



NEICS Spring Meeting

North of England Intensive Care Society

Tuesday 24th March 2015

Wynyard Hall, near Middlesbrough

Cardiac arrest – an update

Gavin Perkins

Professor of Critical Care Medicine

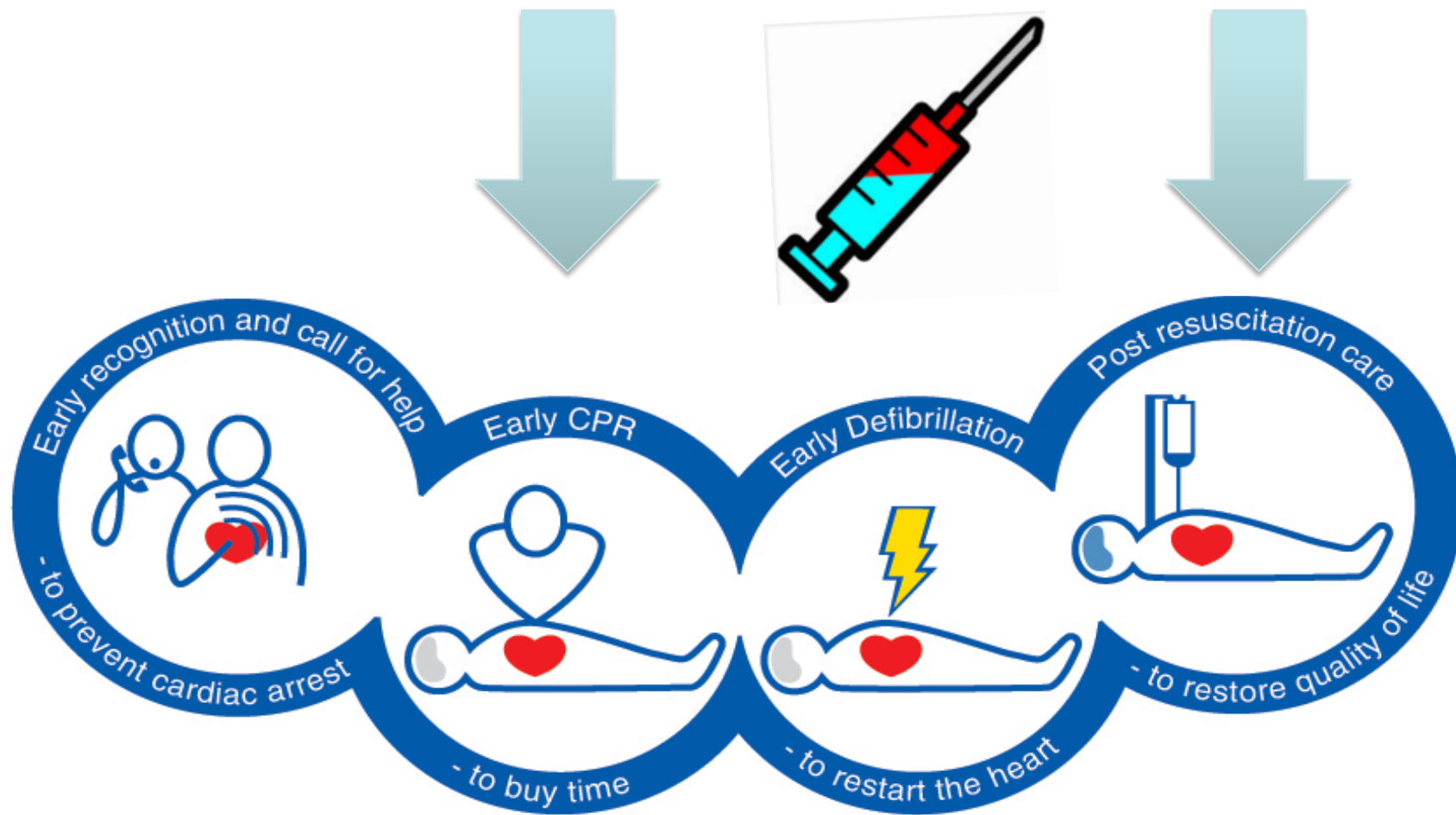
THE UNIVERSITY OF
WARWICK

**HEART of
ENGLAND**
NHS Foundation Trust

**intensive care
foundation**
science saving life

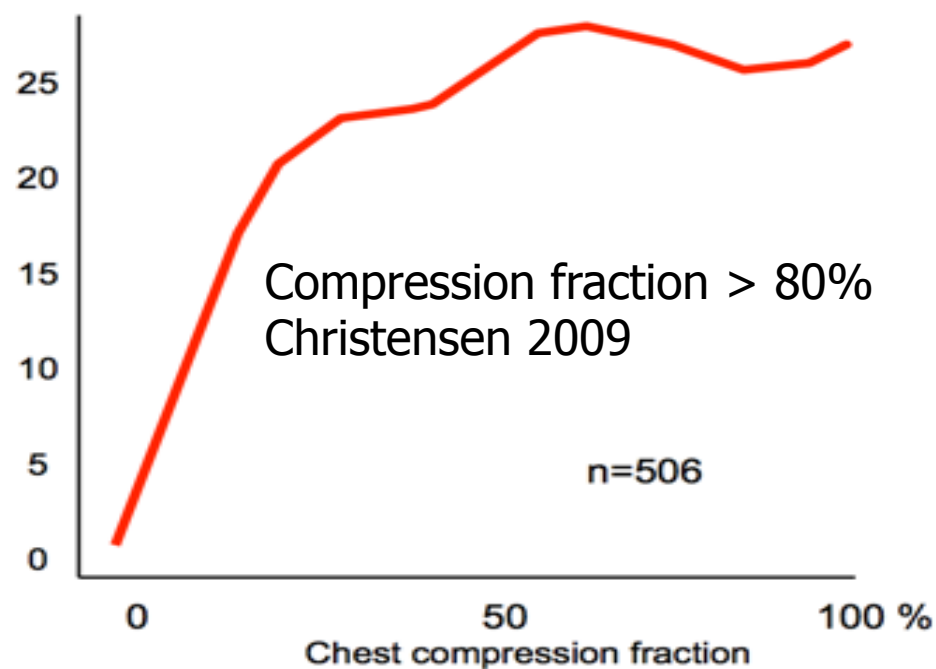
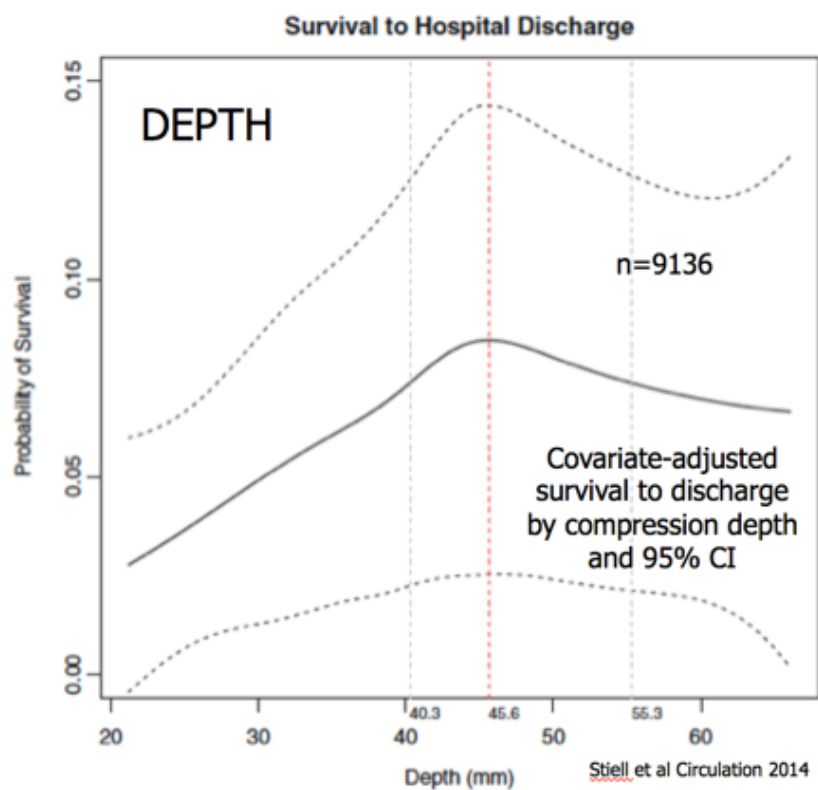
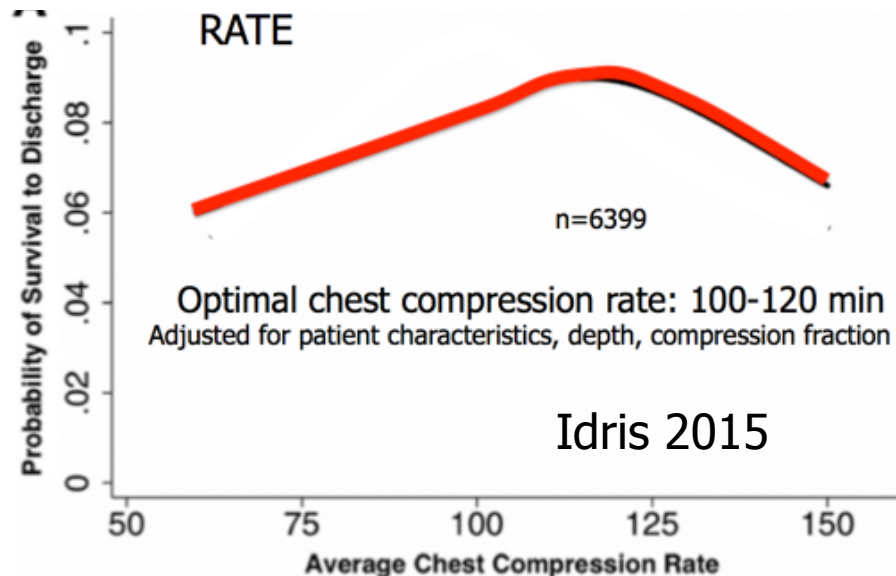
Disclaimer

- The PARAMEDIC trials are funded by the National Institute for Health Research (NIHR) Health Technology Assessment Programme (HTA – 07/37/69 and 12/127/126).
- The views expressed are those of the author(s) and not necessarily those of the NHS, the NIHR, or the Department of Health.



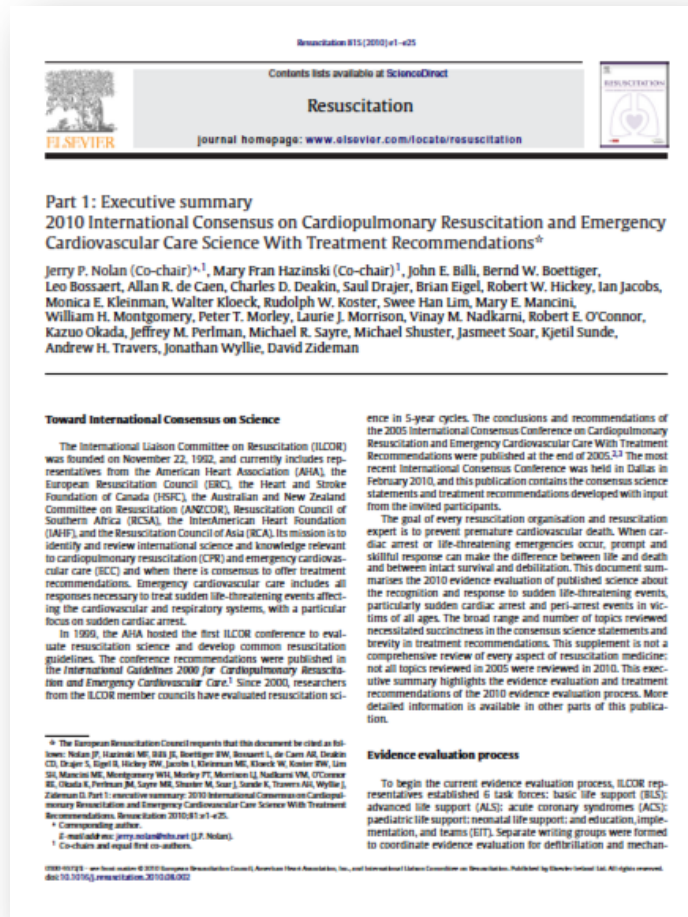
North East Ambulance Service **NHS**

NHS Foundation Trust





ILCOR recommendations 2010



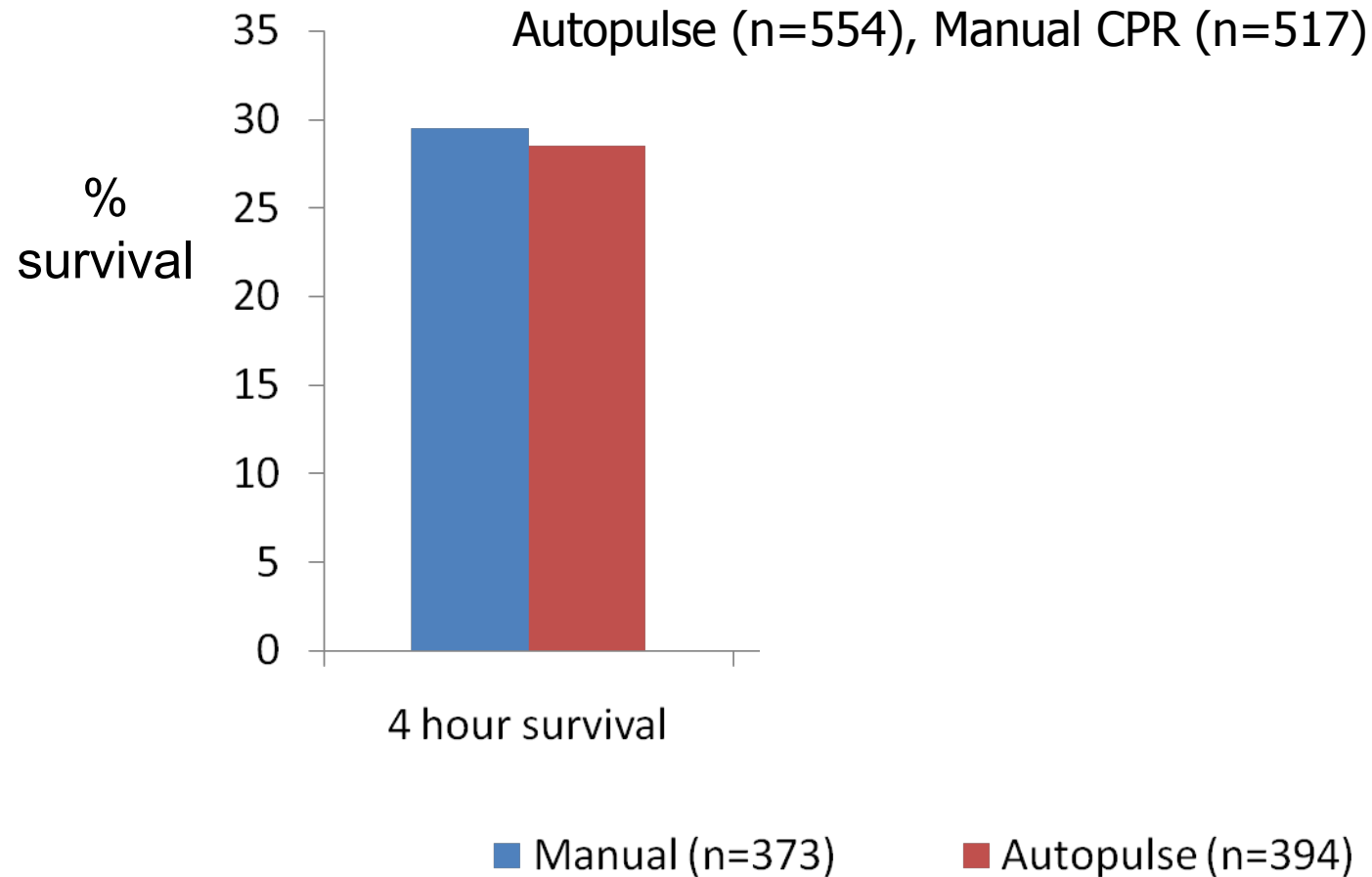
There are insufficient data to support or refute the routine use of load-distributing band CPR, LUCAS CPR, or **mechanical piston CPR, instead of standard CPR.**

On the basis of case reports and case series it may be reasonable to consider LDB or LUCAS CPR to maintain continuous chest compressions while the patient undergoes percutaneous coronary intervention (PCI) or computed tomography (CT) or similar diagnostic studies when provision of manual CPR would be difficult.

