elderly persons of the same age can have very different medical
247 conditions [29]. High frailty and a low performance status have been connected with higher
248 ECPSFNO and mortality in previous studies [30-33]. The increased mortality related to age in this
249 study is indeed partially explained by the pre-admission functional status of the elderly patients; thus,
250 this in combination with age seems to better predict both mortality and neurological outcome.
251 Performance status could be a more precise tool when deciding which patients benefit the most from
252 intensive care and more advanced treatment options.

253 Another possible tool for risk assessment among the elderly seems to be the initial rhythm, as an
254 initially shockable rhythm predicts a better outcome even among the very elderly, where other
255 prognostic factors seem to fail [34]. Supporting the results of previous studies, initial rhythm was one
256 of the factors with the strongest association with outcome among both the younger and elderly
257 patients in this study as well. Interestingly, in our study the elderly had lower incidences of ventricular
258 fibrillation (VF) and ventricular tachycardia (VT) than the younger age group. This could be related
259 to a difference in the aetiology of the arrests or to mechanisms such as faster conversion of VF/VT to
260 asystole owing to the faster depletion of energy in the aged heart. Such an abnormality has been
261 described in mitochondrial metabolism with ageing in the muscle cells [35]. Previous studies have
262 also shown that bradyarrhythmia-related CA patients were generally older than those with
263 tachyarrhythmia-related CA [36]. The significant difference in location of arrest is also something
264 that affects mortality and neurological outcome as CA aetiology differs depending on where the CA
265 occurred [37]. We may speculate that as the elderly had a higher percentage of IHCA, they also had
266 a higher amount of unfavourable pre-arrest comorbidities, which increases mortality as these
267 comorbidities affect CA-aetiology and initial rhythm in an unfavourable manner [38]. Factors in
268 IHCA that may decrease mortality compared to OHCA are shorter times to response and more
269 available treatments, but these do probably not affect mortality as much as the pre-arrest
270 comorbidities seem to do.

271 A major strength of this study is its minimal selection bias owing to socioeconomic factors and
272 personal insurance, as this study was conducted in a setting of government-funded healthcare. A
273 limitation and factor affecting the outcome of this study is that the studied population only included
274 patients with ROSC who were treated in the ICU; this immediately excludes patients in such a bad
275 initial state that they did not survive until admission to the ICU and patients with low chances of
276 benefiting from intensive treatment. This leads to a selection bias especially among the elderly, with
277 those surviving to the ICU having a higher probability of surviving altogether and being in a better
278 initial condition compared to the mean population with the same age.

279 **Conclusion**

280 The interpretation of this study is that treatment intensity for the elderly is lower as a group, the
281 mortality, the risk for a poor neurological outcome is higher and the cost-effectiveness is lower
282 compared to the younger age group. Further studies should focus on the tools for identification of
283 patients who can benefit from a more aggressive treatment approach, enabling an improvement in the
284 resource distribution in post-CA ICU care in the elderly.

285 **Abbreviations**

286 CA: Cardiac arrest; ICU: intensive care unit; TISS: Therapeutic scoring system; CPC: cerebral performance
287 category; CPR: cardiopulmonary resuscitation; FICC: Finnish Intensive Care Consortium; EHR: electronic
288 health records; ROSC: return of spontaneous circulation; WHO/ECOG: World Health Organization/Eastern
289 Cooperative Oncology Group, APACHE: Acute Physiology and Chronic Health Evaluation; LOS: length of
290 stay; ECPSFNO: Effective cost per survivor with favourable neurological outcome; OHCA: out-of-hospital
291 cardiac arrest; IHCA: in-hospital cardiac arrest; ICUCA: in-intensive-care-unit cardiac arrest

292 **Supplementary Information**

293 **Additional file 1.** Table of TISS-point distribution for individual procedures

294 **Additional file 2.** Table of TISS-point distribution for individual procedures (OHCA-cases only)

295 **Additional file 3.** Table of TISS-point distribution for individual procedures (IHCA-cases only)

296 **Additional file 4.** Mean cost in euro based on initial rhythm a) shockable rhythms b) non-shockable rhythms

297 **Additional file 5.** KM-curves based on location of arrest a) all cases during the whole follow up-period, Log
298 Rank $p < 0.001$ b) OHCA, Log Rank $p < 0.001$ c) IHCA, Log Rank $p = 0.003$ d) ICUCA, Log Rank $p =$
299 0.079

300 **Additional file 6.** KM-curves based on initial rhythm a) Shockable rhythm (VF/VT), Log rank $p < 0.001$ b)
301 Non-shockable rhythm, Log Rank $p = 0.062$

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304 **Authors contribution**

305 EH and MBS planned and conceived the study and wrote the first draft. RR assisted with the statistical
306 analysis. All other authors revised the manuscript for intellectual content. All authors approve the
307 submission. EH and MBS take intellectual responsibility for the study findings.

