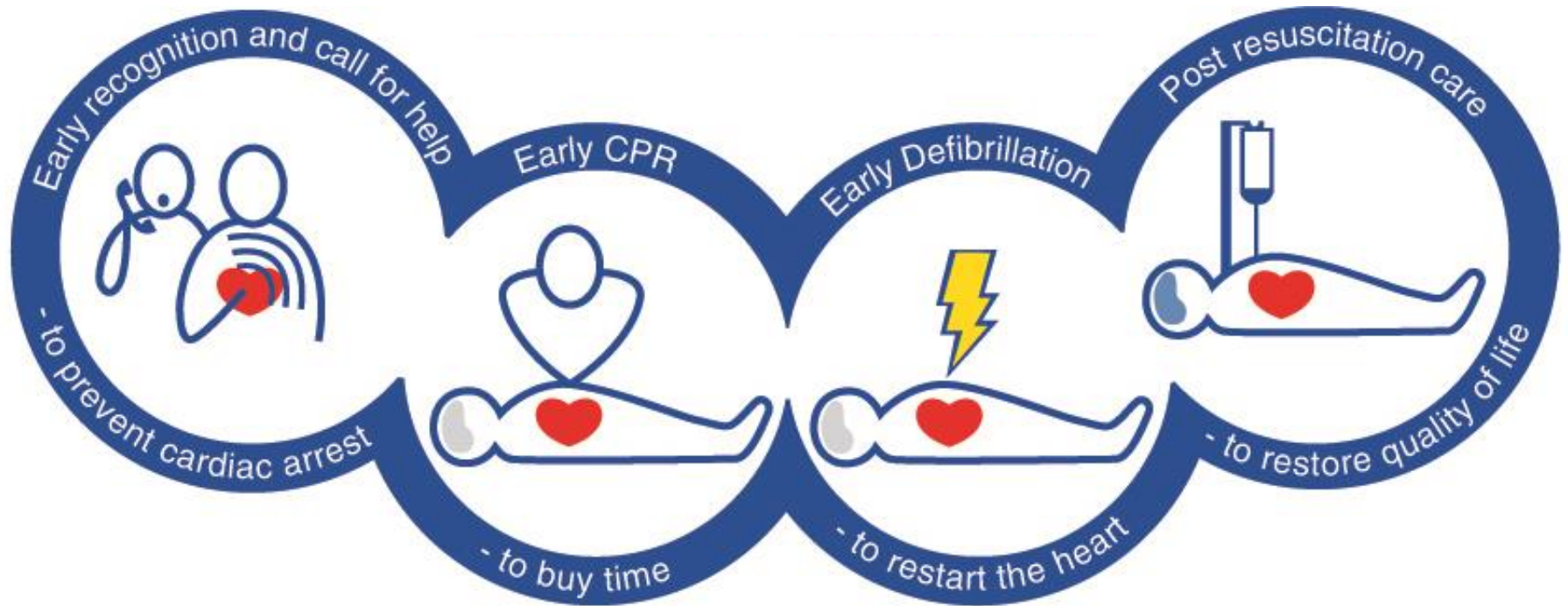


Post Cardiac Arrest Patient in the ICU



Jerry Nolan
Royal United Hospital
Bath

North East Intensive Care Society
Spring Meeting
23 March 2010





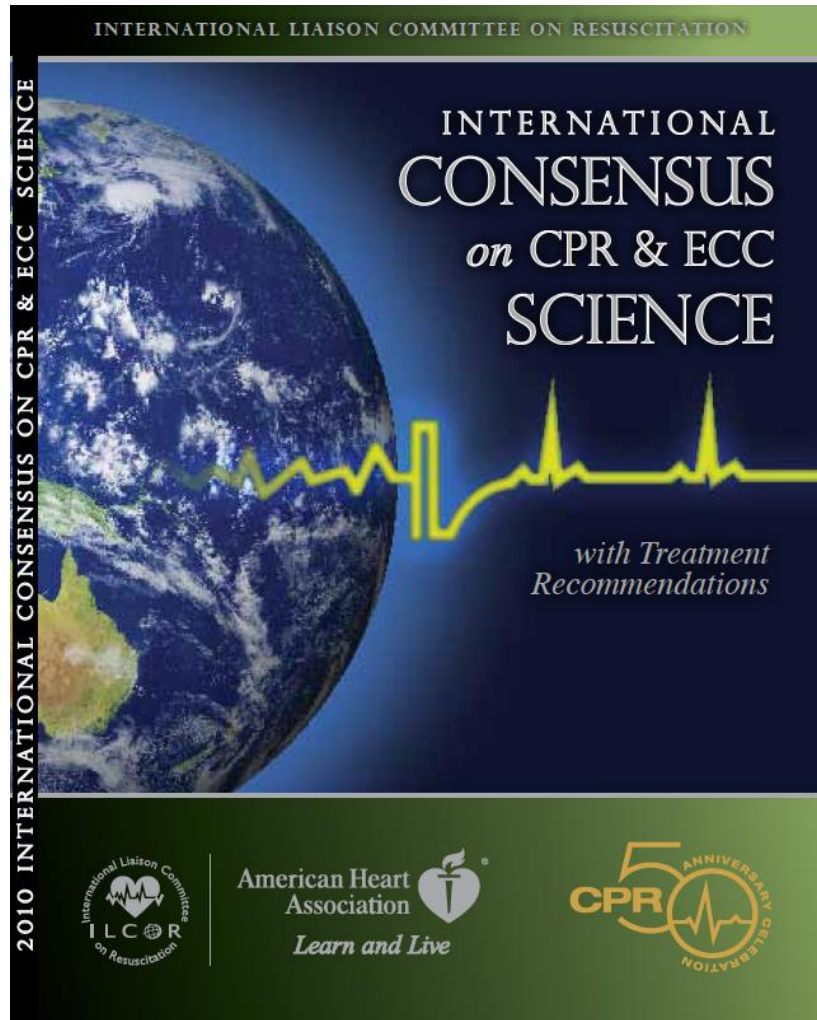
Post cardiac arrest patient in ICU

- Outcome after admission to ITU
- The post cardiac arrest syndrome
- Cardiovascular support
- Neurological support
- Prognostication