Autopulse™



ASPIRE trial (autopulse)





Contents lists available at ScienceDirect

Resuscitation

journal homepage: www.elsevier.com/locate/resuscitation



Clinical Paper

Manual vs. integrated automatic load-distributing band CPR with equal survival after out of hospital cardiac arrest. The randomized CIRC trial^{☆,☆☆}



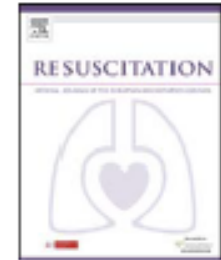
Lars Wik^{a,*}, Jan-Aage Olsen^{a,b}, David Persse^c, Fritz Sterz^d, Michael Lozano Jr.^{e,f}, Marc A. Brouwer^g, Mark Westfall^{h,i}, Chris M. Souders^c, Reinhard Malzer^j, Pierre M. van Grunsven^k, David T. Travis^e, Anne Whitehead^l, Ulrich R. Herken^m, E. Brooke Lernerⁿ



Contents lists available at ScienceDirect

Resuscitation

journal homepage: www.elsevier.com/locate/resuscitation



Clinical paper

Design of the Circulation Improving Resuscitation Care (CIRC) Trial: A new state of the art design for out-of-hospital cardiac arrest research^{☆,☆☆}

E. Brooke Lerner^{a,*}, David Persse^b, Chris M. Souders^b, Fritz Sterz^c, Reinhard Malzer^d, Michael Lozano Jr.^{e,f}, Mark Westfall^{g,h,i,j,k}, Marc A. Brouwer^l, Pierre M. van Grunsven^m, Anne Whiteheadⁿ, Jan-Aage Olsen^o, Ulrich R. Herken^p, Lars Wik^o

- (1) 4 hours training, continuous monitoring for protocol compliance.
- (2) Pre-trial simulation study of provider compliance with the trial protocol.
- (3) Three distinct study phases (infield training, run-in, and statistical inclusion)
- (4) Monitoring of the CPR process
- (5) Randomization at the subject level after the decision to resuscitate
- (6) Use of the Group Sequential Double Triangular Test with sufficient power to determine superiority, inferiority, or equivalence.

Clinical Paper

Manual vs. integrated automatic load-distributing band CPR with equal survival after out of hospital cardiac arrest. The randomized CIRC trial^{☆,☆☆}

	AutoPulse (n = 2099)	Manual (n = 2132)	P
Survival to discharge (%)	9.4	11.0	NS
24-hour survival (%)	21.8	25.0	NS
Compression fraction (%)	80.4	80.2	NS

Adjusted survival to discharge = OR 1.06 (95% CI 0.83 – 1.37)



Wik L. Resuscitation 2014;85:741-8

Results: Neurologic Endpoint

- No difference in mRS scores ≤ 3
 - Adjusted OR 0.80, 95% CI 0.47 - 1.37 (n.s.)

	M-CPR	iA-CPR
Discharge mRS	(n=233)	(n=196)
Score of 0 -3	48.1%	44.4%
Score of 4 -5	26.2%	25.5%
Unknown score	25.8%	30.1%



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LINC trial

A multicenter, randomized, controlled trial
designed to evaluate the efficacy and safety of:

LUCAS concept for resuscitation of OHCA
including defibrillation during ongoing
compressions

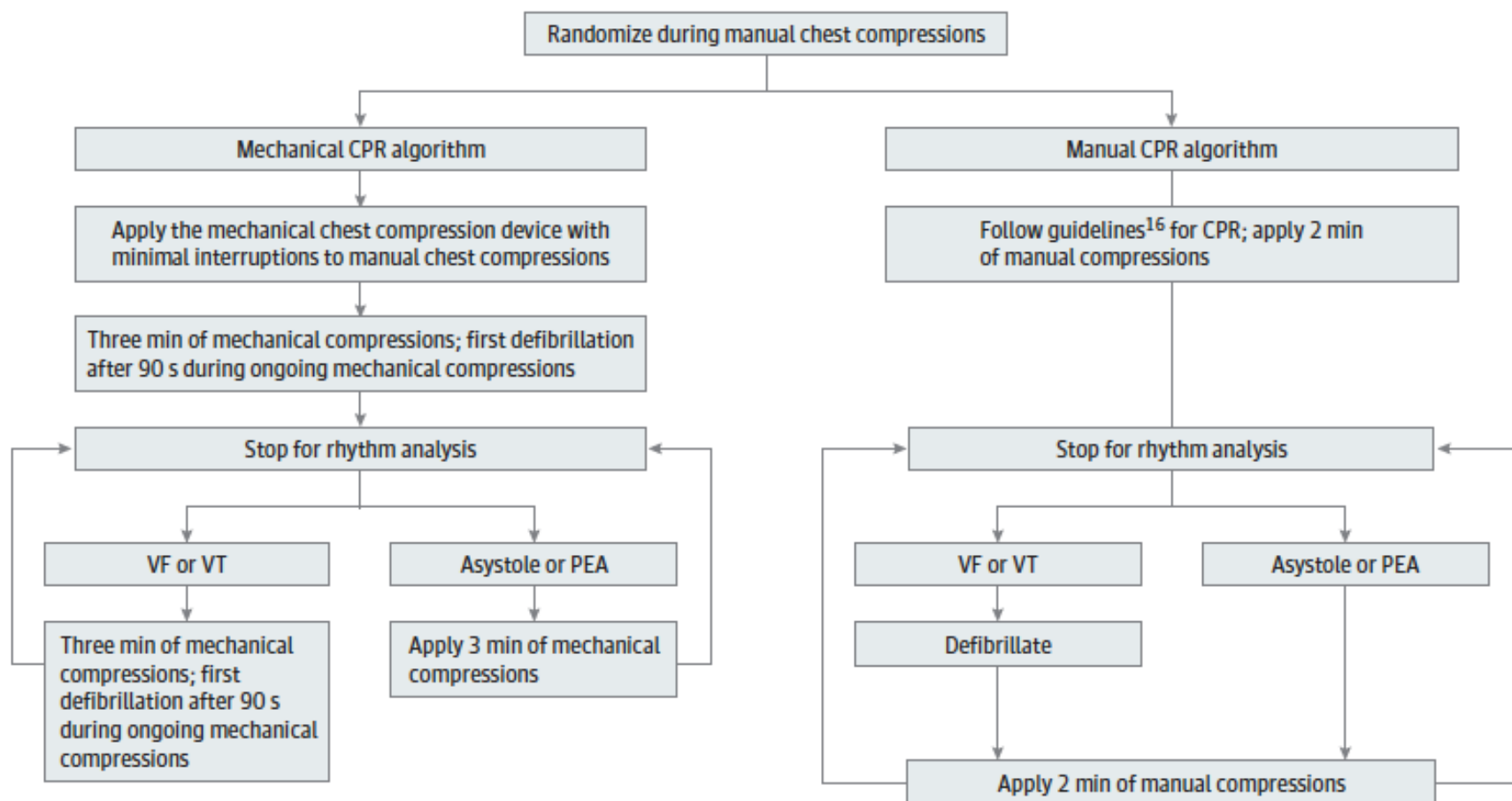
vs.

manual CPR according to 2005 guidelines

Rubertssen JAMA 2013

LINC study

LINC TRIAL

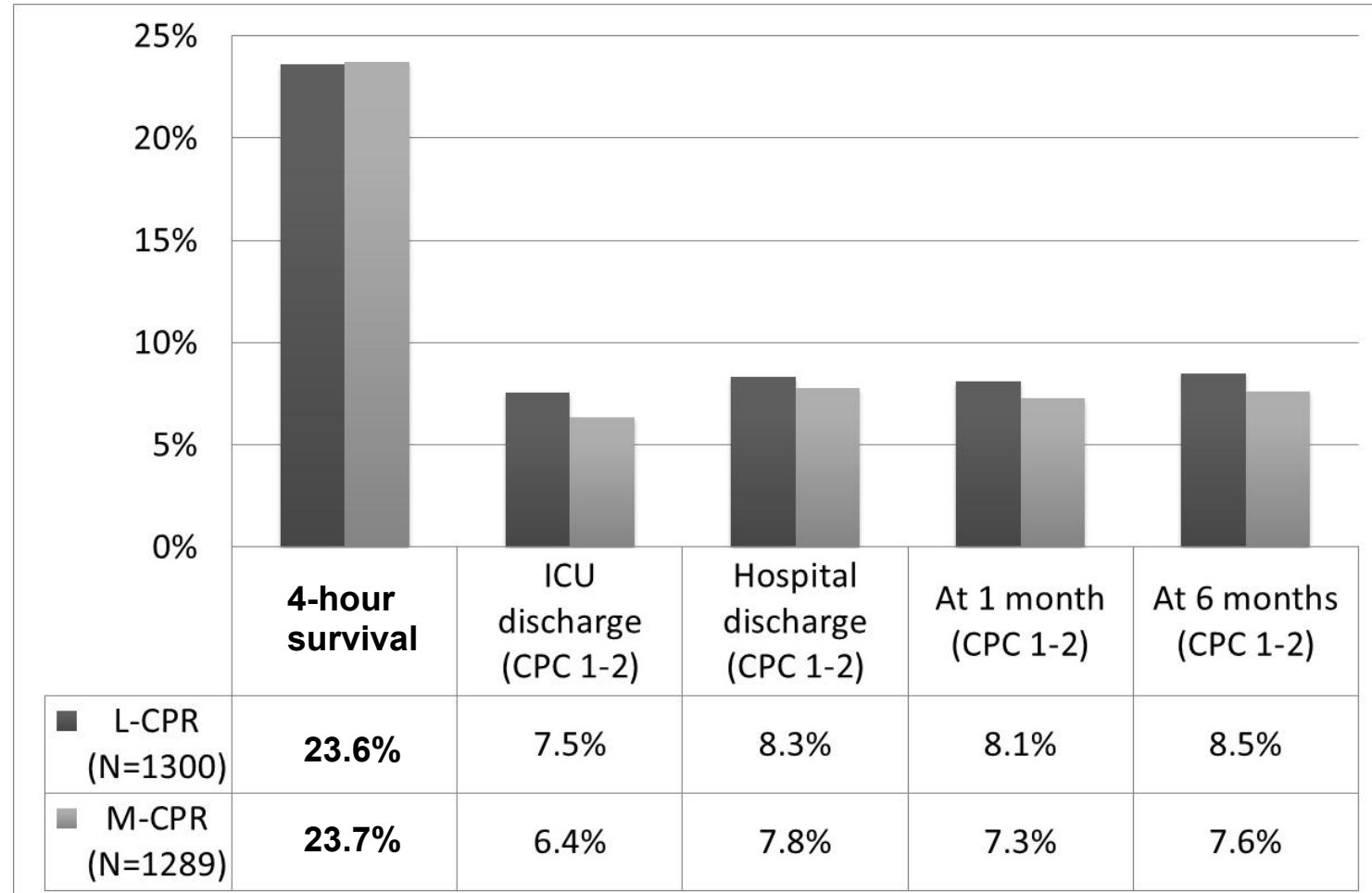




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4-hour survival:
Risk difference -0.05%
95% C.I. -3.32 – 3.23, p=1.00

Outcome





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Authors conclusion

In clinical practice, mechanical CPR using the presented algorithm did not result in improved effectiveness compared with manual CPR

LINC study



PARA♥EDIC



Pre-hospital Randomised Assessment of a Mechanical Compression Device In Cardiac Arrest (PARAMEDIC) Study