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312 decision to submit the manuscript for publication.

313 **Availability of data and materials**

314 Legal restrictions prevent us from making the data publicly available, as it is based on patient registers. The
315 data included in this study are obtained from several databases (The Finnish Intensive Care Consortium, Kela,
316 the Finnish National Institute for Health and Welfare, Statistics of Finland and the five university hospitals in
317 Finland). With appropriate research approval, data can be directly requested from the sources.

318 **Declarations**

319 **Ethics approval and consent to participate**

320 The study was approved by the Finnish National Institute for Health and Welfare (December 2017:
321 THL/2034/5.05.00/2017) and the board of the Hospital District of Helsinki and Uusimaa (March 2018:
322 HUS/26/2018).

323 **Consent for publication**

324 Not applicable.

325 **Competing interests**

326 The authors declare that they have no competing interests.

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370 *Council on Resuscitation, Heart and Stroke Foundation of Canada, InterAmerican Heart*
371 *Foundation, Resuscitation Council of Southern Africa, Resuscitation Council of Asia); and the*
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425 Tables and figures

- 426 **Table 1.** Patients characteristics
- 427 **Table 2.** Intensive care unit-factors
- 428 **Table 3.** Resource use
- 429 **Table 4.** Univariate models and multivariate models for risk factors predicting 12-month unfavourable
- 430 cerebral performance status
- 431 **Table 5.** Univariate models and multivariate models for risk factors predicting 12-month mortality
- 432 **Figure 1.** Mean cost distribution in 2021 euros
- 433 **Figure 2.** Kaplan-Meier survival curve of all cases during the first year, Log Rank $p < 0.001$

434 **Additional Files**

435 **Additional file 1.** Table of TISS-point distribution for individual procedures

Procedure, no (%)	Age <75 (n=1073)	Age ≥75 (n=212)	<i>p</i>
Tracheostomy care	50 (5)	5 (2)	0.130
Controlled ventilation with or without PEEP combined with IMV or assisted ventilation	1055 (98)	197 (93)	<0.001
Cardiac arrest or countershock within 48 h	976 (91)	182 (86)	0.023
Controlled ventilation with intermittent or continuous muscle relaxants	530 (49)	59 (28)	<0.001
Pulmonary artery catheter	271 (25)	46 (22)	0.272
Pacemaker on standby	78 (7)	30 (14)	0.001
Hemofiltration/dialytic techniques	101 (9)	19 (9)	0.837
Induced hypothermia	450 (42)	34 (16)	<0.001
Intra-aortic balloon pressure	51 (5)	16 (8)	0.094
Emergency endoscopy or bronchoscopy	249 (23)	39 (18)	0.125
Vasoactive drug infusion (> 1 drug)	503 (47)	80 (38)	0.015
Intravenous alimentation	286 (27)	43 (20)	0.052
Frequent infusions of blood products (>5 U/24h)	54 (5)	14 (7)	0.350
Vasoactive drug infusion (1 drug)	891 (83)	172 (81)	0.502
Continuous antiarrhythmic infusions	212 (20)	35 (17)	0.273
Cardioversion for arrhythmia	94 (9)	16 (8)	0.564
Arterial line	1071 (100)	211 (100)	0.432
Measurement of cardiac output by any method	390 (36)	48 (23)	<0.001
Active diuresis for fluid overload or cerebral edema	658 (61)	143 (68)	0.092

Active Rx for metabolic acidosis	127 (12)	31 (15)	0.259
Active anticoagulation (initial 48h)	858 (80)	150 (71)	0.003
Rx of seizures	140 (13)	13 (6)	0.004
Central venous pressure	795 (74)	123 (58)	<0.001
Hemodialysis in unstable patient	18 (2)	8 (4)	0.048
Gastrointestinal feedings	337 (31)	53 (25)	0.064
ECG monitoring	1072 (100)	212 (100)	0.657
Hourly vital signs	1071 (100)	212 (100)	0.529
Chronic anticoagulation	679 (63)	114 (54)	0.009
Gastrointestinal decompression	975 (91)	170 (80)	<0.001
PEEP = Positive end-expiratory pressure, IMV = Intermittent mandatory ventilation, ECG = electrocardiogram			

436

437 **Additional file 2.** Table of TISS-point distribution for individual procedures (OHCA-cases only)