2010 International Consensus Conference on CPR Science

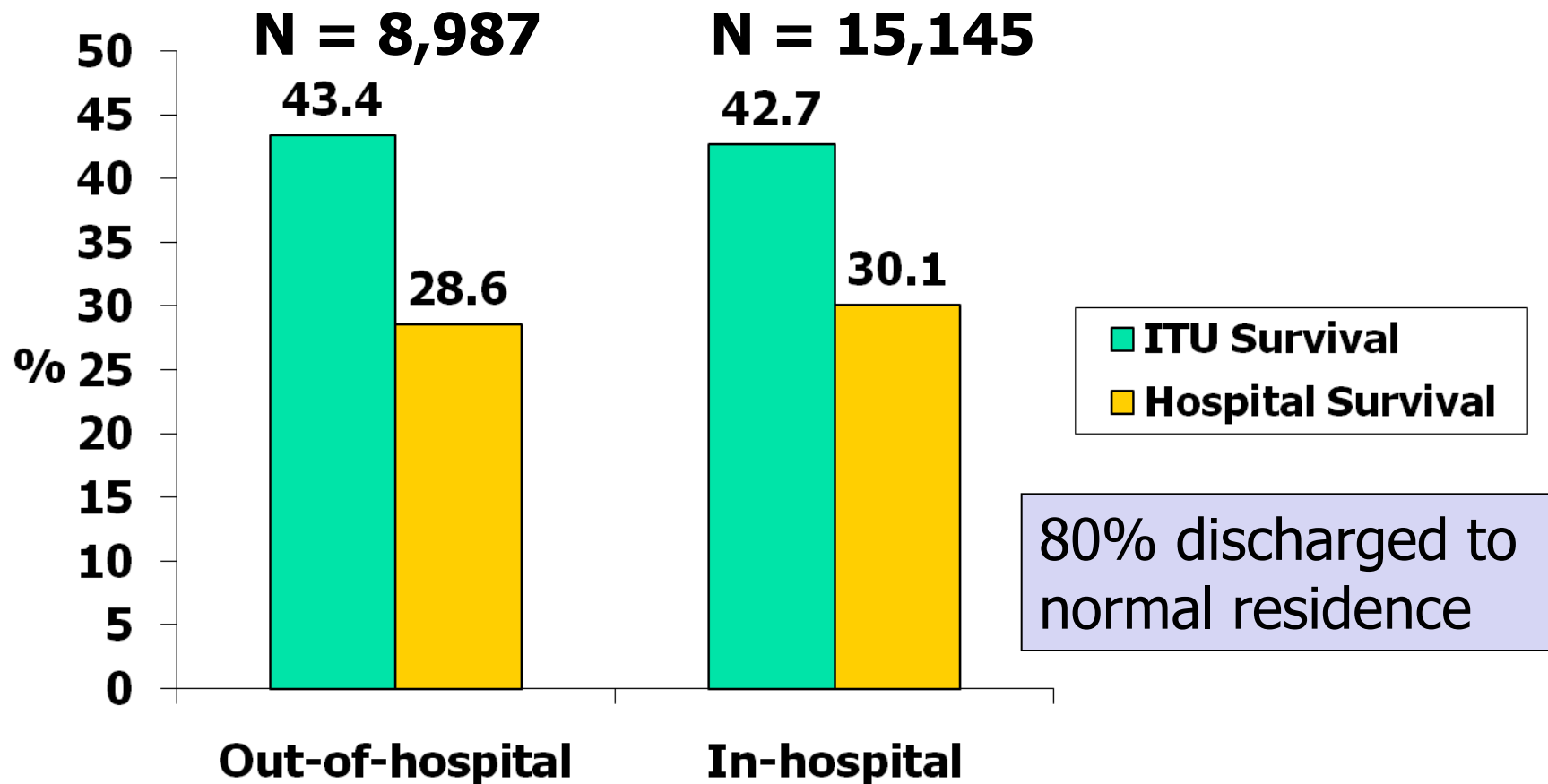


- ILCOR Consensus Conference 1-4 Feb 2010
- 450 evidence-based reviews
- Treatment recommendations based on science
- New AHA, ERC and RC (UK) guidelines online 18/10/10

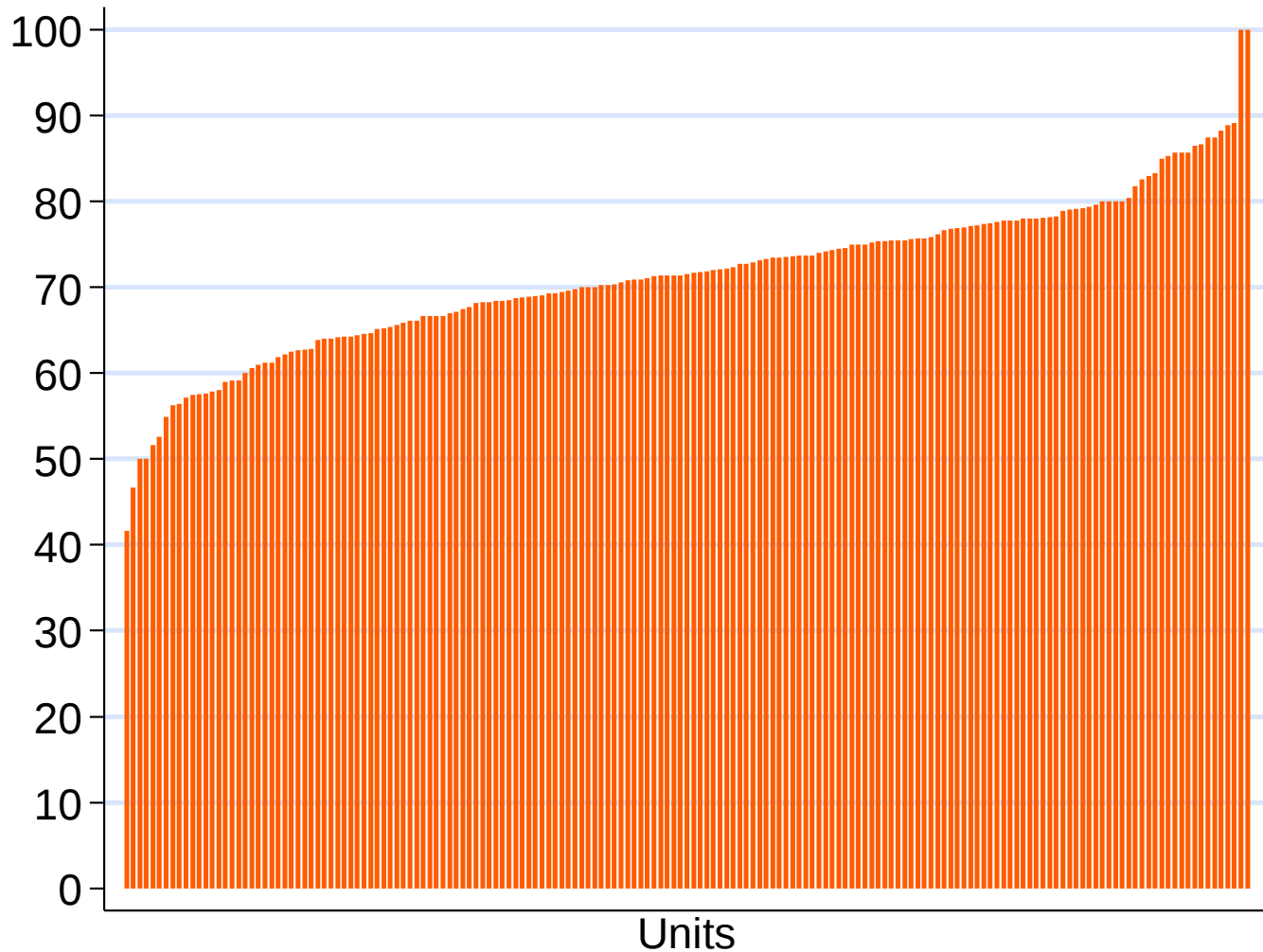


www.ilcor.org

Outcome for cardiac arrest survivors admitted to ITU in UK



Variation in acute hospital mortality – out-of-hospital arrest



Data from ICNARC



Standards for the Management of Patients After Cardiac Arrest

STANDARDS AND GUIDELINES



available at www.sciencedirect.com



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



ILCOR CONSENSUS STATEMENT

Post-cardiac arrest syndrome: Epidemiology, pathophysiology, treatment, and prognostication

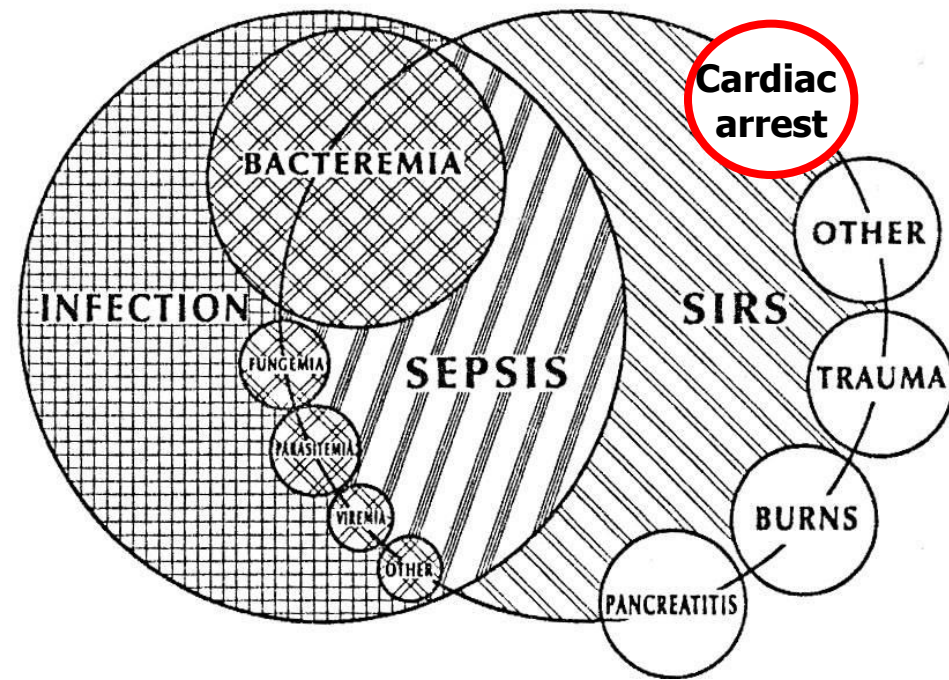
A Scientific Statement from the International Liaison Committee on Resuscitation;