THE UNIVERSITY OF
WARWICK



Study design

Pragmatic, multi-centre, cluster randomised, clinical effectiveness trial

P opulation: Adults, non-traumatic OHCA

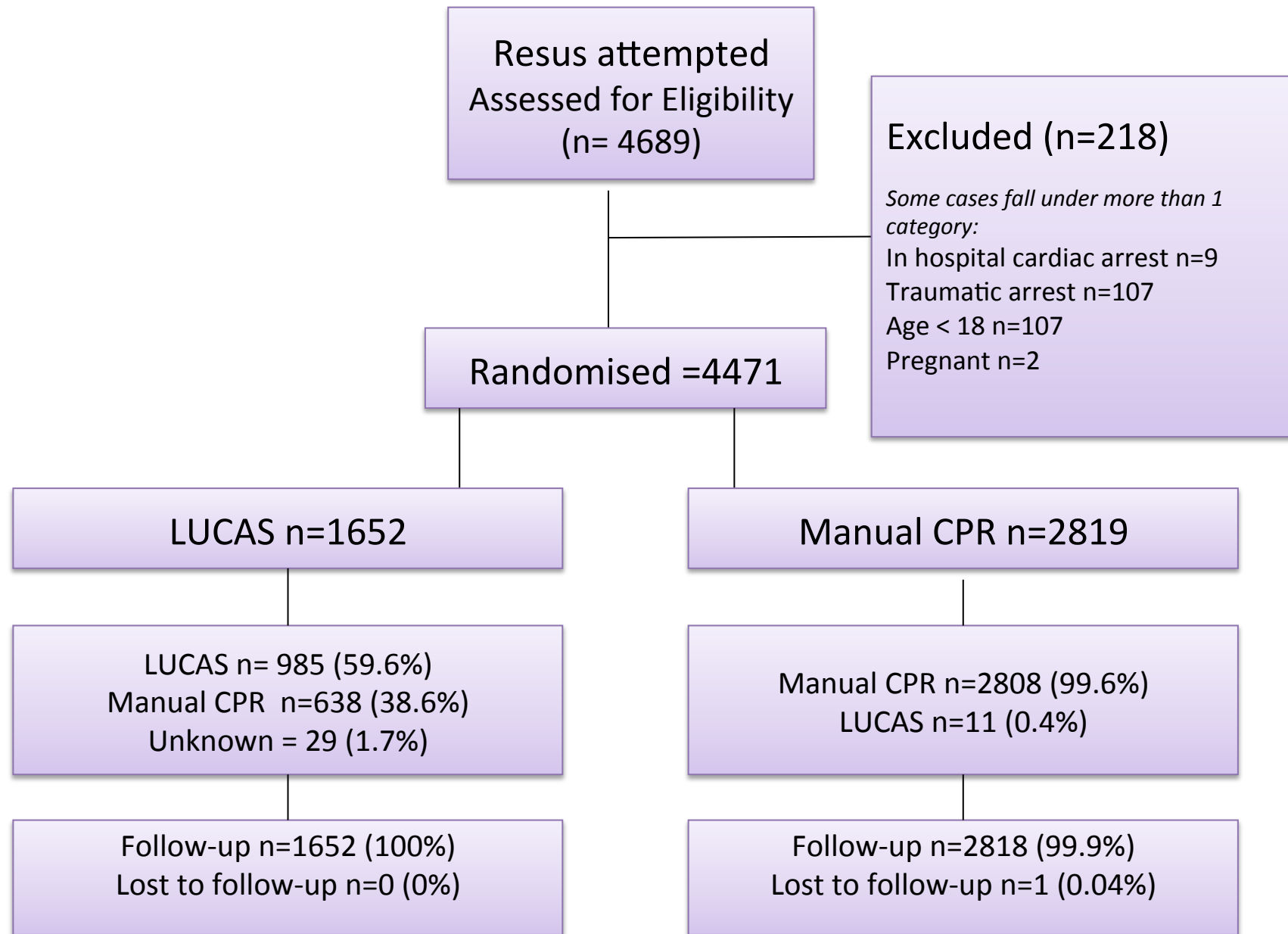
I ntervention: LUCAS CPR

C omparator: Standard CPR

O utcome: 30 day survival, clinical and cost effectiveness



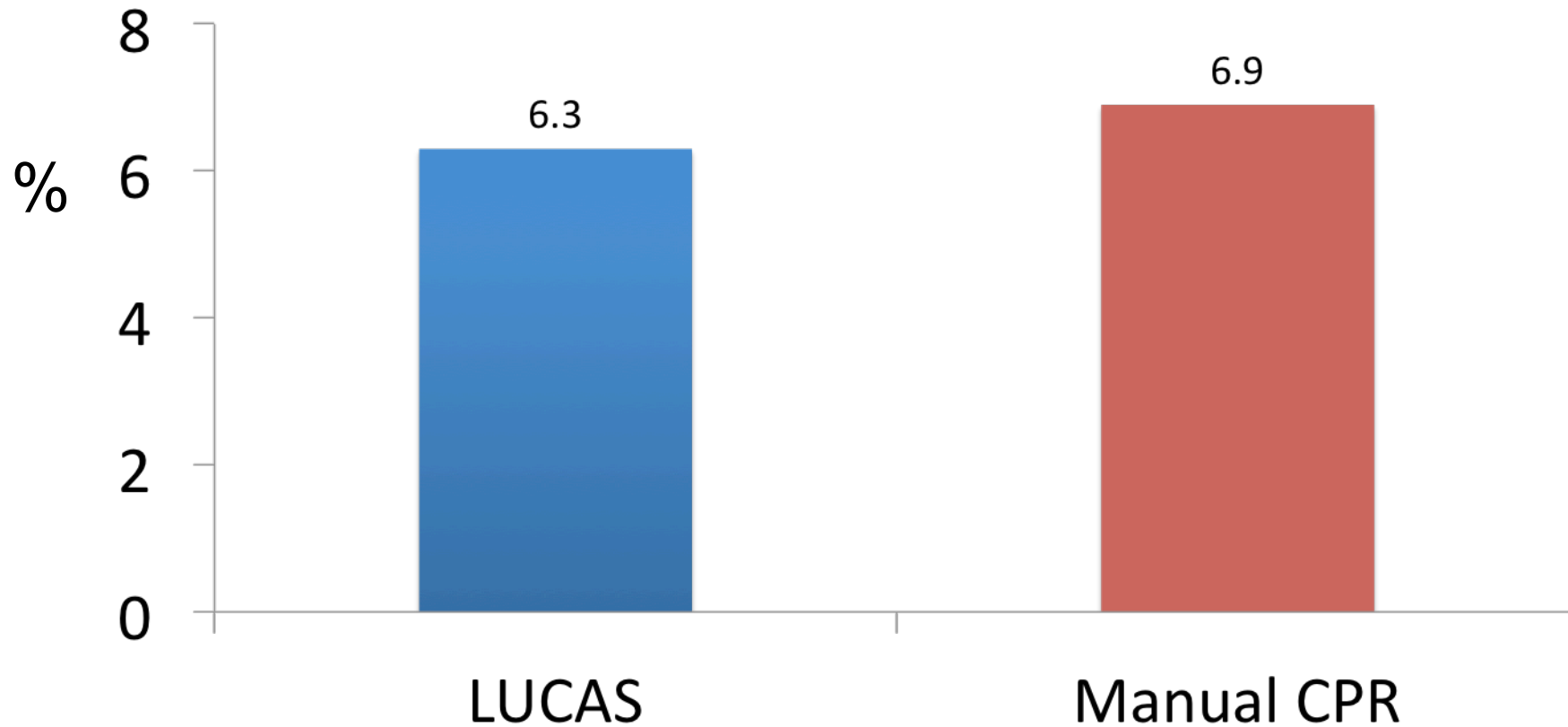
Protocol: Perkins GD *et al* Scand J Trauma Resusc Emerg Med. 2010



No difference in 30 day survival

Unadjusted odds ratio = 0.91 (95% CI: 0.71, 1.17) P=0.473

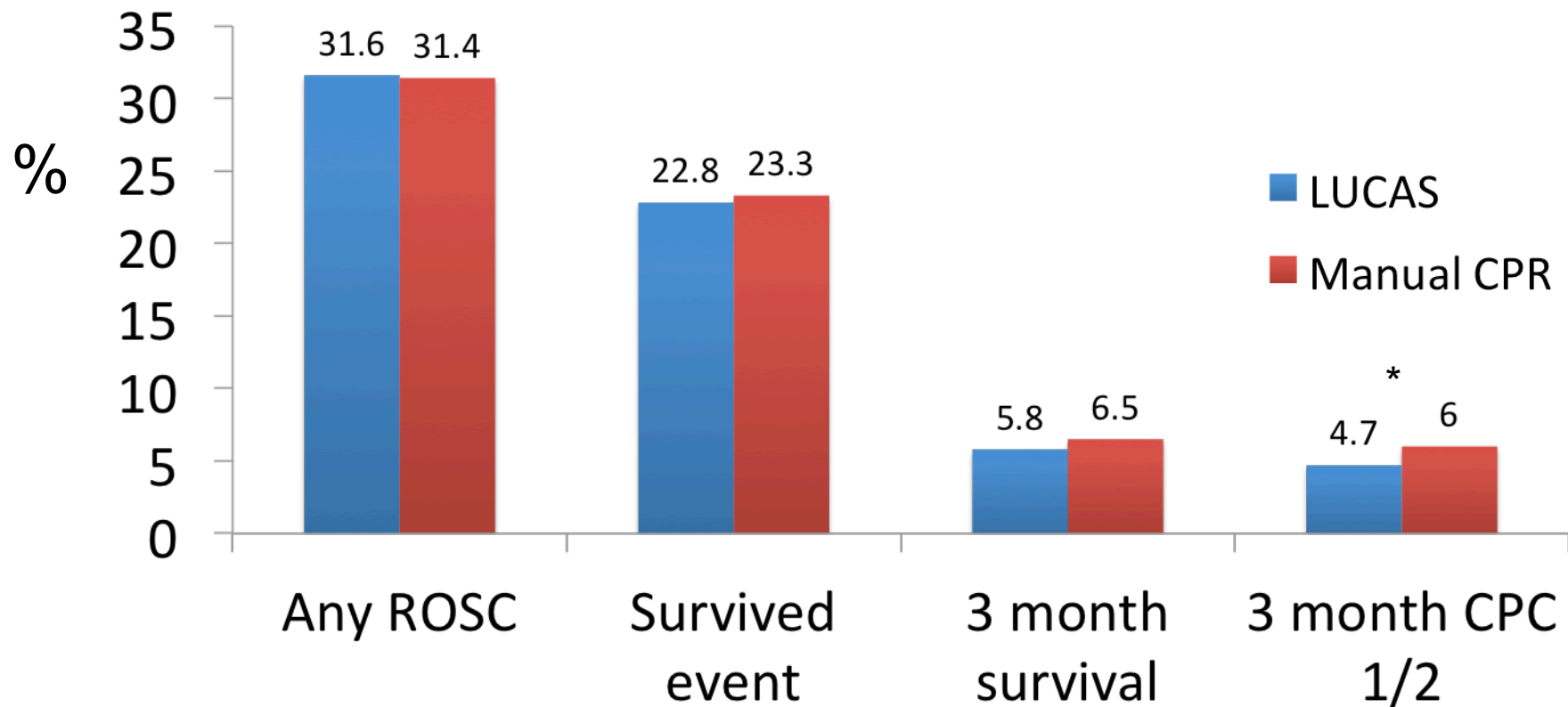
Adjusted odds ratio = 0.86 (95% CI: 0.64, 1.15) P=0.31



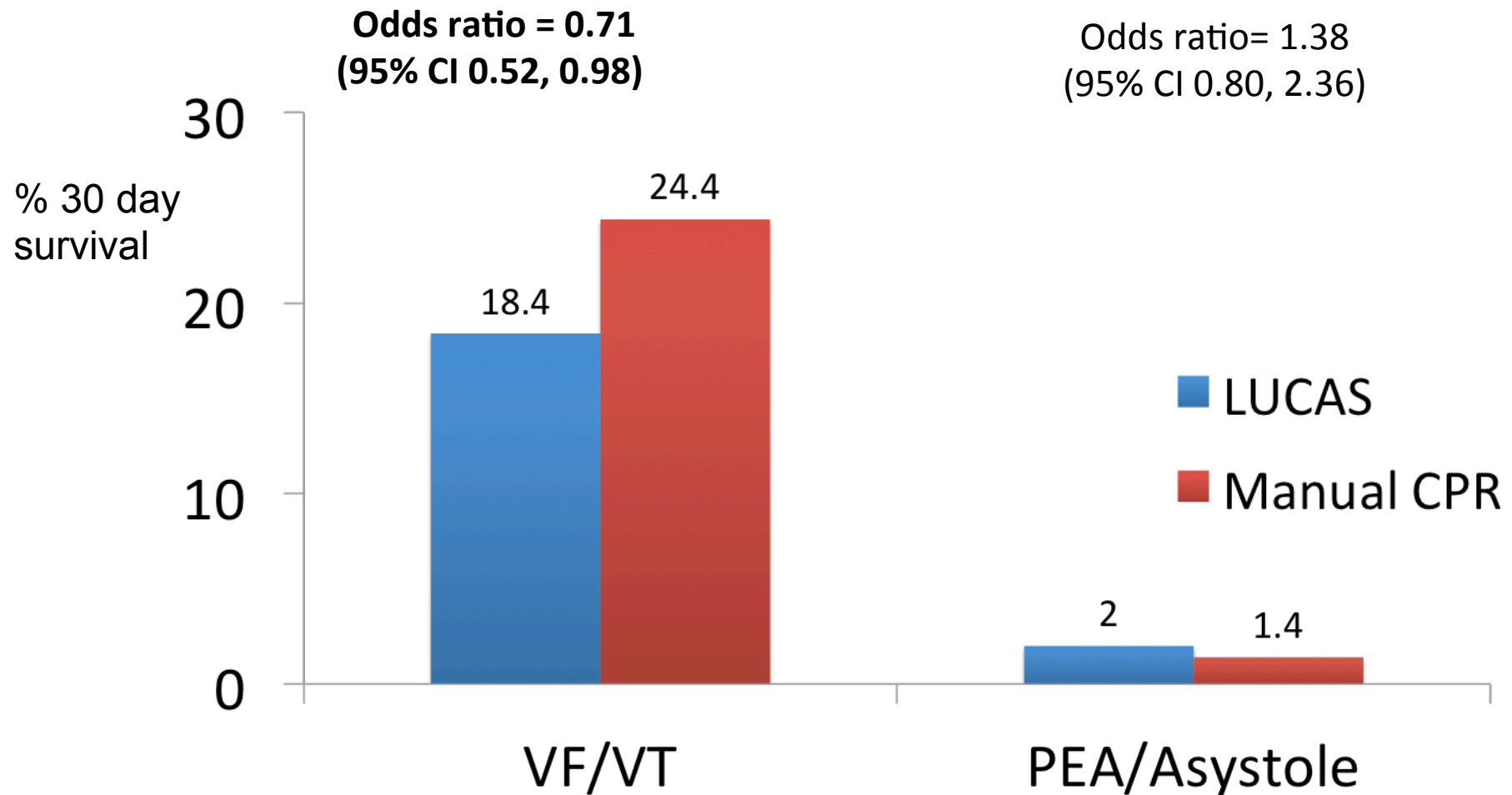
Secondary outcomes

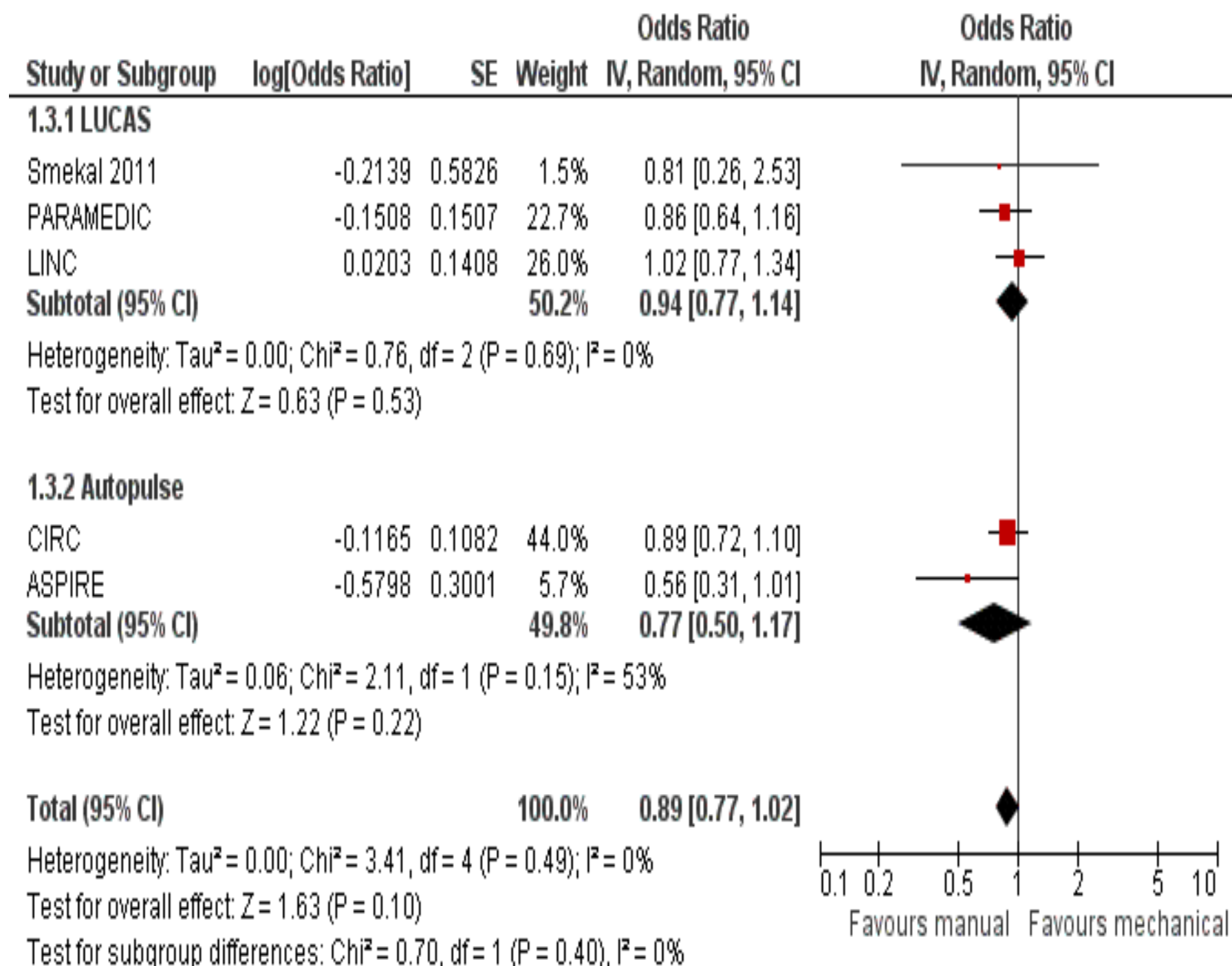
CPC Unadjusted odds ratio = 0.77 (95% CI = 0.59, 1.02) P=0.0667