Procedure, no (%)	Age <75 (n=691)	Age ≥75 (n=92)	p
Tracheostomy care	20 (3)	0 (0)	0.098
Controlled ventilation with or without PEEP combined with IMV or assisted ventilation	688 (100)	89 (97)	0.003
Cardiac arrest or countershock within 48 h	632 (92)	75 (82)	0.002
Controlled ventilation with intermittent or continuous muscle relaxants	421 (61)	34 (37)	<0.001
Pulmonary artery catheter	5 (1)	0 (0)	0.413
Pacemaker on standby	21 (3)	7 (8)	0.027
Hemofiltration/dialytic techniques	23 (3)	0 (0)	0.076
Induced hypothermia	429 (62)	32 (35)	<0.001
Intra-aortic balloon pressure	13 (2)	2 (2)	0.847
Emergency endoscopy or bronchoscopy	133 (19)	13 (14)	0.236
Vasoactive drug infusion (> 1 drug)	316 (46)	38 (41)	0.423
Intravenous alimentation	166 (24)	14 (15)	0.059
Frequent infusions of blood products (>5 U/24h)	15 (2)	1 (1)	0.490
Vasoactive drug infusion (1 drug)	579 (84)	77 (84)	0.981
Continuous antiarrhythmic infusions	100 (15)	11 (12)	0.516
Cardioversion for arrhythmia	43 (6)	5 (5)	0.767
Arterial line	690 (100)	91 (99)	0.093
Measurement of cardiac output by any method	249 (36)	19 (21)	0.003

Active diuresis for fluid overload or cerebral edema	418 (61)	68 (74)	0.013
Active Rx for metabolic acidosis	48 (7)	7 (8)	0.815
Active anticoagulation (initial 48h)	599 (87)	70 (76)	0.007
Rx of seizures	96 (14)	10 (11)	0.426
Central venous pressure	528 (76)	52 (57)	<0.001
Hemodialysis in unstable patient	1 (0)	0 (0)	0.715
Gastrointestinal feedings	204 (30)	26 (28)	0.803
ECG monitoring	691 (100)	92 (100)	-
Hourly vital signs	691 (100)	92 (100)	-
Chronic anticoagulation	453 (66)	50 (54)	0.035
Gastrointestinal decompression	657 (95)	85 (92)	0.277
PEEP = Positive end-expiratory pressure, IMV = Intermittent mandatory ventilation, ECG = electrocardiogram			

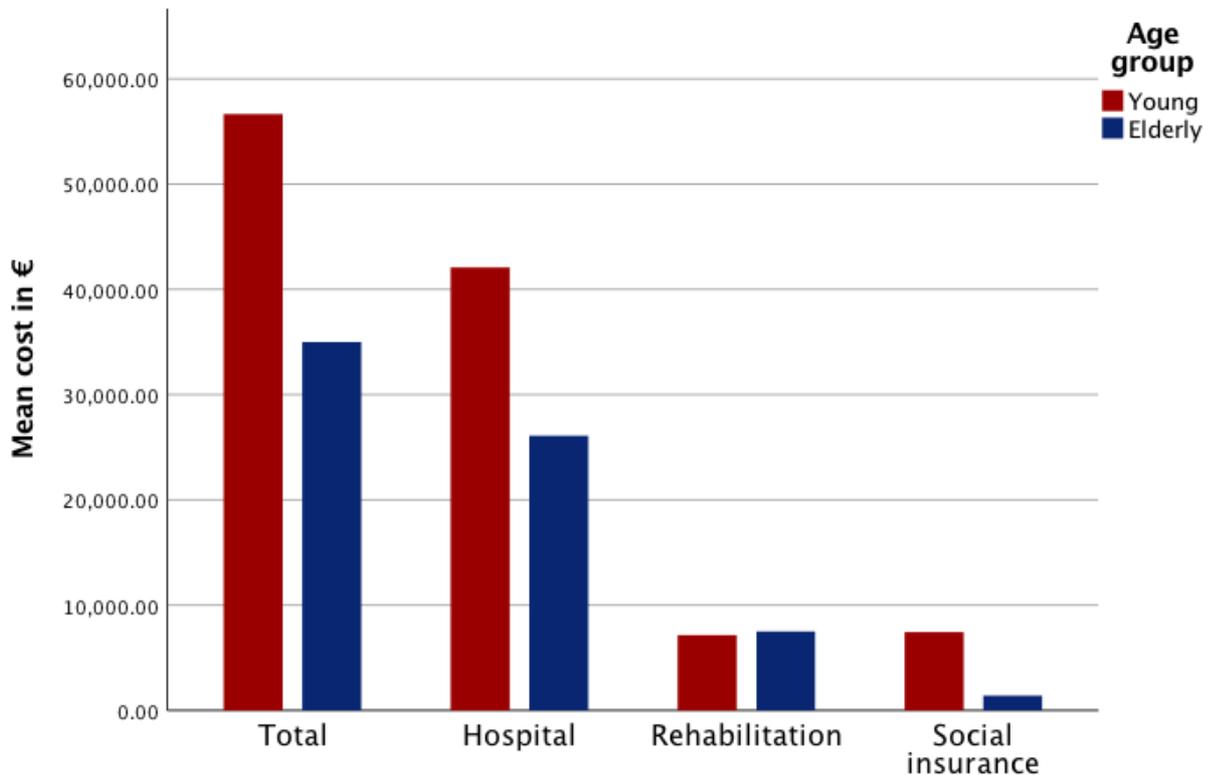
438 **Additional file 3.** Table of TISS-point distribution for individual procedures (IHCA-cases only)
439