Nolan JP, Neumar RW et al. Resuscitation 2008;79:350-79

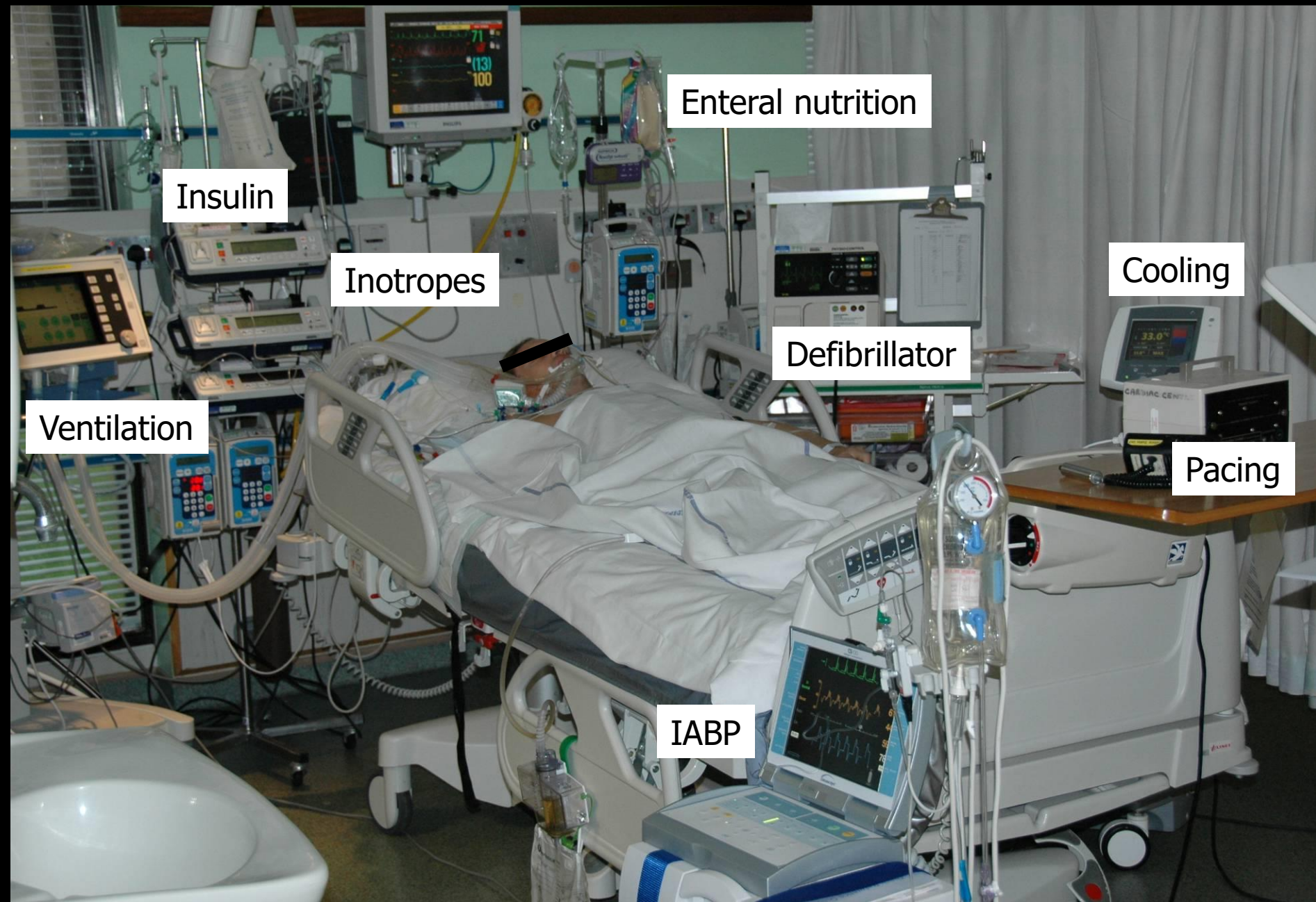
Post-cardiac arrest syndrome

1. Systemic ischaemia/reperfusion response
2. Persistent precipitating pathology
3. Post-cardiac myocardial dysfunction
4. Post-cardiac arrest brain injury

Inflammatory response after cardiac arrest



- High levels of IL-6, IL-8, IL-10, TNF
 - Adrie C. Circulation 2002;106:562-8
- Coagulation and fibrinolysis activated
 - Adrie C. JACC 2005;46:21-8.



Enteral nutrition

Insulin

Inotropes

Cooling

Defibrillator

Ventilation

Pacing

IABP

Early coronary reperfusion: PCI versus thrombolysis



Outcome of PCI after out-of-hospital cardiac arrest & STEMI

STEMI
Emergency angio and PCI
N = 186

Successful PCI
N = 161 (86.5%)

Alive at 6 months
N = 100 (53.8%)

Data from 5 centres with
high-volume PCI in France

Shock	96 (52%)
Intra-aortic balloon pump	80 (43%)
Mechanical ventilation	171 (92%)

Acute angiography after resuscitation from cardiac arrest

	STEMI	Not STEMI
Urgent angiography	192/192 (100%)	50/200 (25%)
Coronary disease	99%	78%
Culprit lesion	94%	36%
Urgent PCI	91%	32%
TIMI 3 Flow	81%	88%