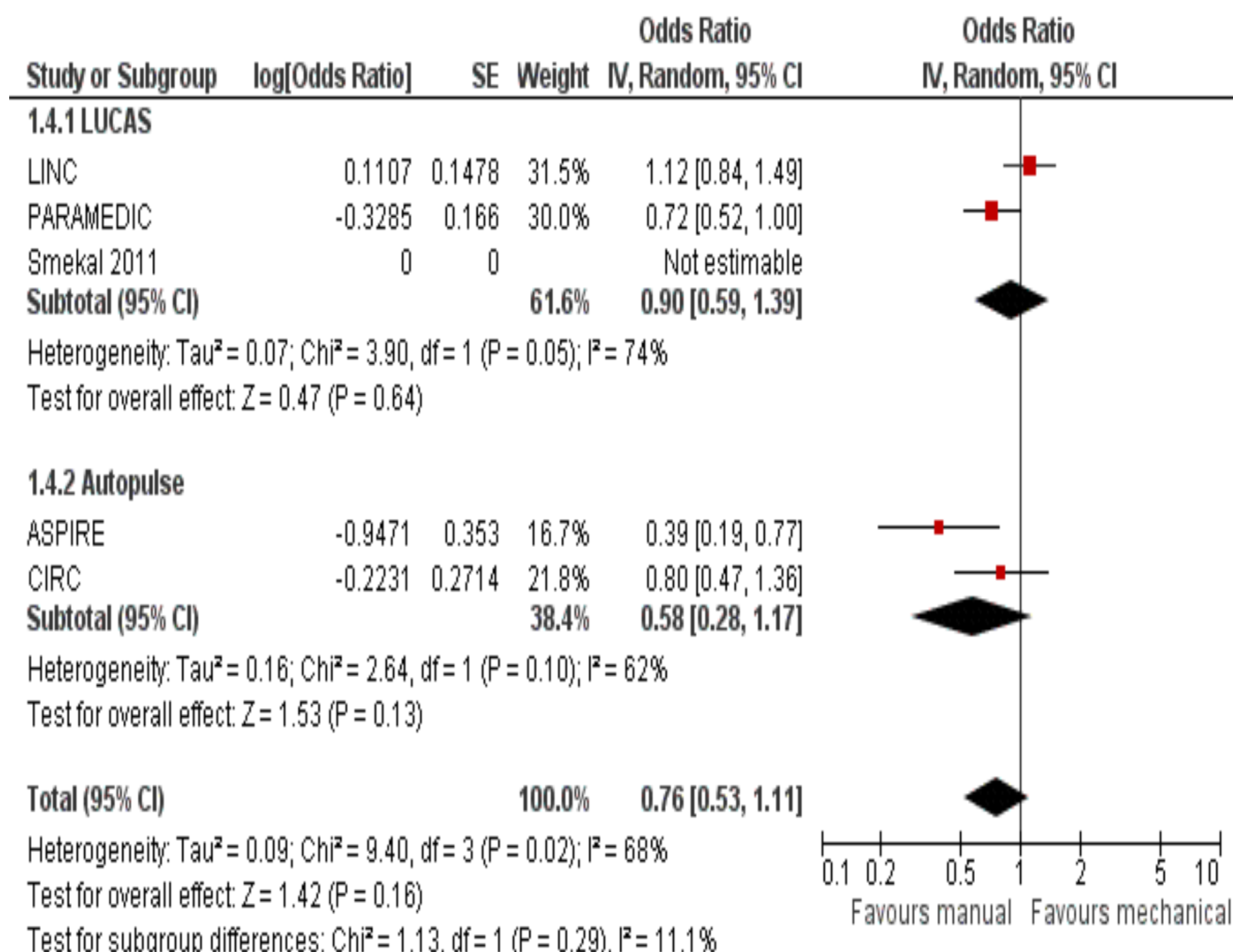
CPC Adjusted odds ratio= 0.72 (95% CI = 0.52, 0.99) P=0.0459



Sub-group analysis (rhythm)







A role for mechanical CPR devices?



- Routine use not recommended
- Consider if manual CPR impractical
 - Prolonged CPR
 - EMS transport
 - PCI
 - Bridge to other interventions (e-CPR, organ donation)

Adrenaline for cardiac arrest

AN EXPERIMENTAL RESEARCH INTO THE RESUSCITATION OF DOGS KILLED BY ANESTHETICS AND ASPHYXIA.

By GEORGE CRILE, M.D., AND DAVID H. DOLLEY, M.D.

(From the Laboratory of Surgical Physiology, Western Reserve Medical School, Cleveland.)

PLATES XLII-XLIX.

In a previous communication by one of us (Crile) resuscitation was attempted by means of both direct and indirect cardiac massage, with and without artificial respiration, with and without intravenous saline infusion, and with and without the addition of adrenalin. The results of these experiments may be summarized as follows: By cardiac massage alone animals were rarely resuscitated at any time after quiescence of the circulation and respiration; by combining either direct or indirect cardiac massage with artificial respiration and the head-down posture a certain percentage of the animals were recovered after the lapse of from one to three minutes. The results, however, were quite uncertain, and in the case of death from chloroform recovery was the exception. By adding to cardiac massage artificial respiration and intravenous saline infusion, resuscitations were in a slightly greater proportion successful. The same procedures, with the addition of adrenalin to the intravenous saline infusion, were markedly more successful in the deaths from asphyxia through rarely so in the cases of death from chloroform. It required in almost every instance a vigorous compression of the thorax over the heart for a considerable time, varying from five to ten minutes, before resuscitation could be accomplished. Even then there were a considerable number of failures. In the majority of these failures autopsy showed

HEART - LUNG RESUSCITATION


I FIRST AID: OXYGENATE THE BRAIN IMMEDIATELY

IF UNCONSCIOUS
Airway - TILT HEAD BACK


IF NOT BREATHING
Breathe - INFLATE LUNGS 3-5 TIMES, MAINTAIN HEAD TILT
MOUTH-TO-MOUTH, MOUTH-TO-NOSE, mouth-to-adjunct, bag-mask

o FEEL PULSE
o IF PRESENT - CONTINUE LUNG INFLATIONS
o IF ABSENT -

Circulate - COMPRESS HEART ONCE A SECOND.
ALTERNATE 2-3 LUNG INFLATIONS WITH 15 STERNAL COMPRESSIONS UNTIL SPONTANEOUS PULSE RETURNS.



Direct lower sternum 1-2 inches



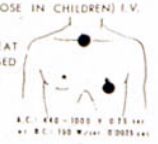
1 or 2 operators

II START SPONTANEOUS CIRCULATION

Drugs - EPINEPHRINE: 1.0mg (10 CC OF 1:1000) I.V. OR 0.5mg INTRACARDIAC. REPEAT LARGER DOSE IF NECESSARY.
SODIUM BICARBONATE: APPROXIMATELY 3.75 G/50 CC (1/2 DOSE IN CHILDREN) I.V. REPEAT EVERY 5 MINUTES IF NECESSARY.

E. K. G. - • FIBRILLATION: EXTERNAL ELECTRIC DEFIBRILLATION REPEAT SHOCK EVERY 1-3 MINUTES UNTIL FIBRILLATION REVERSED
• IF ASYSTOLE OR WEAK BEATS: EPINEPHRINE OR CALCIUM I.V.

Fluids - I.V. PLASMA, DEXTRAN, SALINE
Do not interrupt cardiac compressions and ventilation.
Tracheal intubation only when necessary.
AFTER RETURN OF SPONTANEOUS CIRCULATION USE VASOPRESSORS AS NEEDED.
e.g. NOREPINEPHRINE (Levophed) I.V. DRIP



A.C. 440-1000 x 0.25 sec
M.C. 150-200 x 0.005 sec

III SUPPORT RECOVERY

(physician-specialist)

Gauge EVALUATE AND TREAT CAUSE OF ARREST

Hypothermia START WITHIN 30 MINUTES IF NO SIGN OF CNS RECOVERY

Intensive Care SUPPORT VENTILATION: TRACHEOTOMY, PROLONGED CONTROLLED VENTILATION, GASTRIC TUBE AS NECESSARY
SUPPORT CIRCULATION
CONTROL CONVULSIONS
MONITOR

Figure 1. Heart-lung resuscitation (cardiopulmonary-cerebral resuscitation). First composition in 1961, Pittsburgh, PA. Reproduced with permission from Safar P. Community-wide CPR. J Iowa Medical Society 1964 (Nov); pp 629-635.



Clinical Paper

Wide variability in drug use in out-of-hospital cardiac arrest resuscitation outcomes conference

Benedict M. Glover^a, Siobhan M. O'Connell^b, Lois Van Ottingham^b, Christopher J. Cook^c, the Resuscitation Outcomes Consortium Investigators^d

^a University of Toronto, Toronto, Canada
^b University of Washington, Seattle, WA, USA
^c University of California, San Francisco, CA, USA
^d University of California, San Francisco, CA, USA

Peter J. Kudenchuk^e, Anne L. Atkins^{e,f}

Summary

The available clinical data confirm that epinephrine administration during CPR can increase short-term survival (return of pulses), but point towards either no benefit or even harm of this drug for more patient-centred outcomes (long-term survival or functional recovery). Prospective trials are needed to determine the correct dose, timing and patients for epinephrine in cardiac arrest.

Conclusions: Epinephrine use during cardiac arrest is not associated with improved survival to hospital discharge. Observational studies with a lower-risk for bias suggest that it may be associated with decreased survival.

Neither lidocaine nor amiodarone was associated with a survival benefit while there was an inverse relationship between the administration of epinephrine, atropine and sodium bicarbonate and survival to hospital discharge.

Conclusions: There is considerable variability among Emergency Medical Services agencies in their use of pharmacological therapy for out-of-hospital cardiac arrests which may be resolved by performing large randomised controlled trials examining effects on survival.

With this important addition to the “adrenaline in cardiac arrest” puzzle, the resuscitation community eagerly awaits new randomised controlled trials to provide more definitive answers

a treatment for out

Line¹, Peter Cottrell medical student², Simon Gates

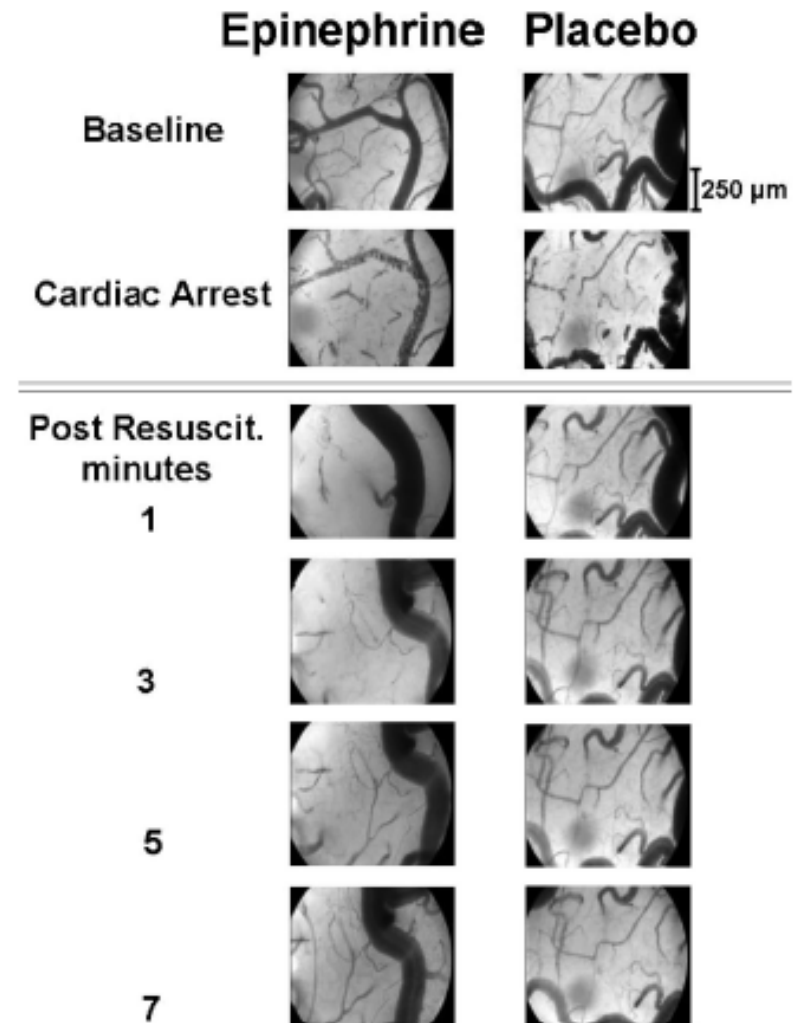


Epinephrine reduces cerebral perfusion during cardiopulmonary resuscitation*

Crit Care Med 2009;37:1408-15

Giuseppe Ristagno, MD; Wanchun Tang, MD, FCCM; Lei Huang, MD; Alain Fymat, MD; Yun-Te Chang, MD; Shijie Sun, MD, FCCM; Carlos Castillo, MSEE; Max Harry Weil, MD, PhD, FCCM

- Pig study
- Cerebral oxygen tension (PbO_2) and microcirculatory imaging
- Adrenaline increased arterial pressure but reduced PbO_2 and microcirculatory flow.

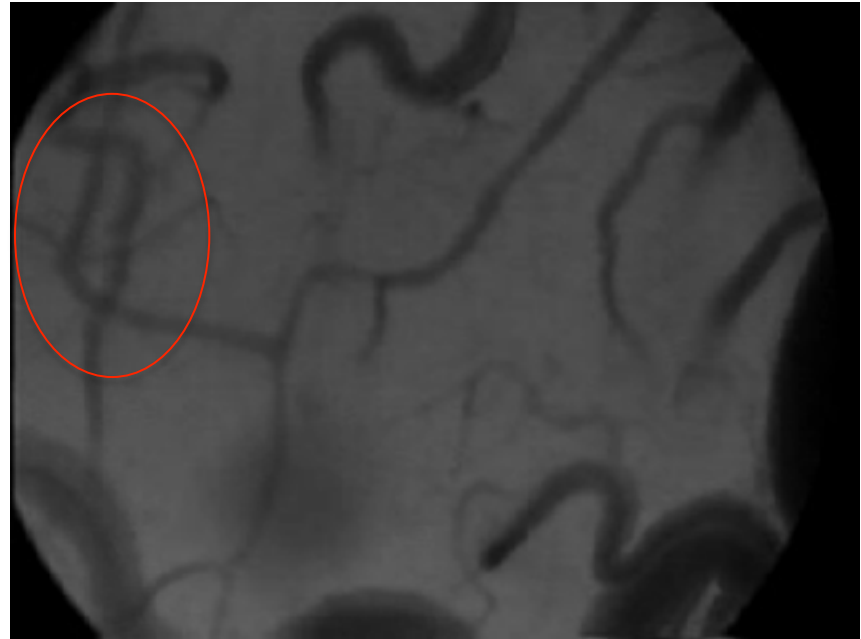


Blood flow 3 min after ROSC

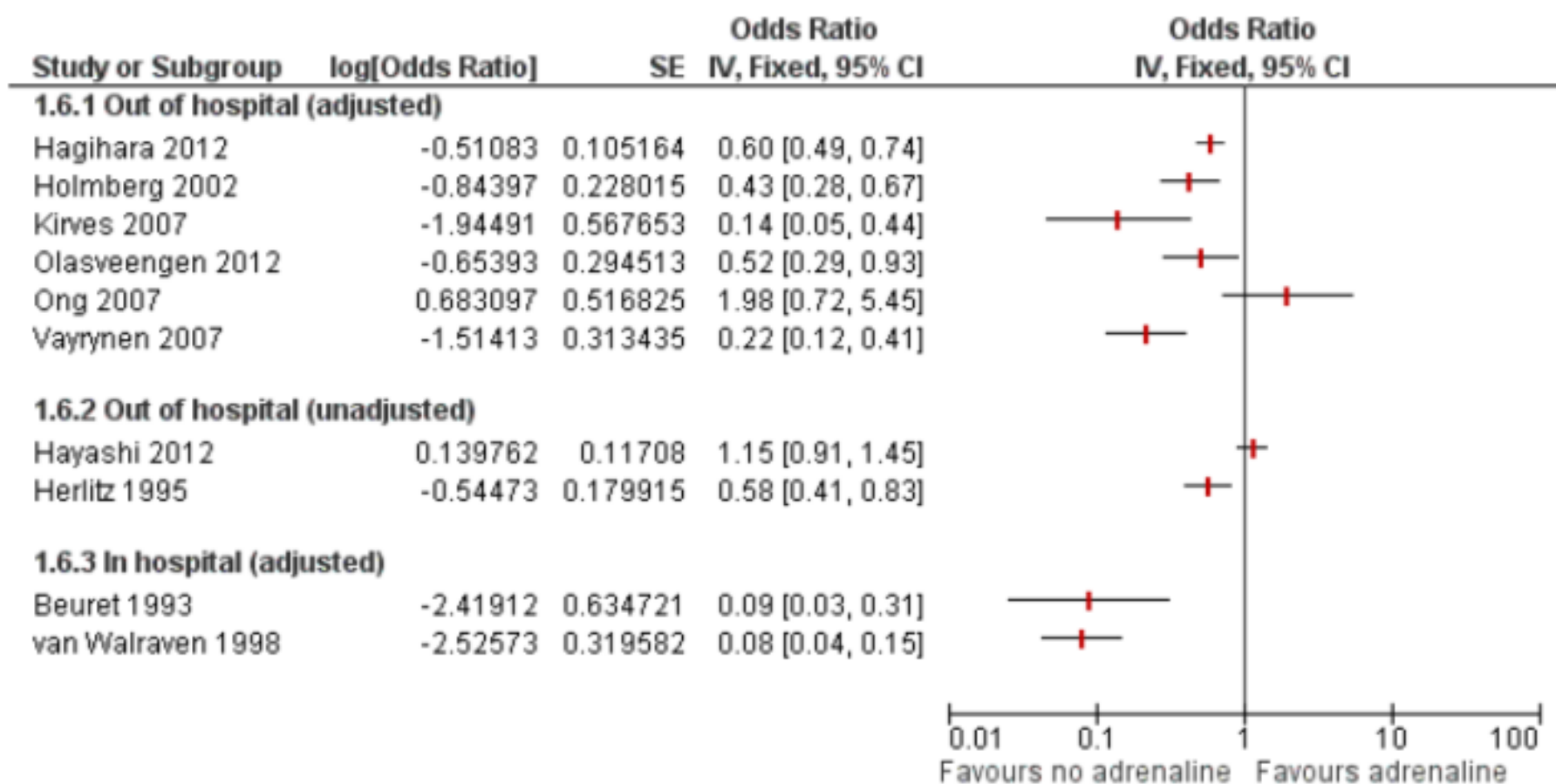
Adrenaline arm



Saline (placebo) arm



Films courtesy of Giuseppe Ristagno,



Prehospital Epinephrine Use and Survival Among Patients With Out-of-Hospital Cardiac Arrest