Procedure, no (%)	Age <75 (n=382)	Age ≥75 (n=120)	p
Tracheostomy care	30 (8)	5 (4)	0.167
Controlled ventilation with or without PEEP combined with IMV or assisted ventilation	367 (96)	108 (90)	0.010
Cardiac arrest or countershock within 48 h	344 (90)	107 (90)	0.779
Controlled ventilation with intermittent or continuous muscle relaxants	109 (29)	25 (21)	0.096
Pulmonary artery catheter	5 (1)	0 (0)	0.208
Pacemaker on standby	57 (15)	23 (19)	0.268
Hemofiltration/dialytic techniques	78 (20)	84 (16)	0.267
Induced hypothermia	21 (6)	2 (2)	0.080
Intra-aortic balloon pressure	38 (10)	14 (12)	0.590
Emergency endoscopy or bronchoscopy	116 (30)	26 (22)	0.065
Vasoactive drug infusion (> 1 drug)	187 (49)	42 (35)	0.007
Intravenous alimentation	120 (31)	29 (24)	0.130
Frequent infusions of blood products (>5 U/24h)	39 (10)	13 (11)	0.845
Vasoactive drug infusion (1 drug)	312 (82)	95 (80)	0.541
Continuous antiarrhythmic infusions	112 (29)	24 (20)	0.045
Cardioversion for arrhythmia	51 (13)	11 (9)	0.224
Arterial line	381 (100)	120 (100)	0.575

Measurement of cardiac output by any method	141 (37)	29 (24)	0.010
Active diuresis for fluid overload or cerebral edema	240 (63)	75 (63)	0.948
Active Rx for metabolic acidosis	79 (21)	24 (20)	0.872
Active anticoagulation (initial 48h)	259 (68)	80 (67)	0.817
Rx of seizures	44 (12)	3 (3)	0.003
Central venous pressure	267 (70)	71 (59)	0.029
Hemodialysis in unstable patient	17 (5)	8 (7)	0.330
Gastrointestinal feedings	133 (35)	93 (23)	0.012
ECG monitoring	381 (100)	120 (100)	0.575
Hourly vital signs	380 (100)	120 (100)	0.427
Chronic anticoagulation	226 (59)	64 (53)	0.259
Gastrointestinal decompression	318 (83)	85 (71)	0.003
PEEP = Positive end-expiratory pressure, IMV = Intermittent mandatory ventilation, ECG = electrocardiogram			

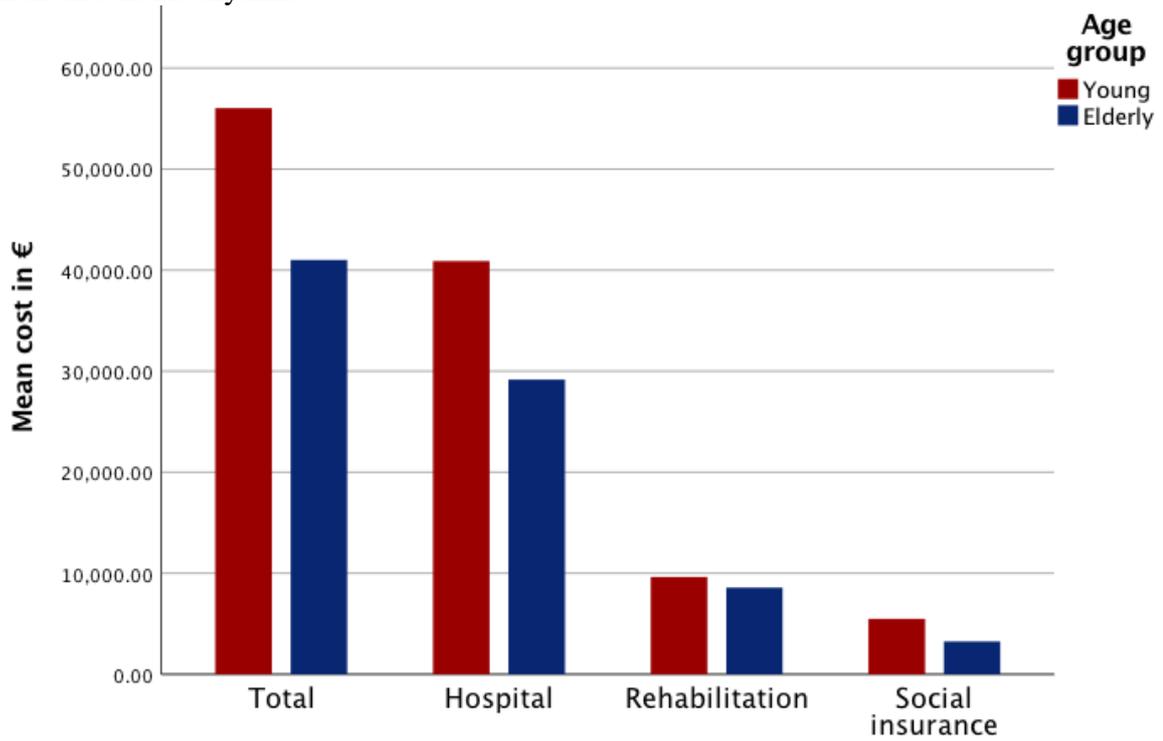
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Additional file 4. Mean cost in euro based on initial rhythm
a) shockable rhythms