Jan 2003 – Dec 2007
Ljubljana, Slovenia

Radsel P. Presented at ReSS 2008

Reversible myocardial dysfunction in survivors of out-of-hospital cardiac arrest

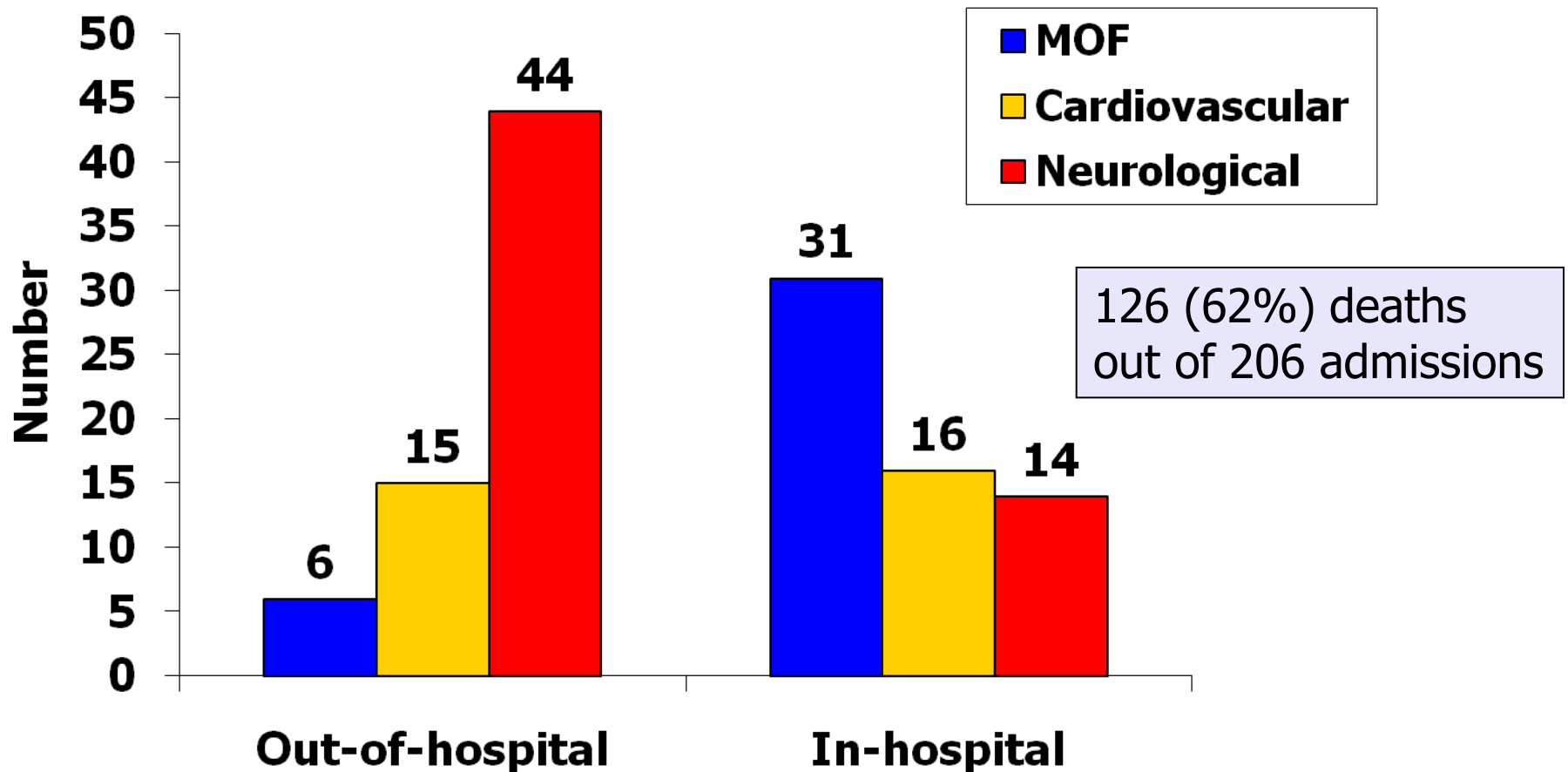
(n = 73)	8h	12 h	24 h	67 h
Adrenaline (mg h ⁻¹)	1.0	1.3	1.5	0.4
CI (l min ⁻¹ m ⁻²)	2.05	2.61	3.19	3.69
SVRI	2,908	1,936	1,672	1,518
Temp (°C)	36.6	37.3	37.6	37.8

Cardiovascular support

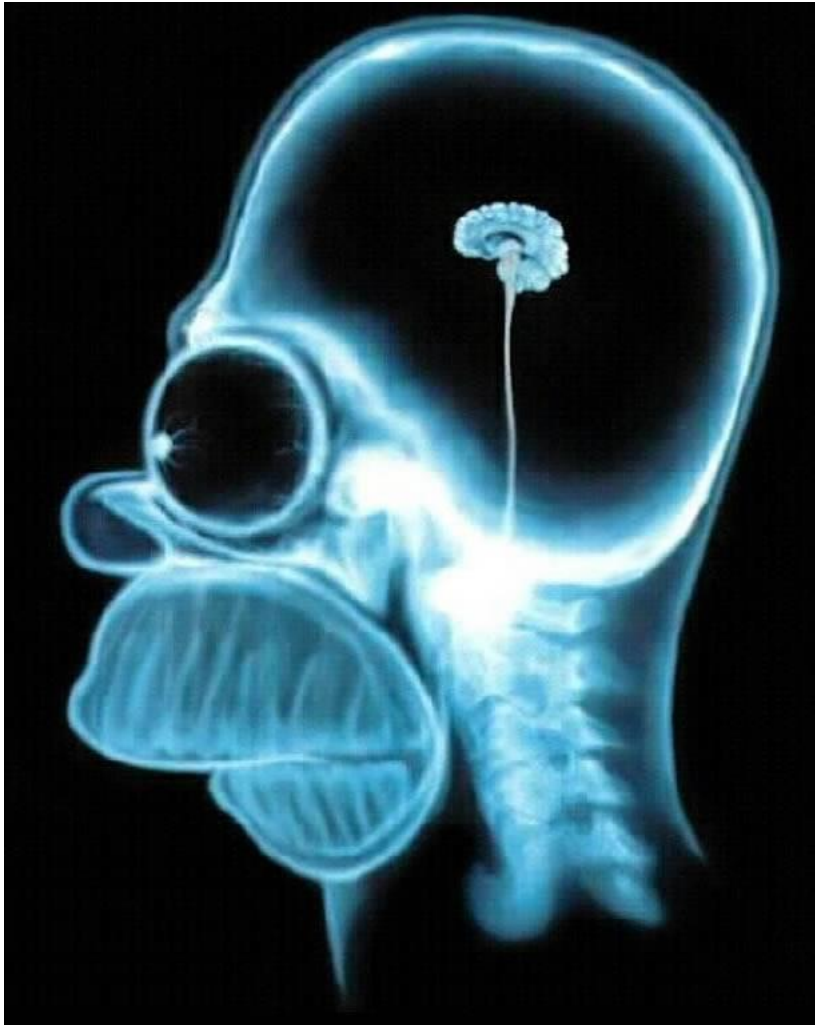


- Intra-aortic balloon pump
- Inotropes
 - Noradrenaline
 - Dobutamine
- Anti-arrhythmic therapy
- Pacing

Mode of death after admission to ITU following cardiac arrest



Improving neurological outcome after cardiac arrest

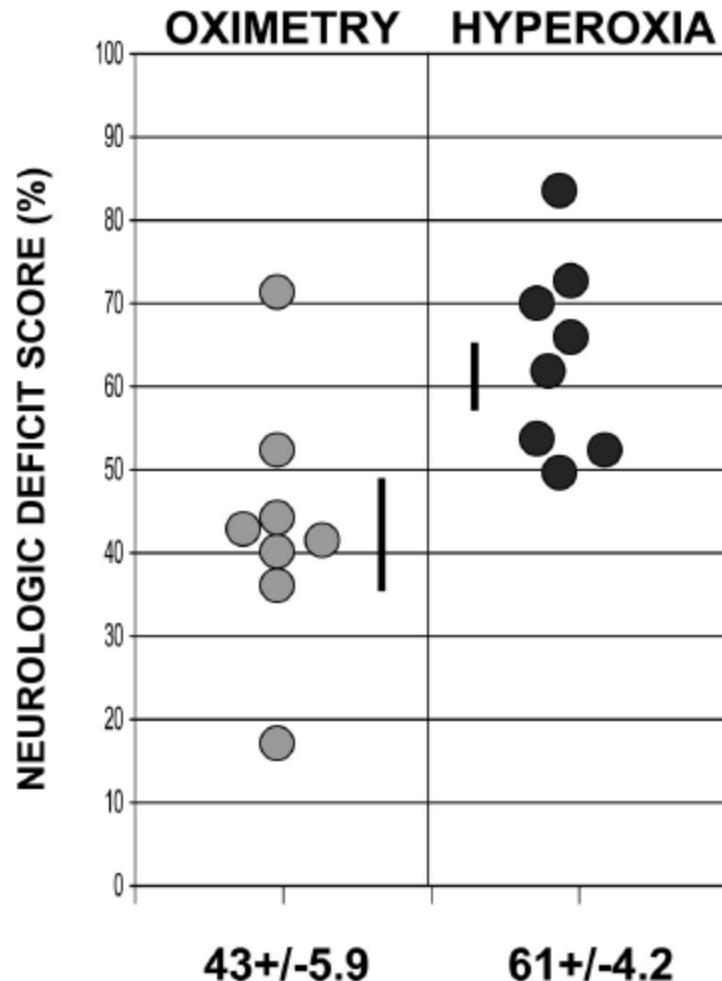


- Controlled re-oxygenation
- Cerebral perfusion
- Sedation
- Control of seizures
- Glucose control
- Temperature control

Hyperoxia after return of spontaneous circulation

- Hyperoxia during early reperfusion may harm postischaemic neurones
 - Vereczki V. J Cerebral Blood Flow Metab 2006;26:821-5
 - Richards EM. Stroke 2007; 38: 1578-84
 - Zwemer CF. Resuscitation 1994; 27: 159-70.
 - Liu Y. Stroke 1998;29: 1679-86.