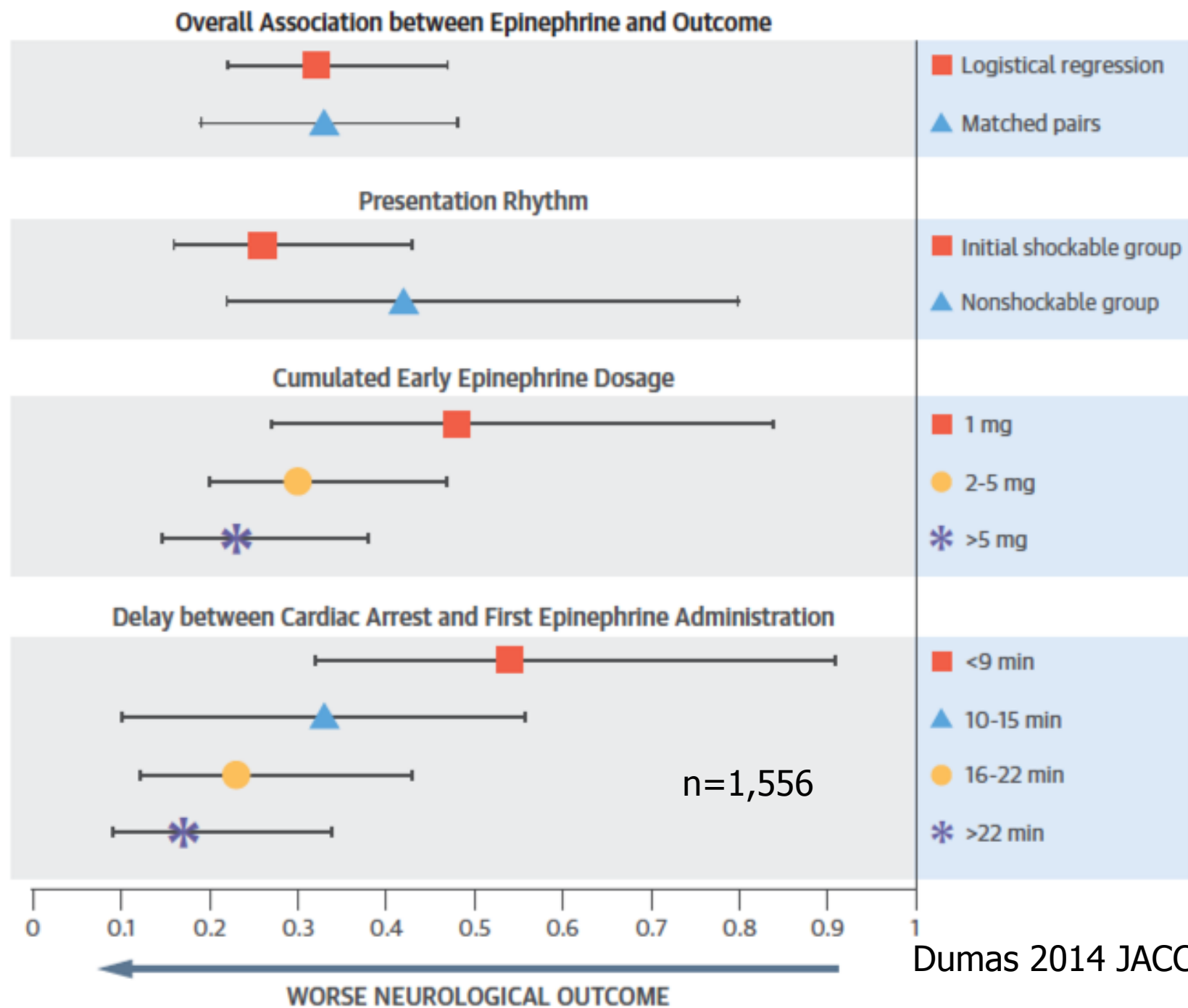
Hagihara A. JAMA 2012;307:1161-8

Whole of Japan - 2005 -2008

**417 188 cases analysed; 15 030 received epinephrine;
13 401 propensity-matched pairs**

Table 4. Conditional Logistic Regression Analyses of Outcome in Epinephrine Group (vs No-Epinephrine Group) Among Propensity-Matched Patients With Out-of-Hospital Cardiac Arrest (n = 26 802)

Analysis	Odds Ratio (95% CI) ^a			
	ROSC	1-Month Survival	CPC 1 or 2	OPC 1 or 2
Unadjusted	1.91 (1.78-2.05)	0.71 (0.64-0.79)	0.41 (0.34-0.49)	0.43 (0.36-0.51)
Adjusted for propensity	2.01 (1.83-2.21)	0.71 (0.62-0.81)	0.41 (0.33-0.52)	0.43 (0.34-0.54)
Adjusted for propensity and selected variables ^b	2.24 (2.03-2.48)	0.60 (0.49-0.74)	0.40 (0.26-0.63)	0.43 (0.28-0.66)
Adjusted for propensity and all covariates ^c	2.51 (2.24-2.80)	0.54 (0.43-0.68)	0.21 (0.10-0.44)	0.23 (0.11-0.45)



Prehospital Assessment of the Role of Adrenaline: Measuring the Effectiveness of Drug administration In Cardiac arrest

- Placebo versus adrenaline in out-of-hospital cardiac arrest
- Primary end point 30-day survival
- Target sample size = 8000
- Pilot started Dec 2014



Population

Intervention

Outcome

Persistent or
recurrent VF/VT*
after ≥ 1 shock

Vascular Access

RANDOMIZE

Amiodarone

Lidocaine

Neither
(saline placebo)

SURVIVAL TO
HOSPITAL DISCHARGE

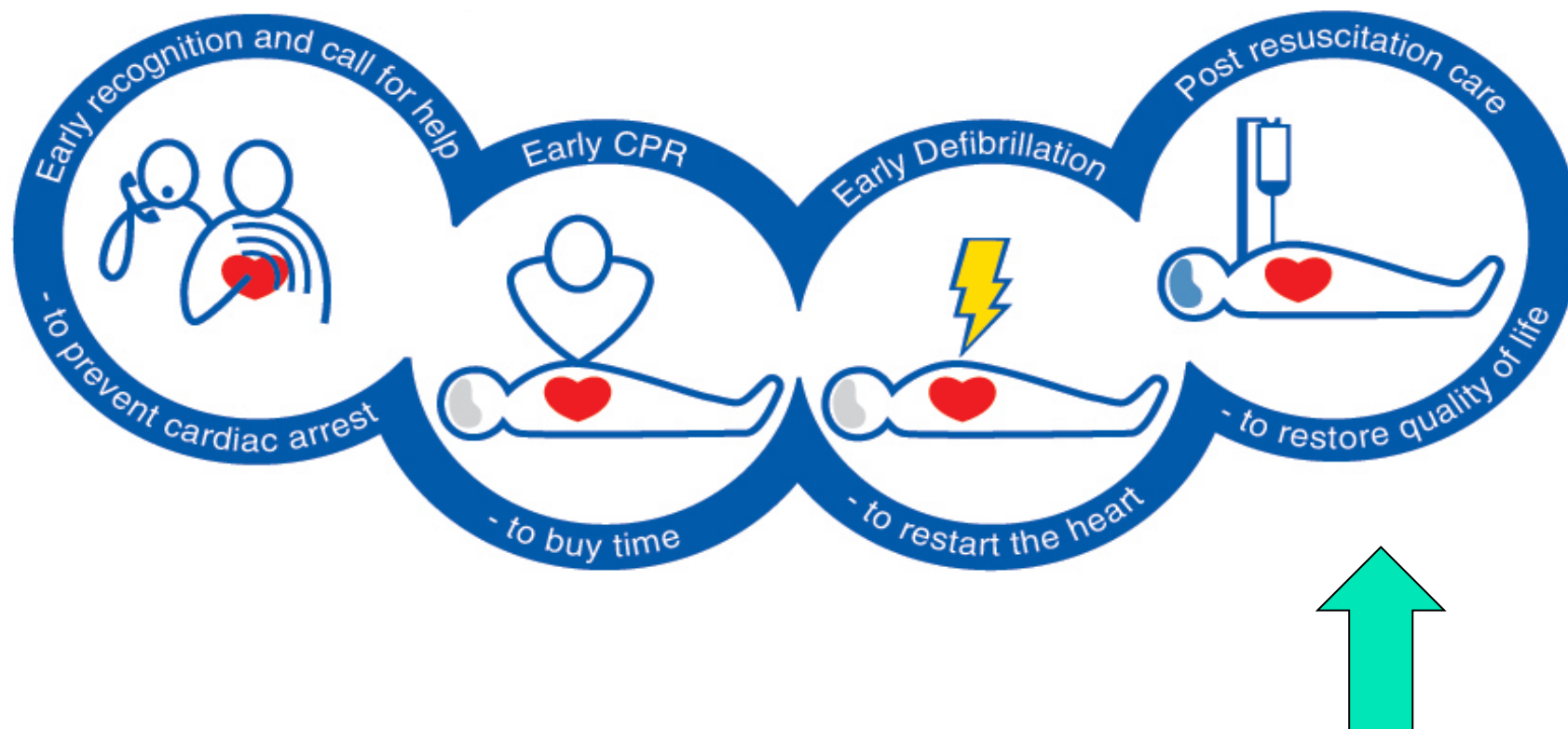
ROC ALPS

Amiodarone, Lidocaine, or Placebo Study

Version 24: Rev. 2012-4-24

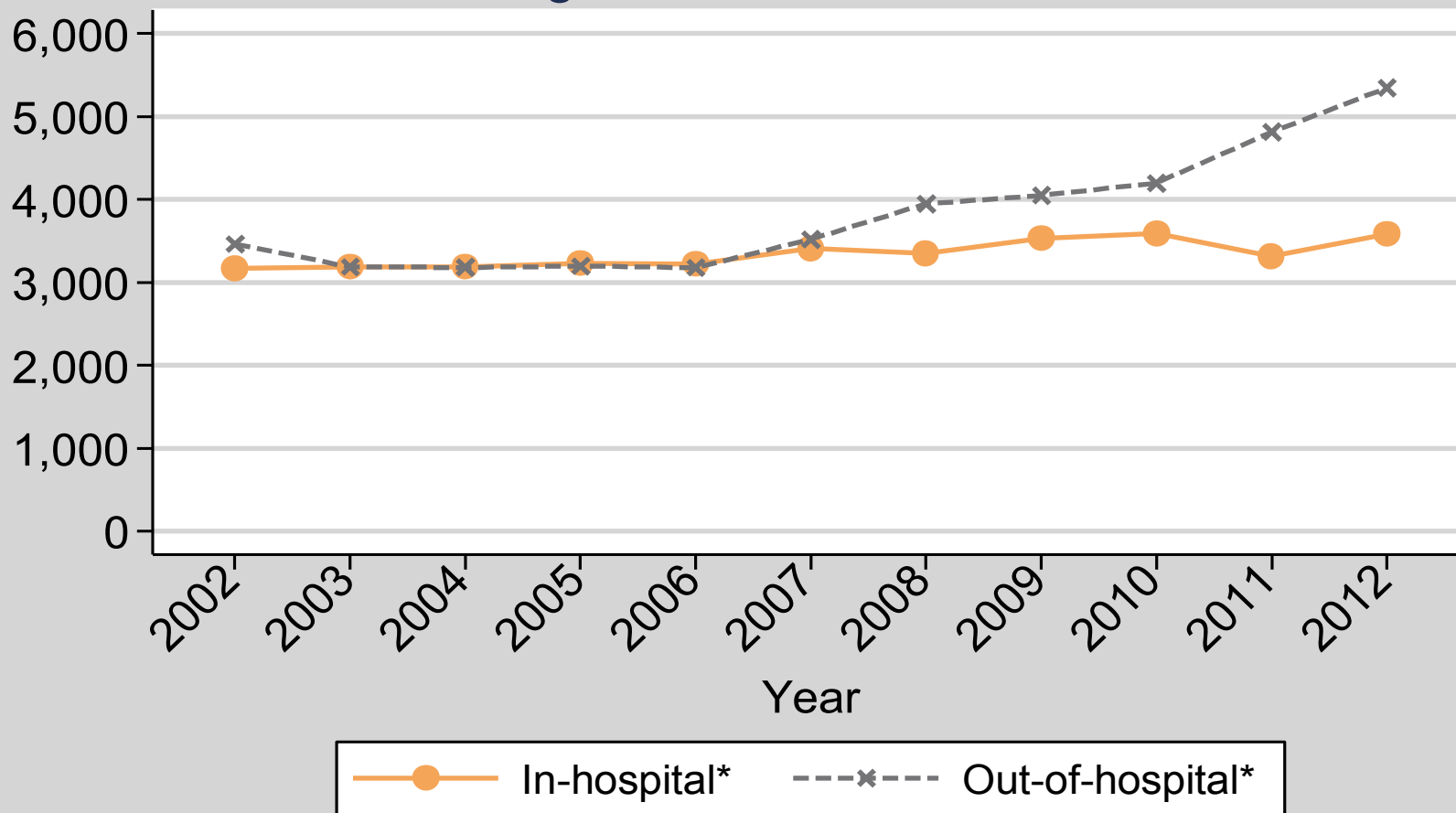
RESUSCITATION OUTCOMES CONSORTIUM

Chain of survival



Admissions following cardiac arrest

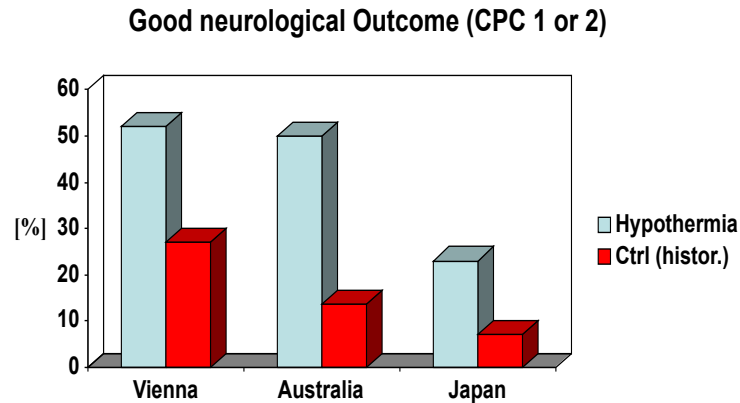
Extrapolated number of admissions to critical care following cardiac arrest in England, Wales and Northern Ireland



* location of cardiac arrest is approximated using location immediately prior to critical care admission