444

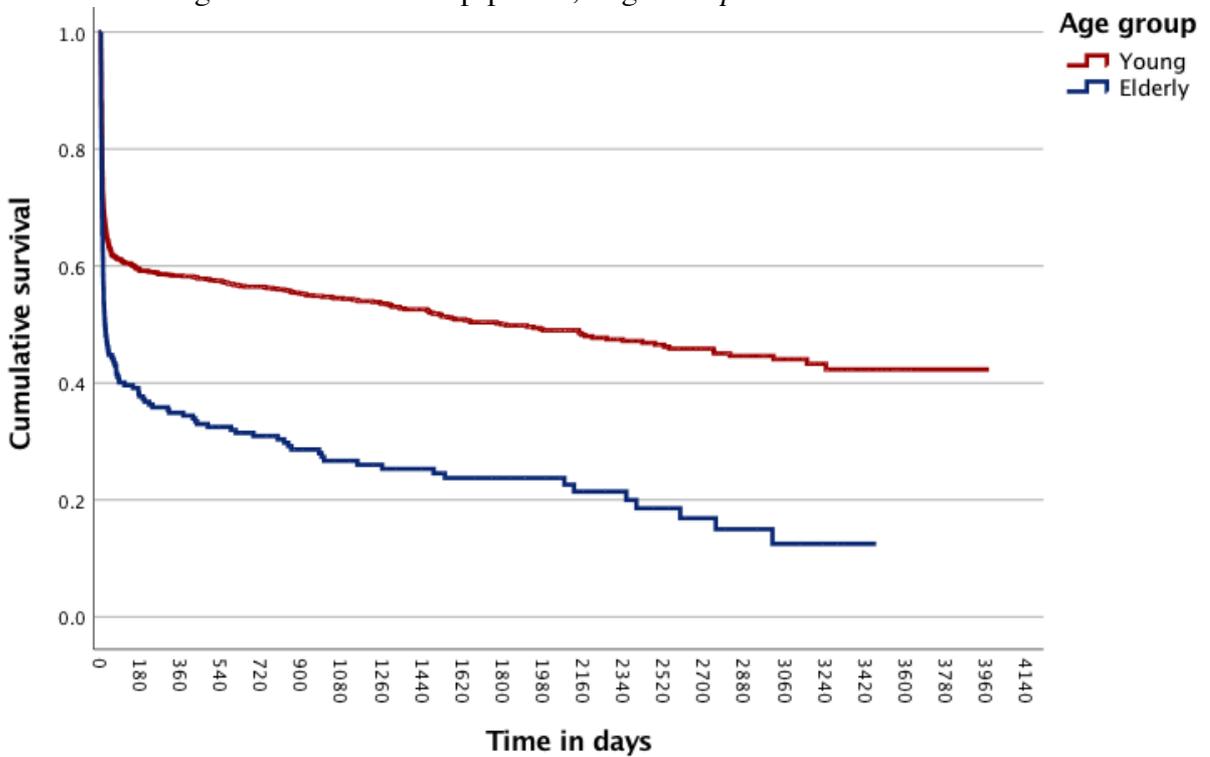
445 b) non-shockable rhythms



446

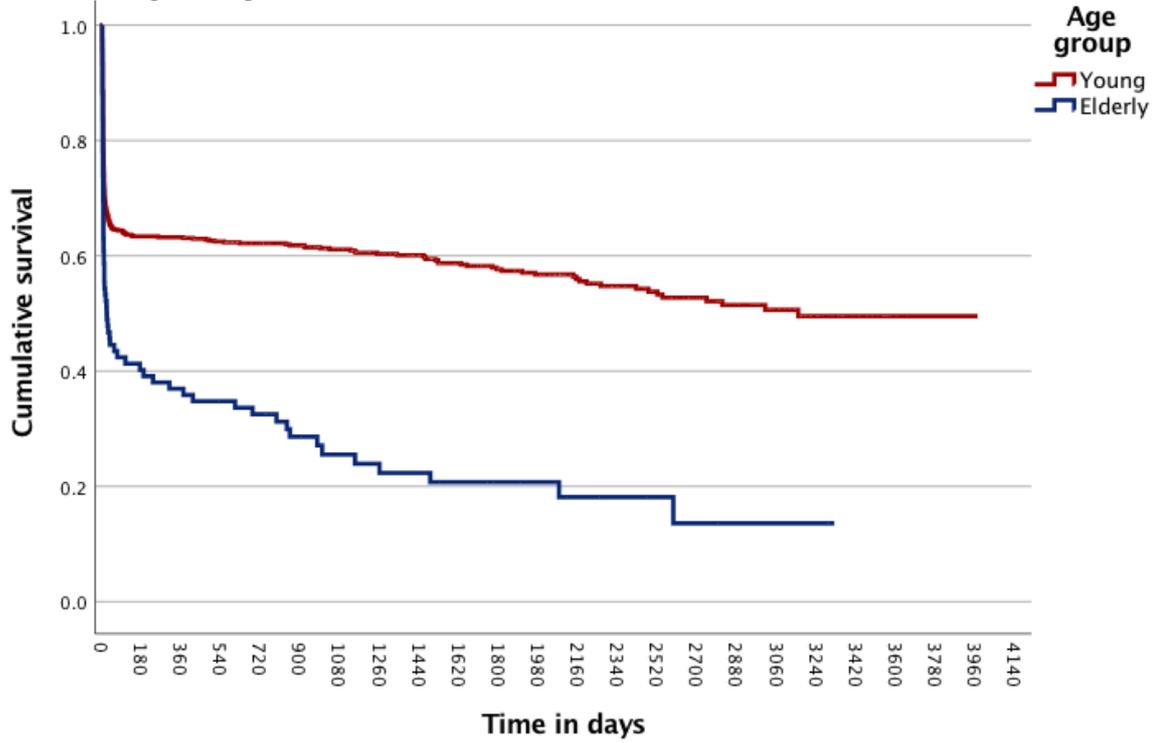
447 **Additional file 5.** KM-curves based on location of arrest