Oximetry-guided re-oxygenation improves outcome after cardiac arrest



- 8 dogs per group
- 10 min cardiac arrest
- 1 hour ventilation:
 - 100% O₂ versus
 - SpO₂ 94 – 96%

Brain – cerebral perfusion

- Cerebral hyperaemia (initially), but rarely high ICP except asphyxial arrest
- Impaired autoregulation
 - Nishizawa H. Acta Anaesthesiol Scand 1996;40:1149-53.
- Maintain adequate cerebral perfusion pressure
- MAP 80-100 mmHg??
 - Bell DD. Can J Anaesth 2005;52:309-322

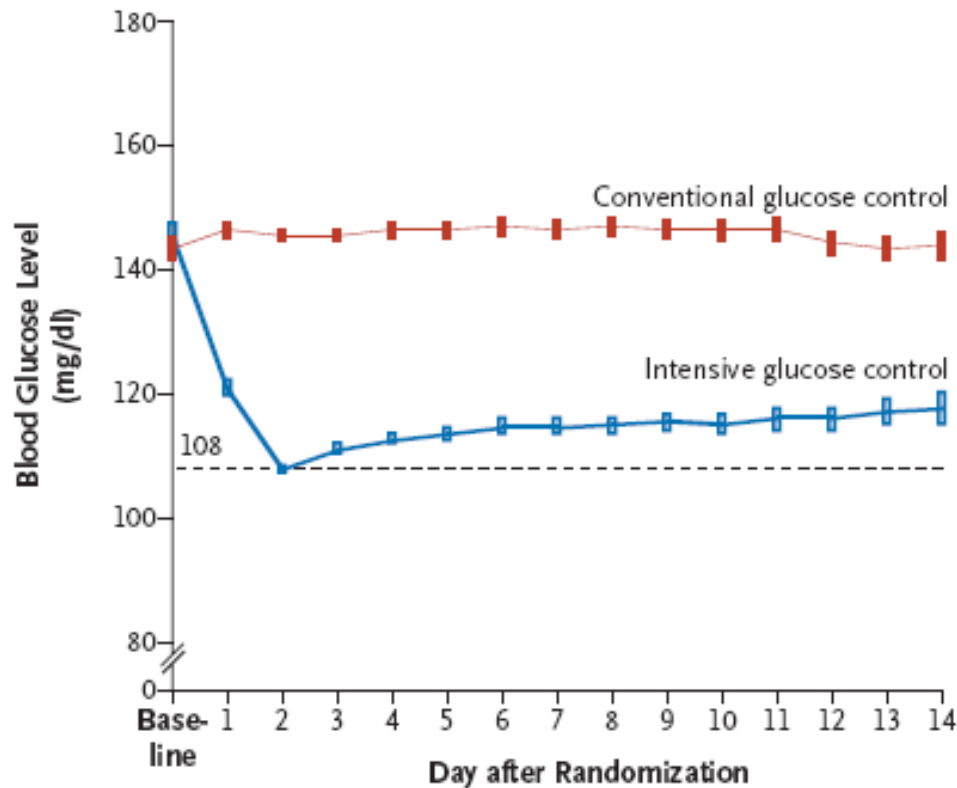
Brain – control of seizures

- Occur in 5-15% of those who achieve ROSC and in 40% of those remaining comatose
- Increase cerebral oxygen consumption
- Drugs
 - No data on the use of prophylactic anti-convulsants
- Monitoring; especially with NMBs?

Strict versus moderate glucose control after VF cardiac arrest

- Out-of-hospital VF cardiac arrest
- 90 patients – all cooled (intravascular)
- Insulin for 48 h; randomised:
 - Glucose 4 – 6 mmol l⁻¹ (SGC)
 - Glucose 6 – 8 mmol l⁻¹ (MGC)
- 30-day mortality and hypos (<3 mmol)
 - SGC 13/39 (**33%**) **18%**
 - MGC 18/51 (**35%**) **2%**

NICE-SUGAR Study



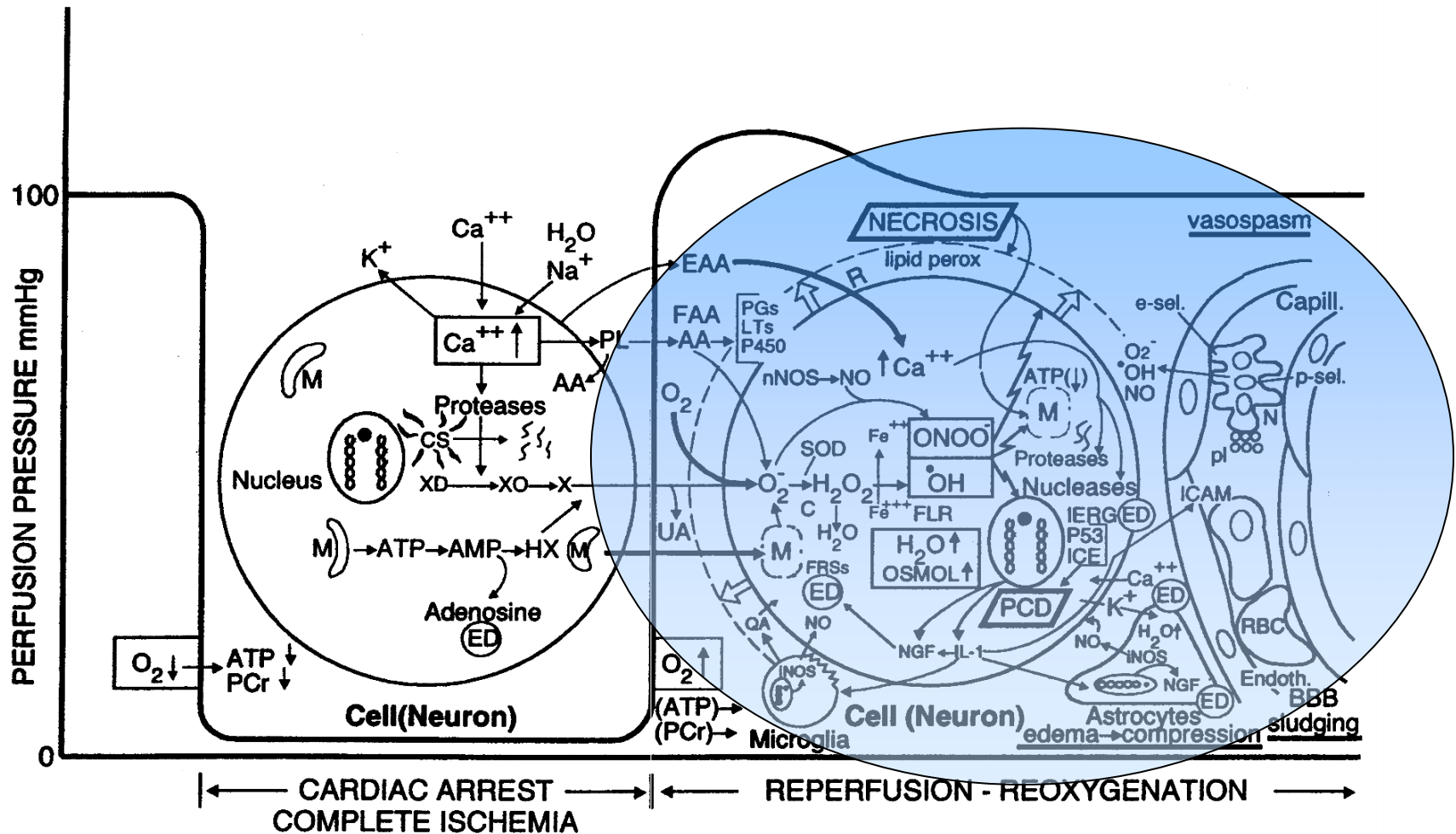
- 6104 patients predicted ICU stay > 3 days:
 - 4.5 – 6.0 mmol L⁻¹
 - < 10.0 mmol L⁻¹
- 90-day mortality
 - Intensive 27.5%
 - Conventional 24.9%
 - P = 0.02