© ICNARC 2013

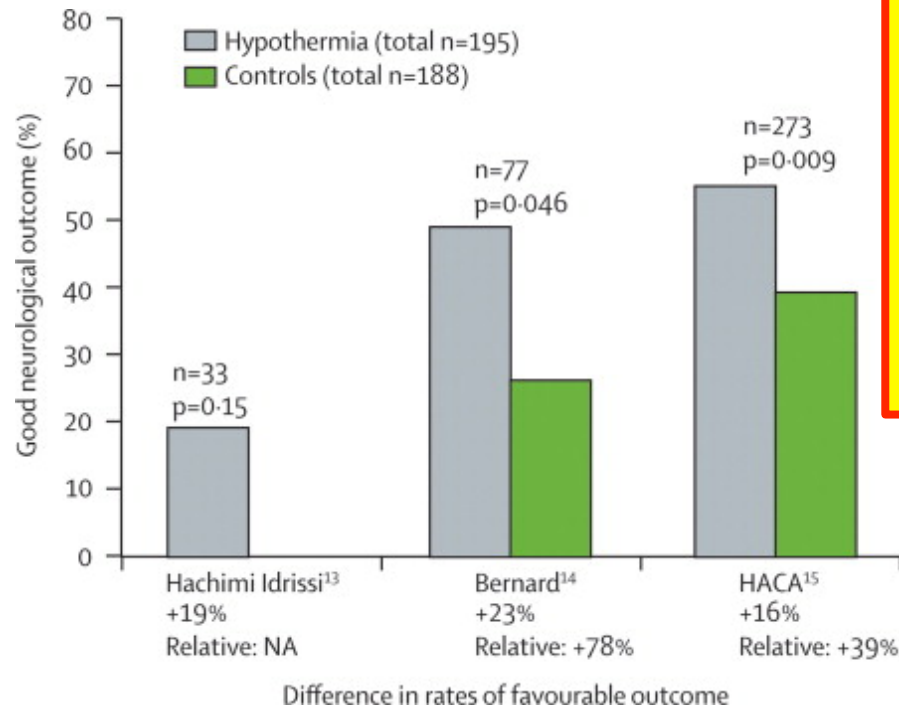
TH - clinical studies



Zeiner et al. *Stroke* 2000;31:86-94

Bernard et al. *Ann Emerg Med* 1997;30:146-153

Yanagawa et al. *Resuscitation* 1998;39:61-66



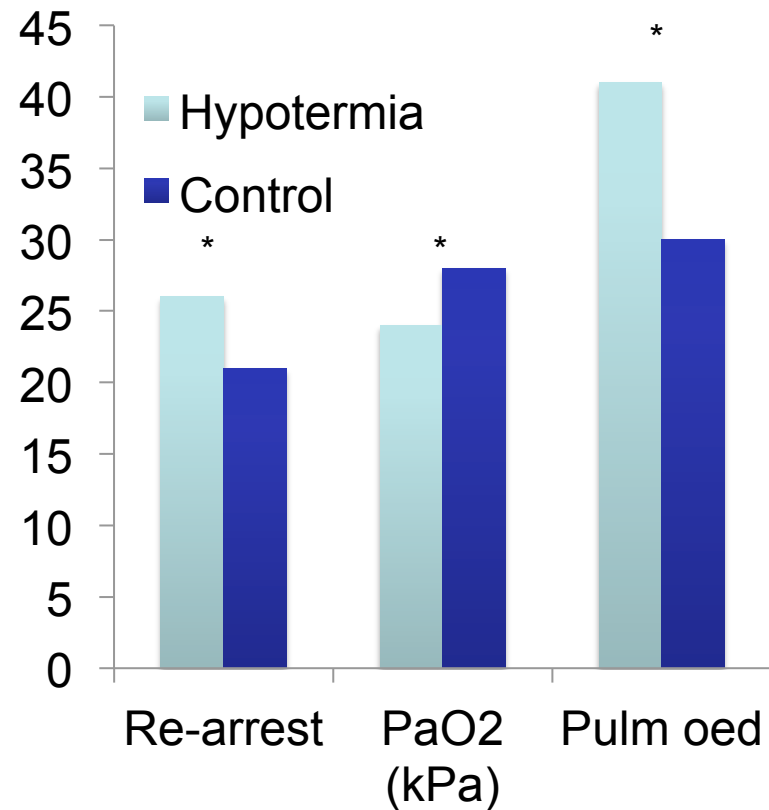
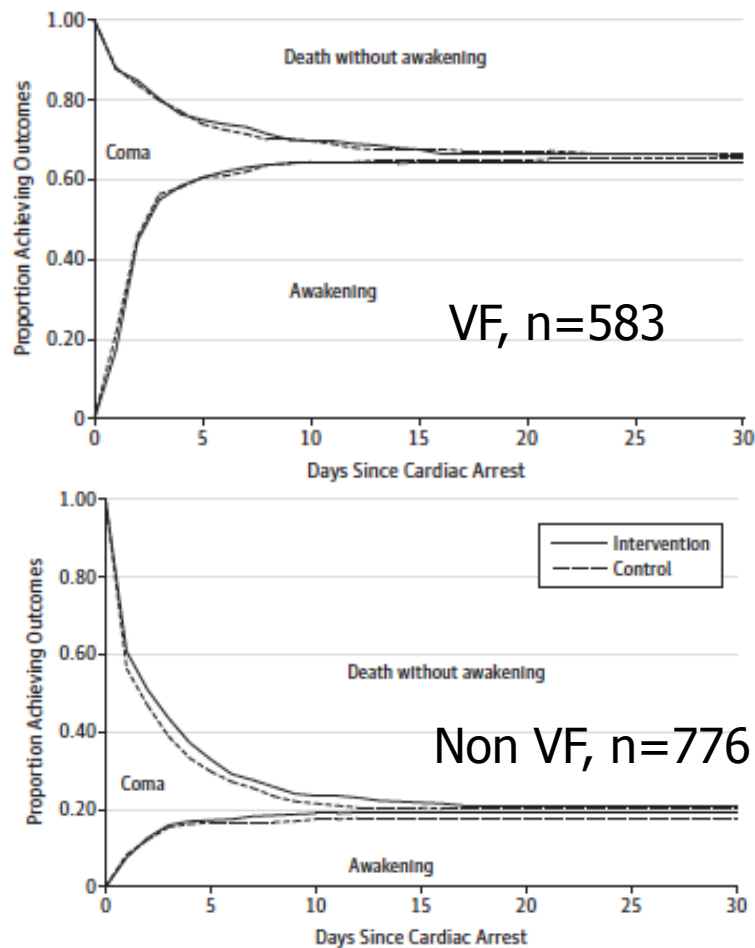
ILCOR Advisory Statement (2003)

Unconscious adult patients with spontaneous circulation after out-of-hospital cardiac arrest ***should be cooled*** to 32-34°C for 12-24 hrs when the initial rhythm was VF.

For any other rhythm, or cardiac arrest inhospital, such cooling may also be beneficial.

Original Investigation

Effect of Prehospital Induction of Mild Hypothermia on Survival and Neurological Status Among Adults With Cardiac Arrest A Randomized Clinical Trial



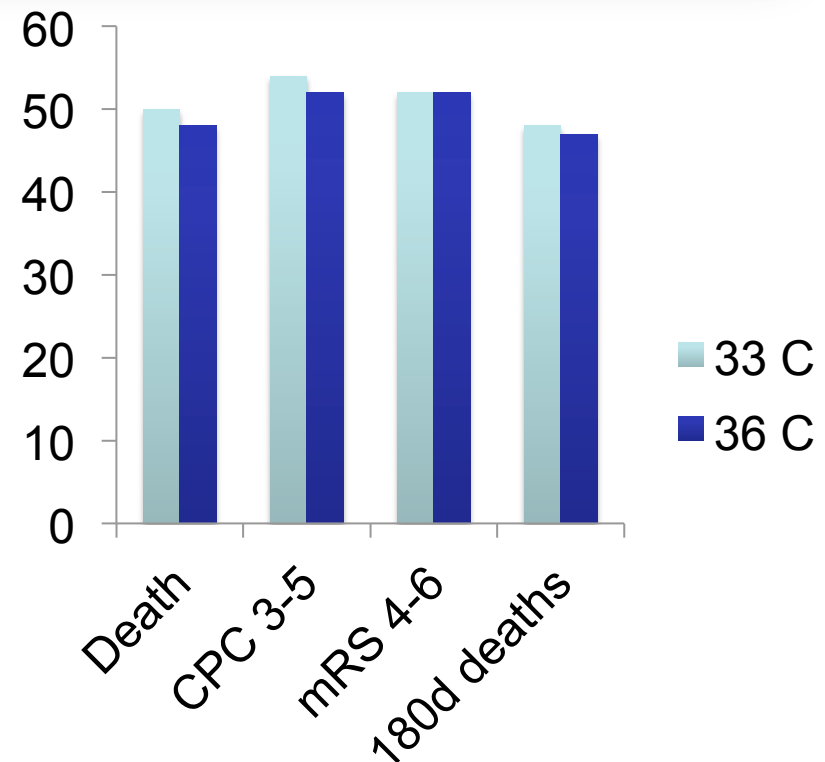
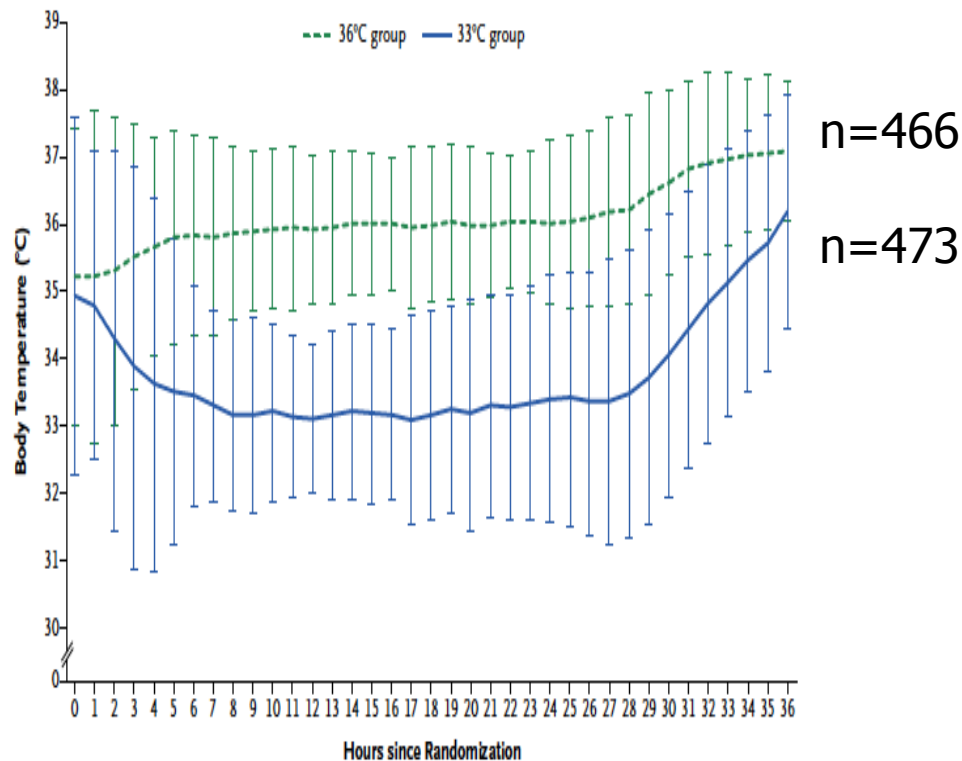
2L Ice cold saline

* P < 0.05

ORIGINAL ARTICLE

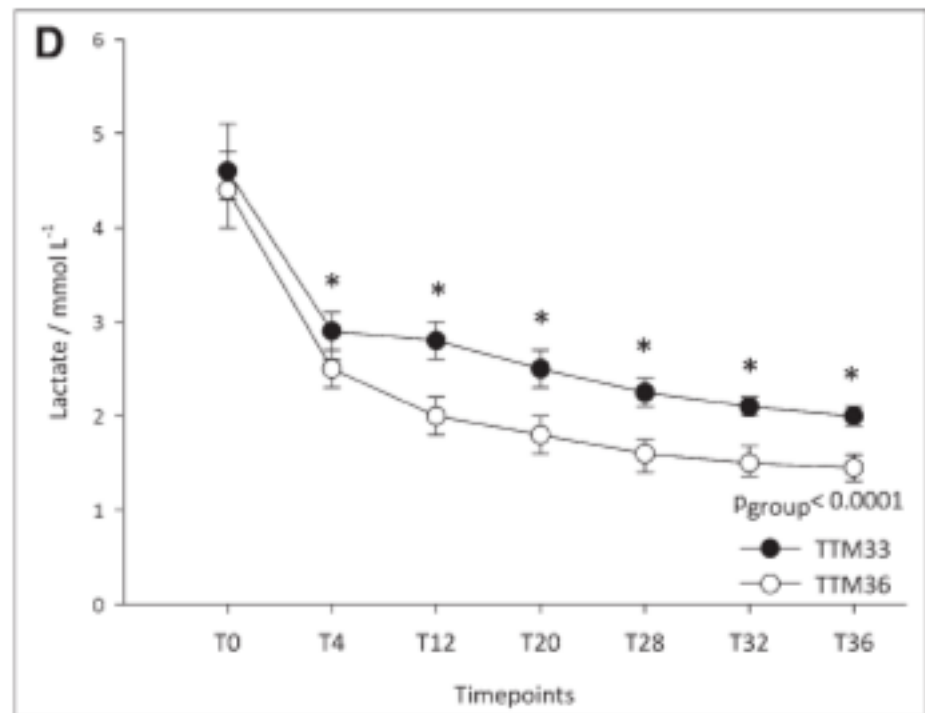
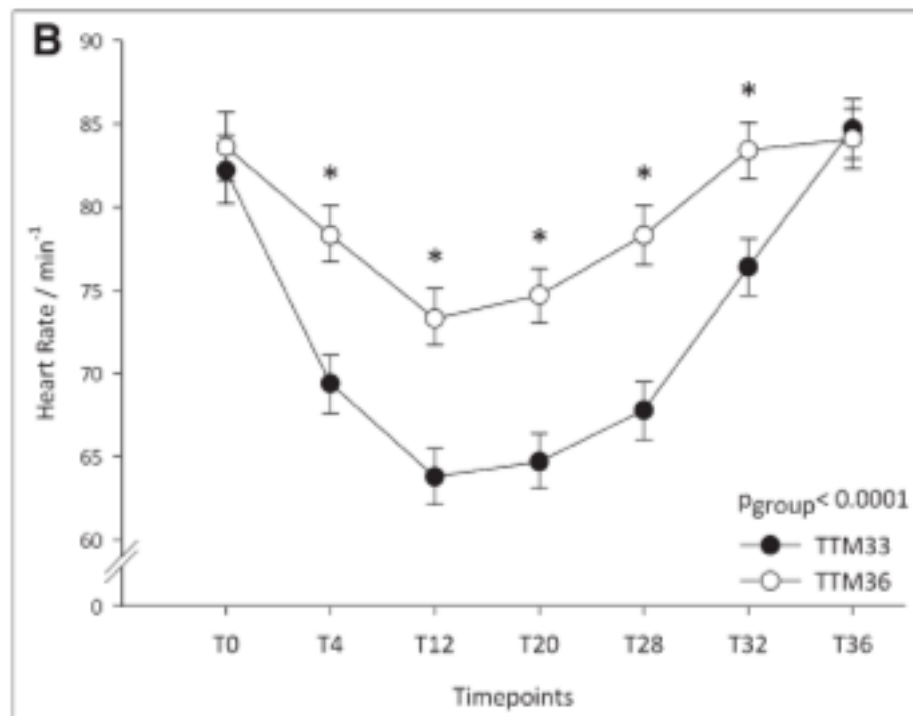
Targeted Temperature Management at 33°C versus 36°C after Cardiac Arrest

Niklas Nielsen, M.D., Ph.D., Jørn Wetterslev, M.D., Ph.D., Tobias Cronberg, M.D., Ph.D.,