448 a) All cases during the whole follow up-period, Log Rank $p < 0.001$



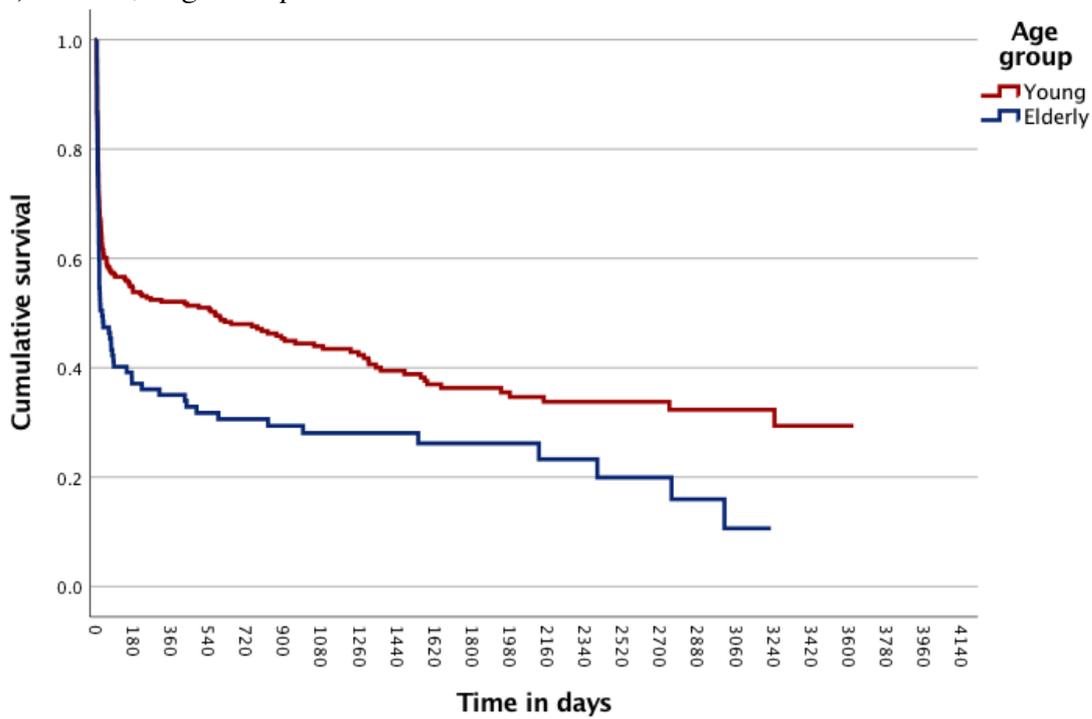
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450 b) OHCA, Log Rank $p < 0.001$