Brain – temperature control

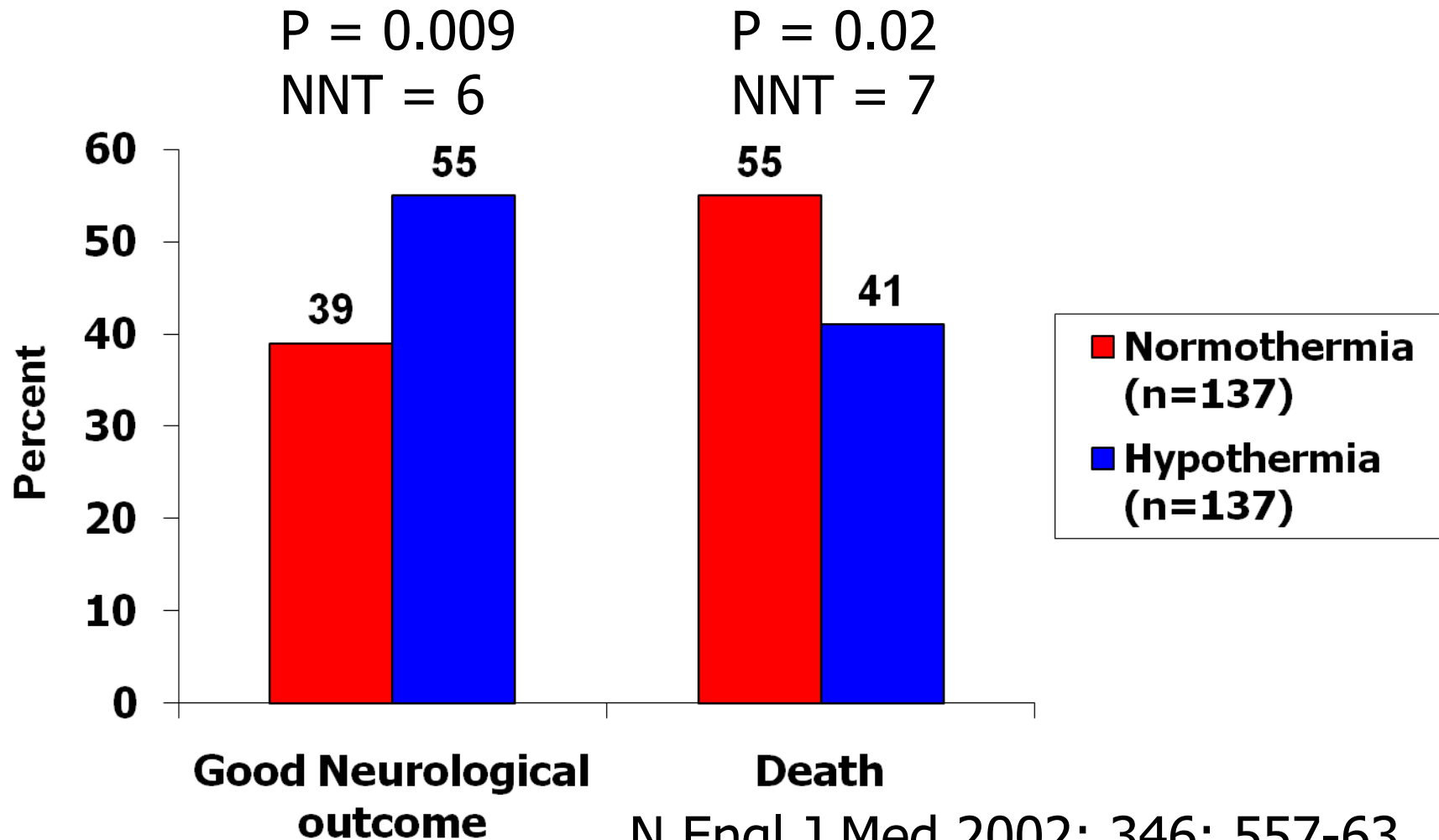
- Prevention of hyperthermia
 - Hyperthermia common for 2-3 days
 - Takino M. Intensive Care Med 1991;17:419-20
 - Hyperthermia associated with poor outcome
 - Zeiner A. Arch Intern Med 2001;161:2007-12
 - Hickey RW. Crit Care Med 2003;31:531-5
- Therapeutic hypothermia

Hypothermia:

Mechanism of action?



The Hypothermia After Cardiac Arrest (HACA) Study Group



N Engl J Med 2002; 346: 557-63

Therapeutic hypothermia after cardiac arrest

An Advisory Statement by the ALS Task Force of the
International Liaison Committee on Resuscitation (ILCOR)

- Unconscious adult patients with spontaneous circulation after out of hospital cardiac arrest should be cooled to 32-34°C for 12-24 hours when the initial rhythm was VF
- For any other rhythm, or cardiac arrest in hospital, such cooling may also be beneficial

Cooling techniques

External

- Ice packs, wet linen, fans
- Cooling blankets
 - Air, e.g. Polar Air
 - Water, e.g. Blanketrol
- Pre-refridgerated cooling pads
- Hydrogel-coated pads
- Cold water immersion

Internal

- Cold i.v. saline
- Intravascular catheters
 - Intravascular balloons
 - Metal catheter
 - Helix system