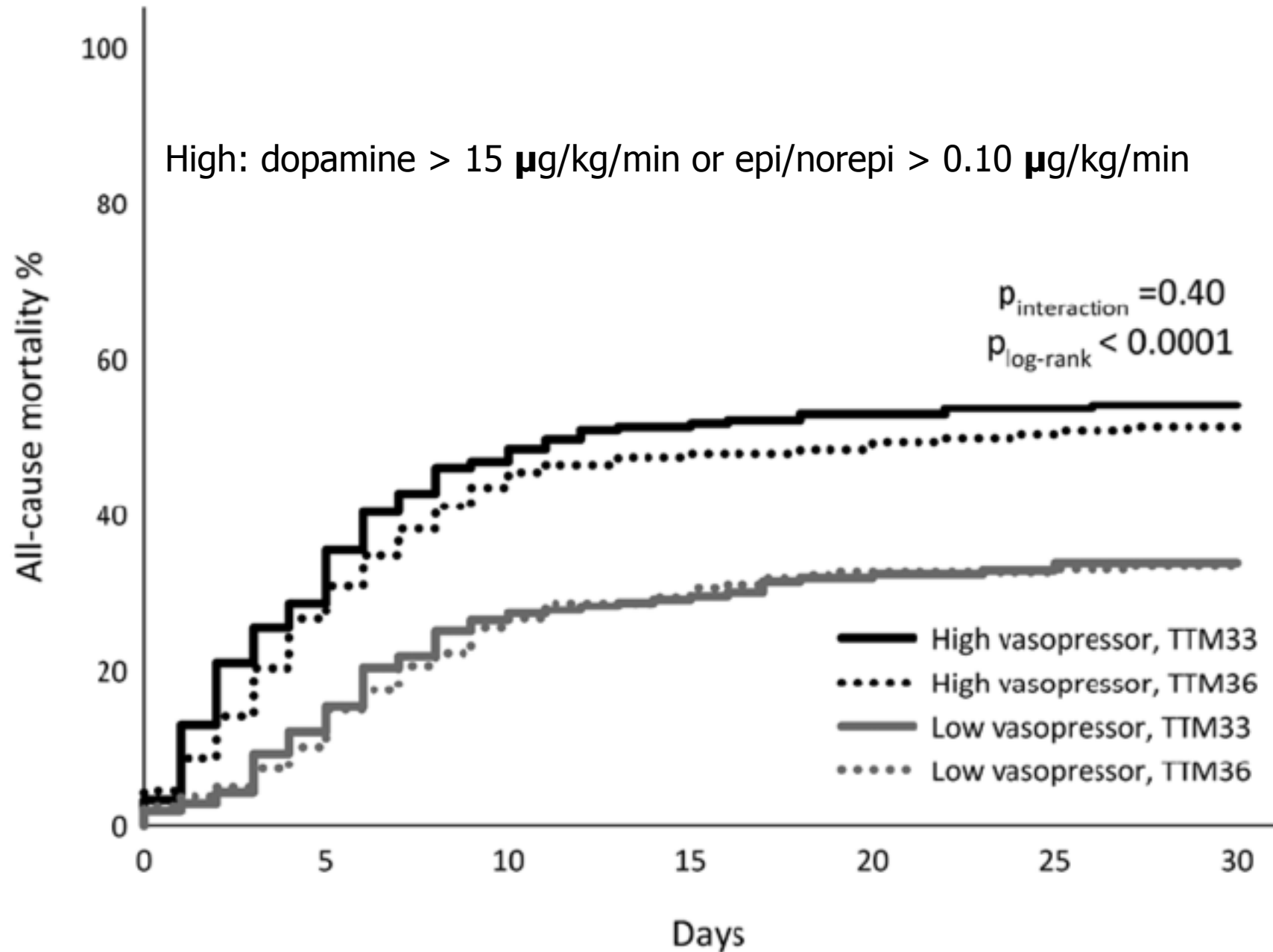


Death at discharge OR 1.06 (0.89-1.28)

Hemodynamics and Vasopressor Support During Targeted Temperature Management at 33°C Versus 36°C After Out-of-Hospital Cardiac Arrest: A Post Hoc Study of the Target Temperature Management Trial*



High: dopamine > 15 $\mu\text{g/kg/min}$ or epi/norepi > 0.10 $\mu\text{g/kg/min}$



Interpretation

- Consider targeted temperature management in comatose survivors of out of hospital cardiac arrest
- Temperature target 33 or 35 acceptable
- Avoid pyrexia
- Delay prognostication until 72 hours

Claudio Sandroni
Alain Cariou
Fabio Cavallaro
Tobias Cronberg
Hans Friberg
Cornelia Hoedemaekers
Janneke Horn
Jerry P. Nolan
Andrea O. Rossetti
Jasmeet Soar

Prognostication in comatose survivors of cardiac arrest: An advisory statement from the European Resuscitation Council and the European Society of Intensive Care Medicine

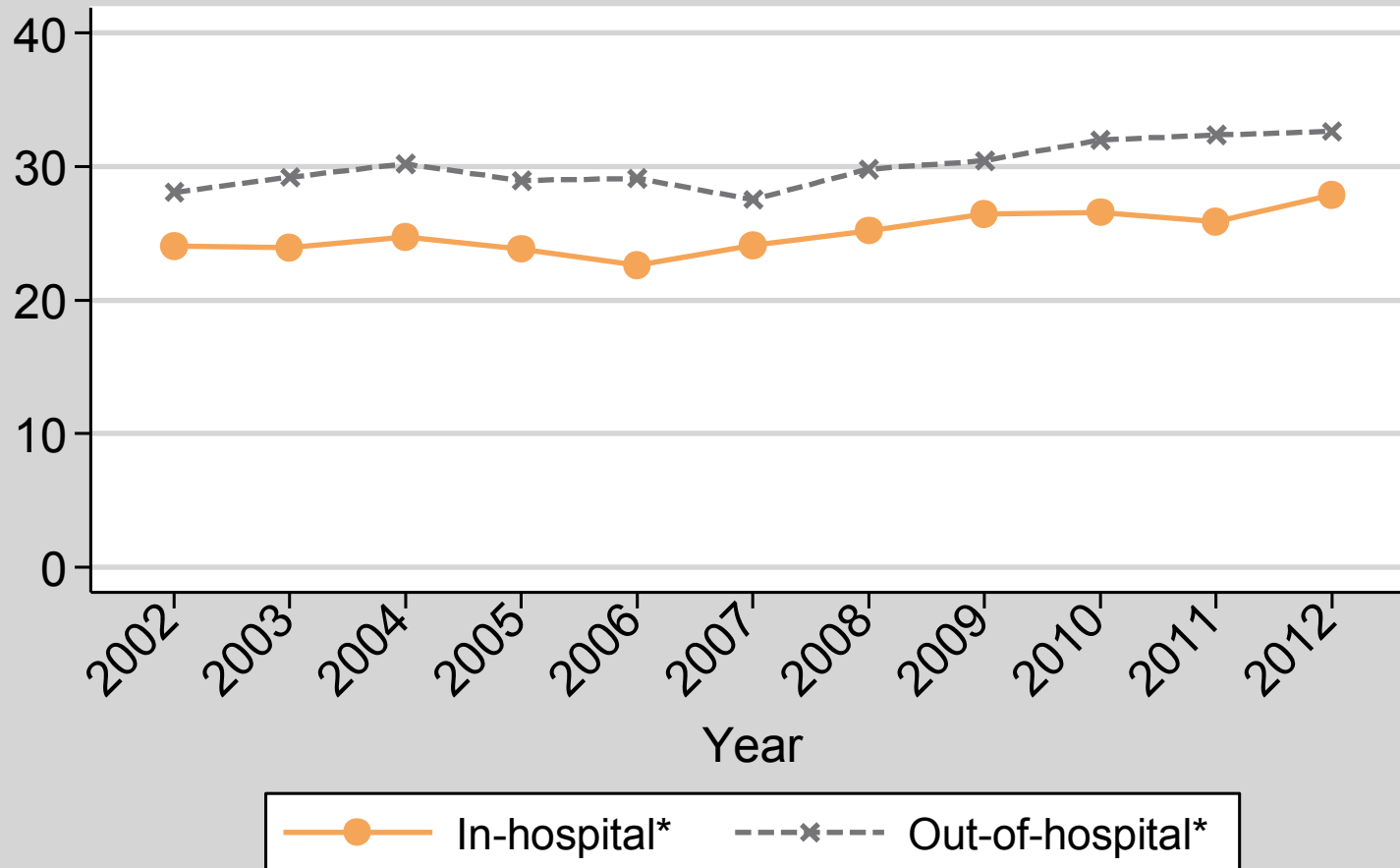
Prognostication in comatose survivors of cardiac arrest: An advisory statement from the European Resuscitation Council and the European Society of Intensive Care Medicine[☆]

Claudio Sandroni^{a,*}, Alain Cariou^b, Fabio Cavallaro^a, Tobias Cronberg^c, Hans Friberg^d, Cornelia Hoedemaekers^e, Janneke Horn^f, Jerry P. Nolan^g, Andrea O. Rossetti^h, Jasmeet Soarⁱ

Resuscitation 2014;85:1779-89

Active treatment withdrawn

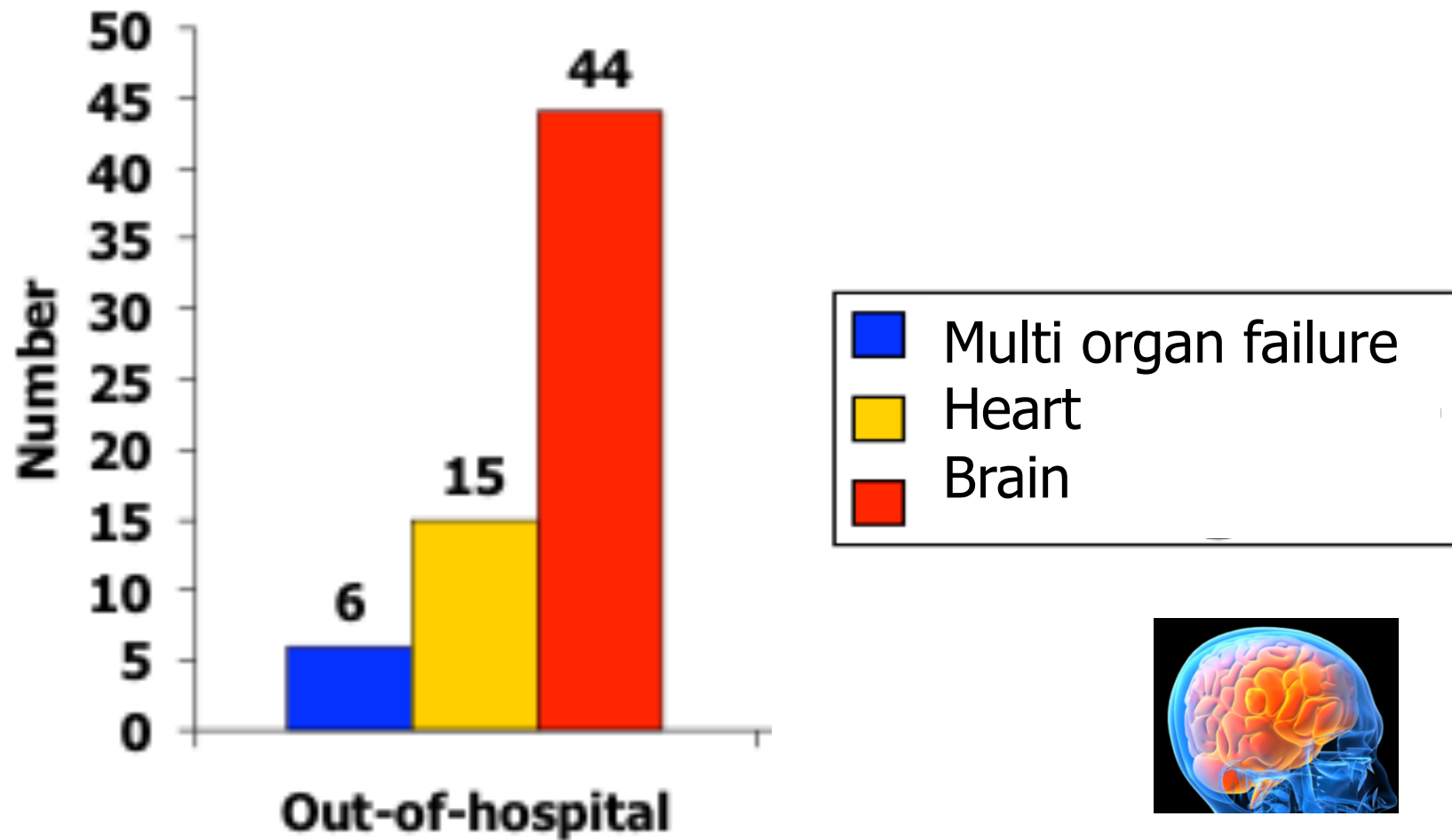
Percentage of admissions to critical care following cardiac arrest, with active treatment withdrawn

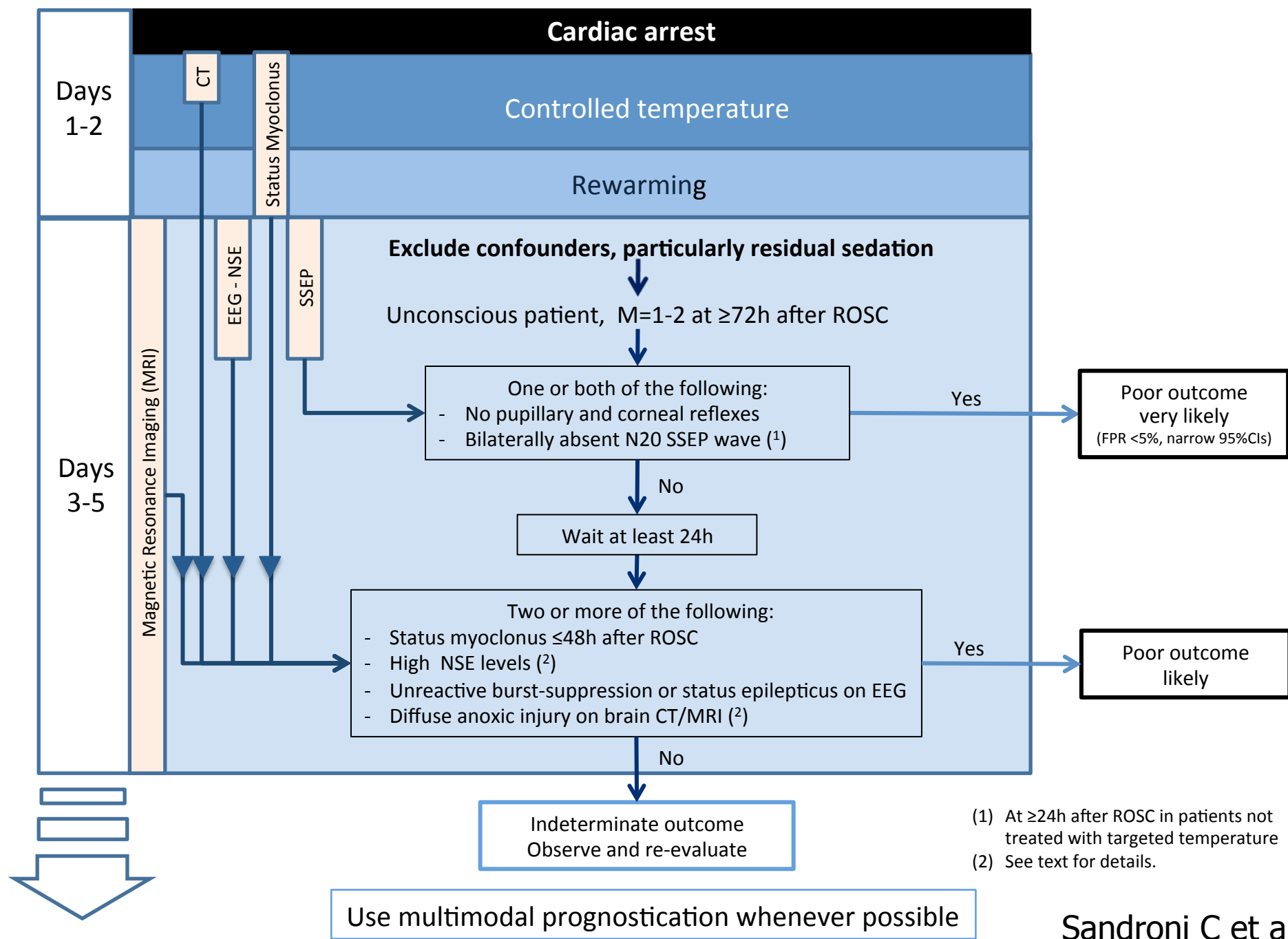


* location of cardiac arrest is approximated using location immediately prior to critical care admission