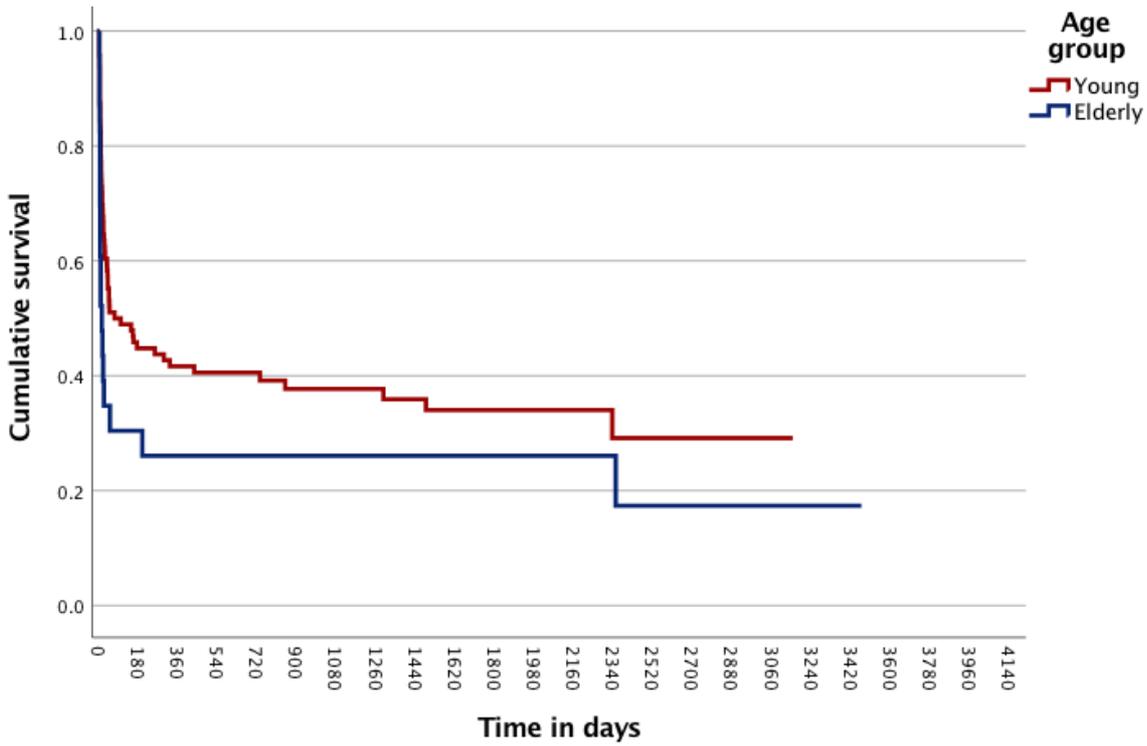
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452 c) IHCA, Log Rank $p = 0.003$