Start cooling
with cold i.v.
fluid

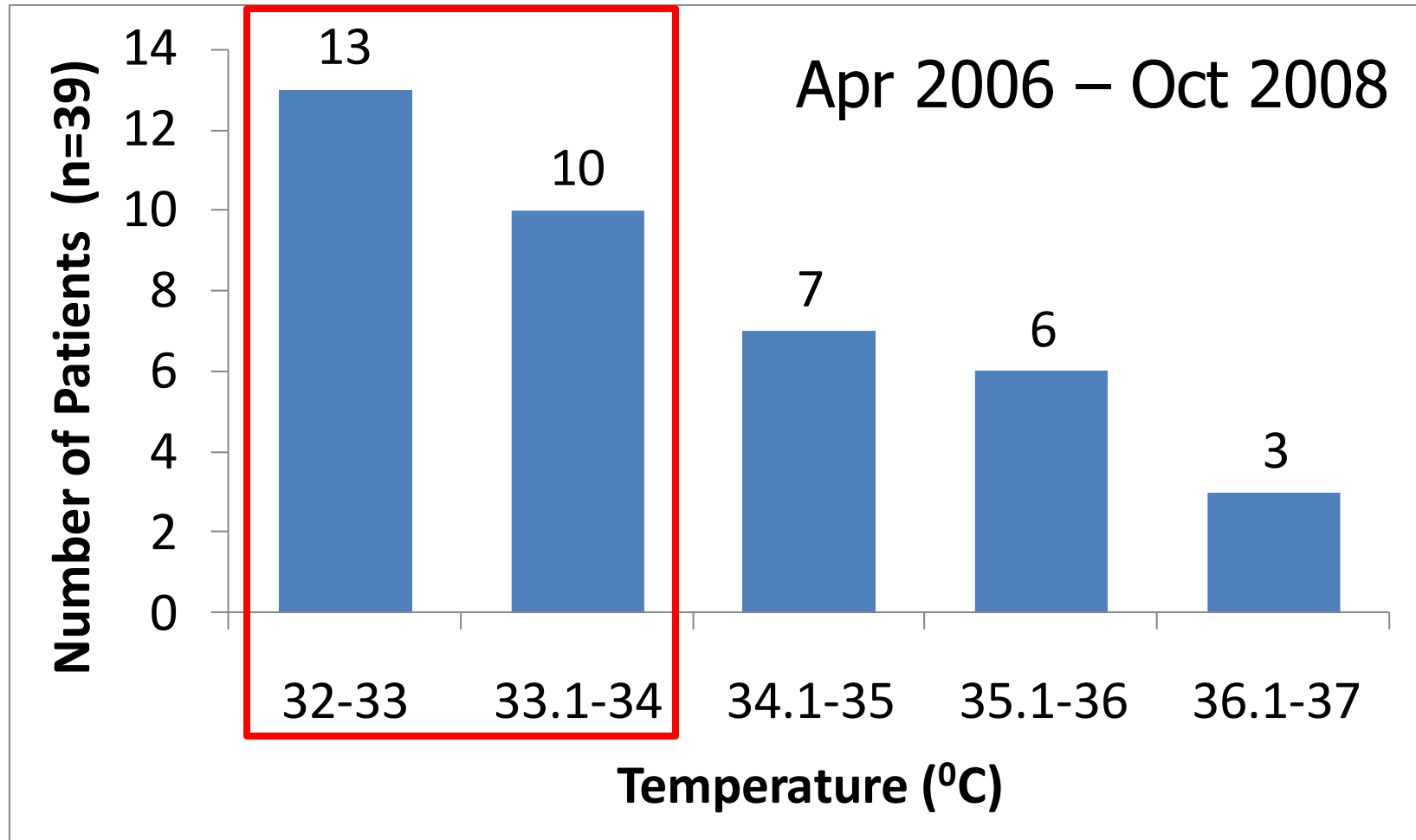
Cooling with ice-cold i.v. fluid

Study	N	T °C	Comments
Bernard S. Resuscitation 2003;56:9-13	22	1.7	30 ml kg ⁻¹ Hartmann's at 4°C. No pulmonary oedema
Kliegel A. Resuscitation 2005;64:347	26	1.8	2 pts mild pulmonary oedema
Virkkunen I. Resuscitation 2004;62:299	13	1.8	Prehospital – 2 more trials underway
Kim F. Circulation 2007;115:3064-70	125	1.2	RCT prehospital – half given cold fluid. Pilot study
Bruel C. Crit Care 2008;12:R31	33	2.1	1 pt pulmonary oedema

RCT of prehospital cooling (n=1200) due to complete Nov 2011
[ClinicalTrial.gov NCT00391469](https://clinicaltrials.gov/ct2/show/study/NCT00391469)

Kim F. J Neurotrauma 2009;26:359-63

Out-of-hospital cardiac arrests: temp on admission to ITU



Does more rapid cooling improve outcome?

- 49 patients intravascularly cooled (78% OHCA; 84% VF/VT) **Wolff B. Int J Cardiol 2009;133:223-8**
 - Multivariate analysis: time to target temperature = OR 0.69 (0.51 – 0.98) for good outcome per hour
- Hypothermia network study of 986 cooled patients **Nielsen N. Acta Anaesthesiol Scand 2009;53:926-34**
 - Time to target, duration of cooling, target temp = no association with outcome

Prehospital cooling



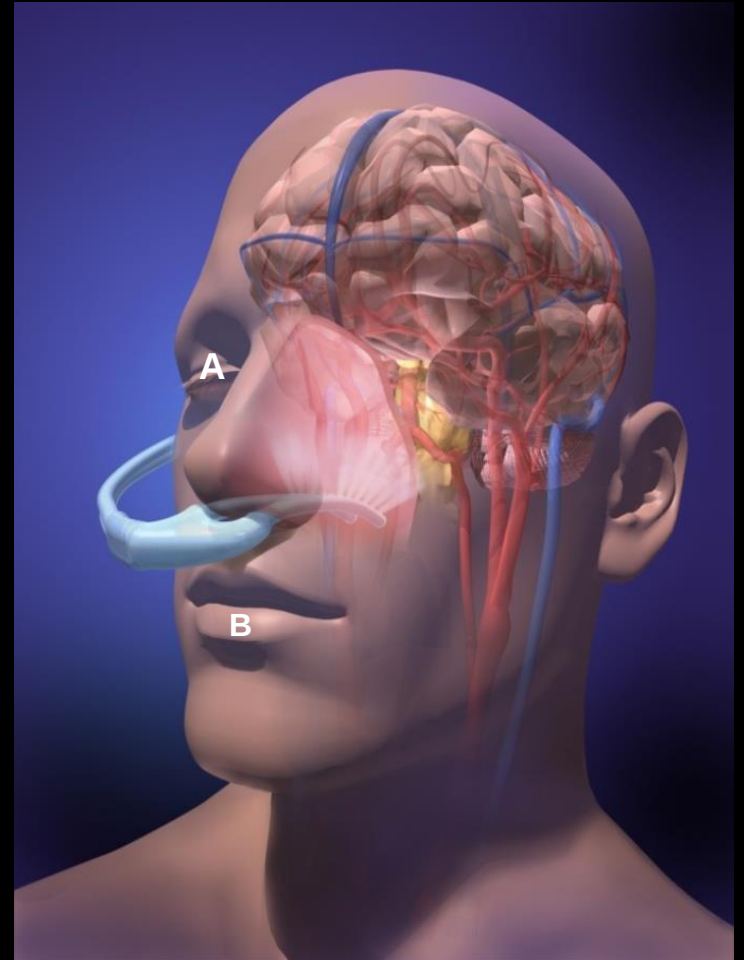
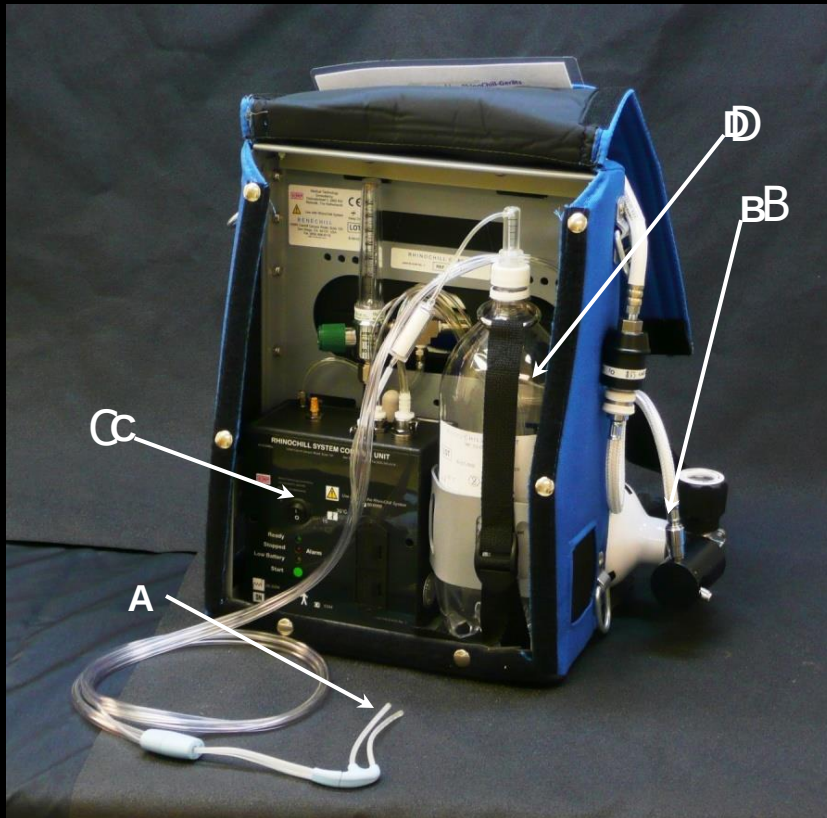
Prehospital cooling versus emergency department cooling

- VF cardiac arrest (n = 234)
- 2 litres cold saline prehospital vs. ED
- Temperature on ED arrival
 - Prehospital cooled 34.4°C
 - Control 35.9°C
- Survival to discharge
 - Prehospital cooled 48%
 - Control 51%

Cooling during CPR?