© ICNARC 2013

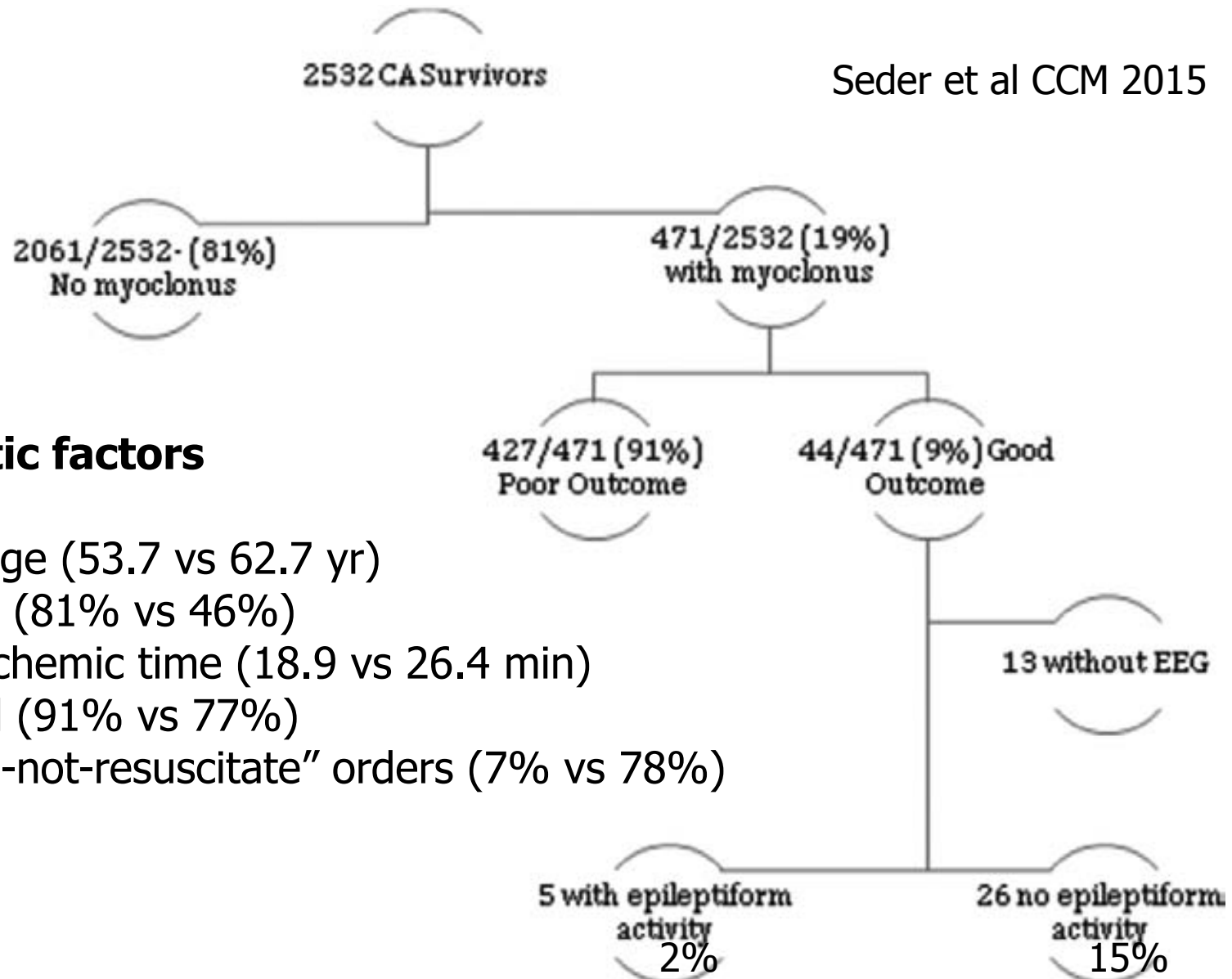
Treatment withdrawn in ITU – 30%





Neurologic Outcomes and Postresuscitation Care of Patients With Myoclonus Following Cardiac Arrest

Seder et al CCM 2015



Prognostic factors

Younger age (53.7 vs 62.7 yr)

Shockable (81% vs 46%)

Shorter ischemic time (18.9 vs 26.4 min)

Witnessed (91% vs 77%)

Fewer "do-not-resuscitate" orders (7% vs 78%)

Clinical examination

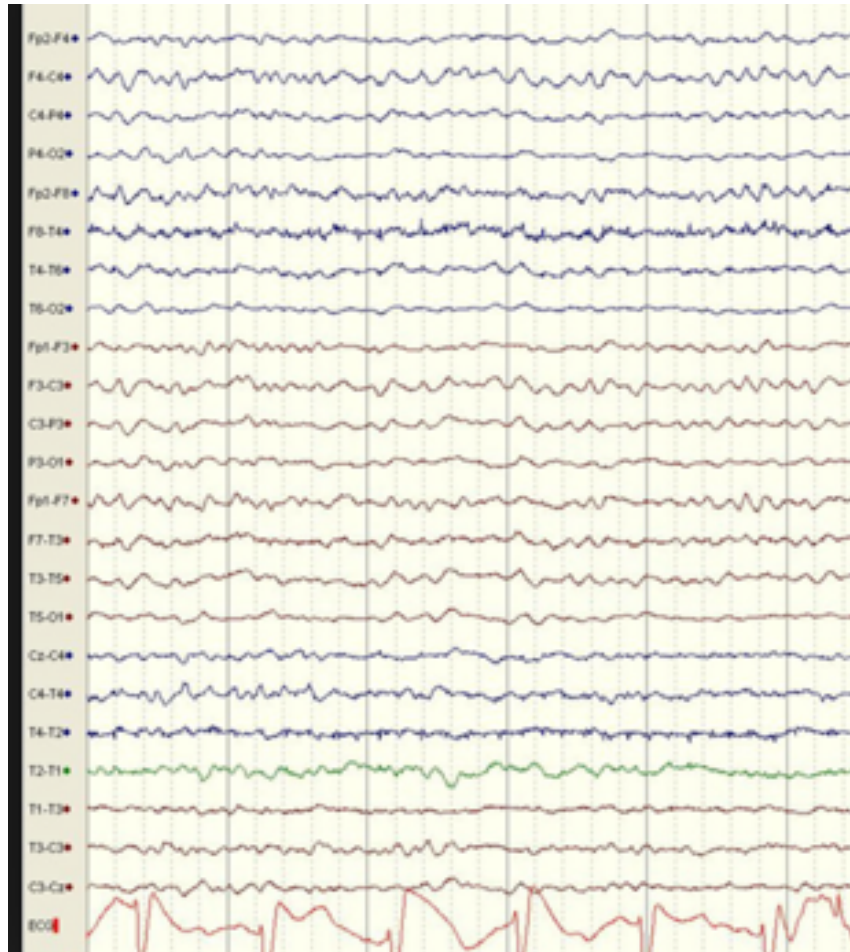
	Sensitivity	False positive rate
Motor response to pain	74%	27% (12-48%)
Bilateral absence of pupillary reflex at 24 hours		8%(1-25%)
Bilateral absence pupillary reflex at 72 hours (TH)	24%	0%(0-2%)
Bilateral absence pupillary reflex at 72 hours (NTH)	18%	0%(0-8%)
Bilateral absence corneal reflexes (NTH)	29%	4%(1-7%)

Brain imaging



- Reduced GM/WM ratio
- Sulcal effacement on brain CT within 24 h
- Sens 81%, FPR 8% (0-38%)
- MRI extensive reduction in diffusion (at 2-5 days)

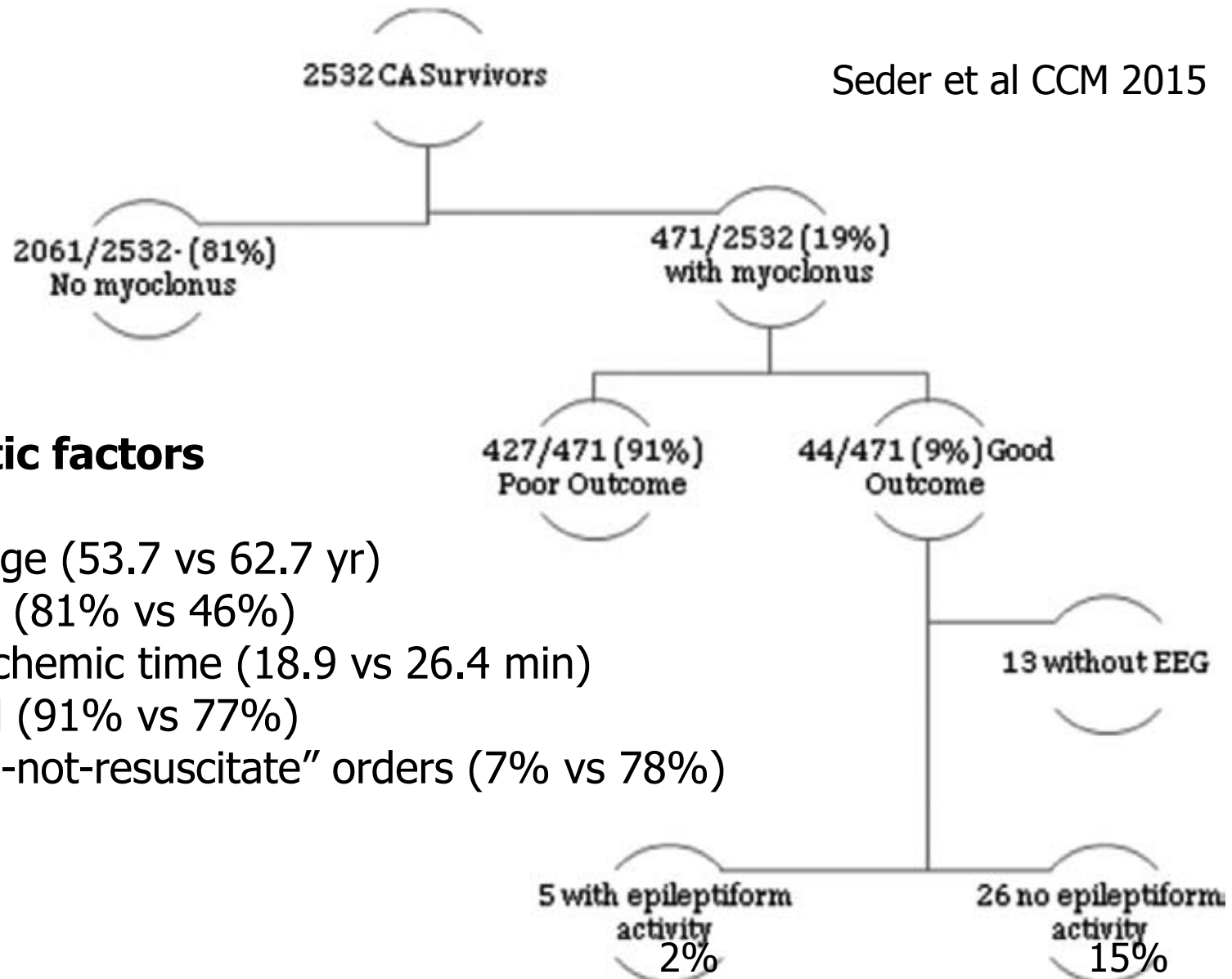
Neurophysiology



- Bilateral absence of SSEP at > 72 h
 - Sensitivity 45%,
 - FPR 0(0-2%)
- EEG
 - Absence of reactivity
 - Burst-suppression
 - Status epilepticus

Neurologic Outcomes and Postresuscitation Care of Patients With Myoclonus Following Cardiac Arrest

Seder et al CCM 2015



Prognostic factors

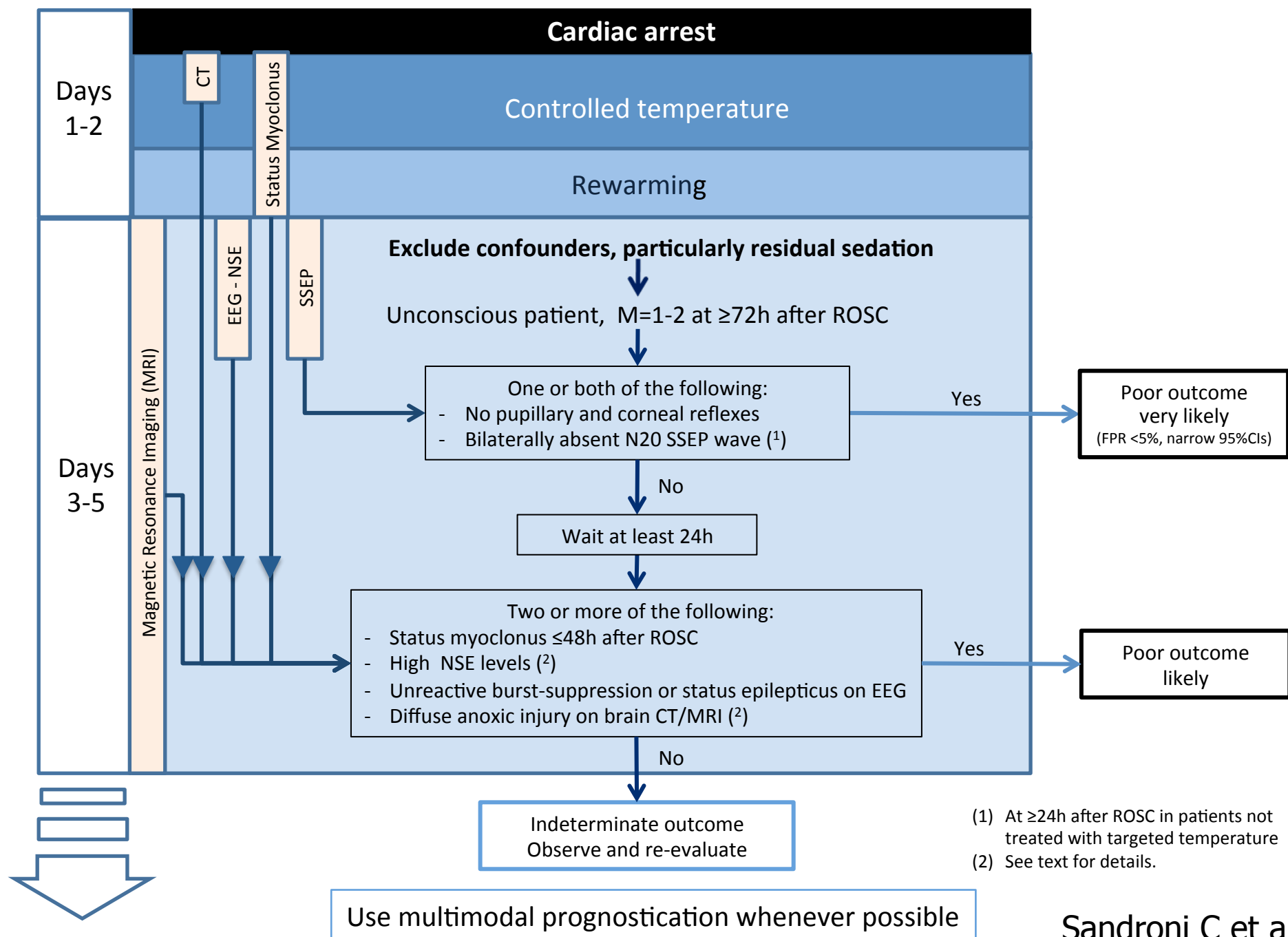
Younger age (53.7 vs 62.7 yr)

Shockable (81% vs 46%)

Shorter ischemic time (18.9 vs 26.4 min)

Witnessed (91% vs 77%)

Fewer "do-not-resuscitate" orders (7% vs 78%)



International Consensus Conference on ECC and CPR Science with Treatment Recommendations 2005, 2010... and 15th Oct 2015



Summary

- Update on new science and controversies in cardiac arrest
- Focus on chain of survival
 - High quality CPR
 - Uncertainty about pharmacological treatments
 - Targeted temperature management
 - Neuroprognostication



Scientific
Symposium 2015







Highlights

- ✓ Guidelines 2015
- ✓ The science behind the changes
- ✓ Hot topics



Save the date!