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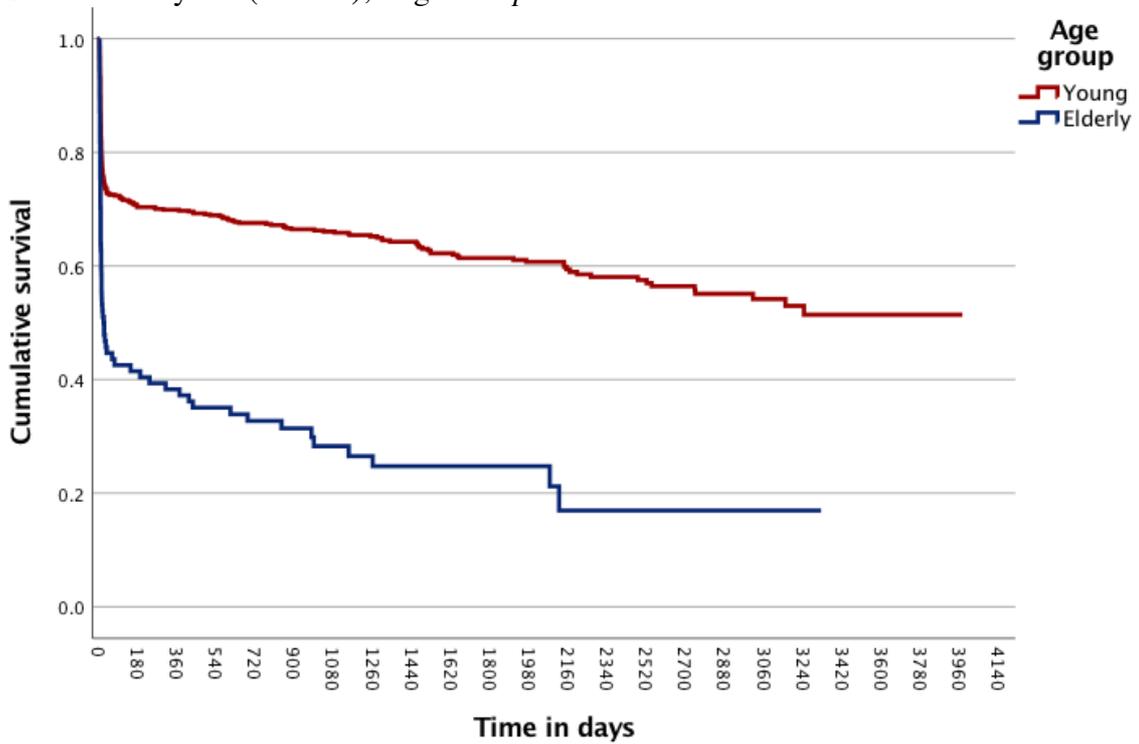
454 d) ICUCA, Log Rank $p = 0.079$



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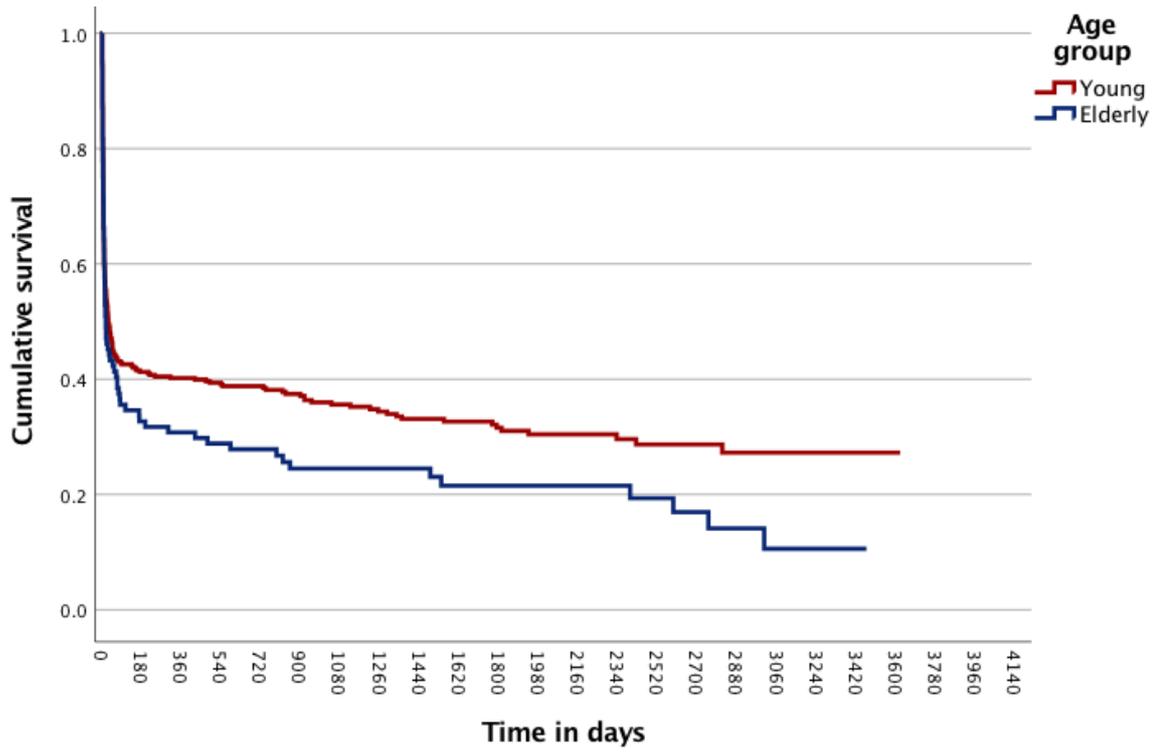
456 **Additional file 6.** KM-curves based on initial rhythm

457 a) Shockable rhythm (VF/VT), Log Rank $p < 0.001$



458

459 b) Non-shockable rhythm, Log Rank $p = 0.062$



460