- Animal data indicate better outcomes
 - Kuboyama K. Crit Care Med 1991 & 1993
 - Nordmark J. Resuscitation 2005;66:357-65.
- Clinical 'pilot study' (n = 5)
 - Kamarainen A. Resuscitation 2008;76:360-3.
- Impact of fluid infusion during CPR??
- Cooling with cardiopulmonary bypass
 - Nagao K. Circ J 2010; 74: 77-85

RhinoChill™ intra-nasal cooling system



- A: Nasal Catheter
- B: Oxygen Tank
- C: Control Unit
- D: Coolant Bottle

External cooling

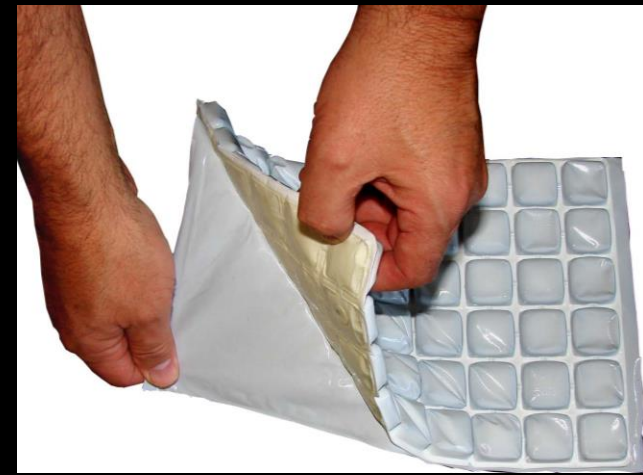


Larsson I-M. Resuscitation 2010;81:15-9

Laerdal MediCool



EMCOOLS



Uray T. Resuscitation 2008;77:331-8

Circulating cold water blankets



Water-circulating cooling device (Artic Sun[®])



Heard KJ. Resuscitation 2010; 81: 9-14

Convective-immersion surface cooling: ThermoSuit[®]



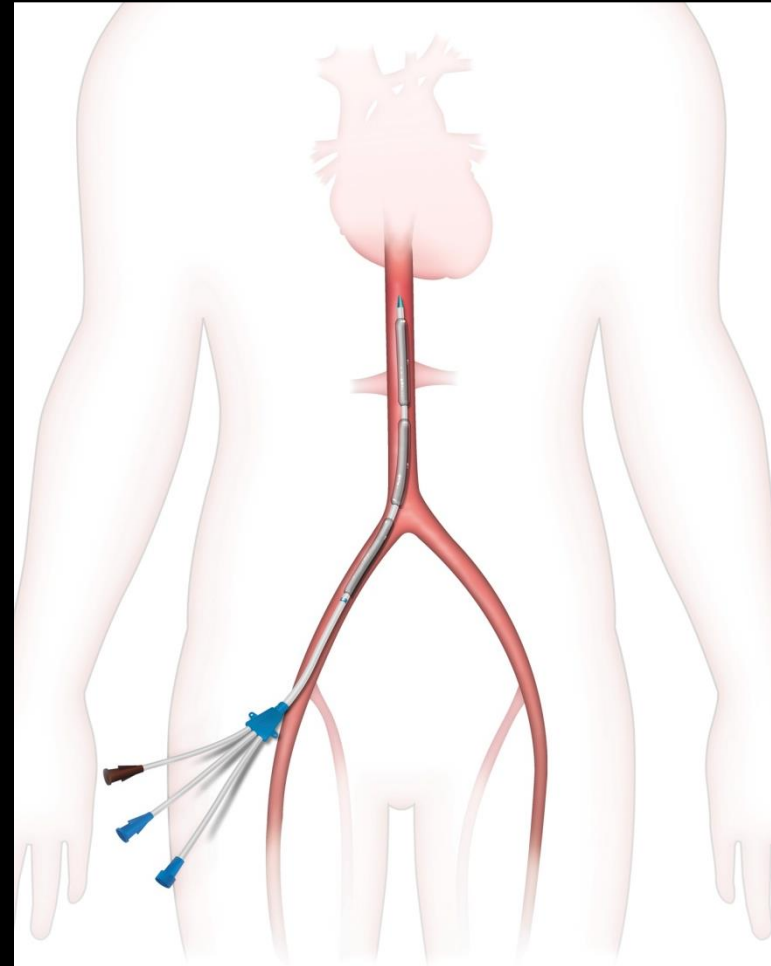
- 24 patients
- Median cooling rate = $3.0^{\circ}\text{C h}^{-1}$
- Faster with propofol
- Median time to $< 34^{\circ}\text{C}$ = 37 min

Life
Recovery
Systems

Photograph courtesy
of Dr M. Foedisch

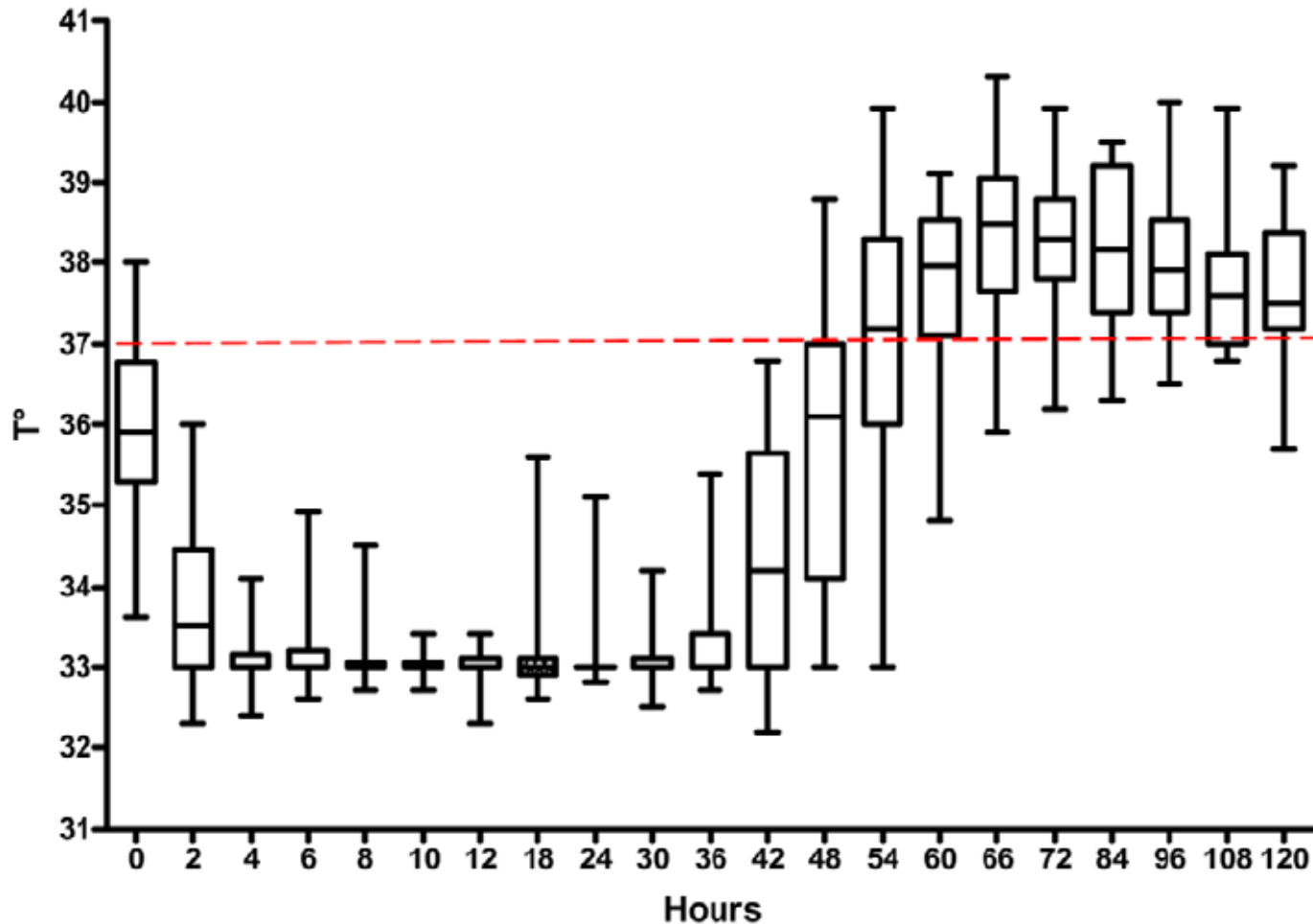


Endovascular cooling

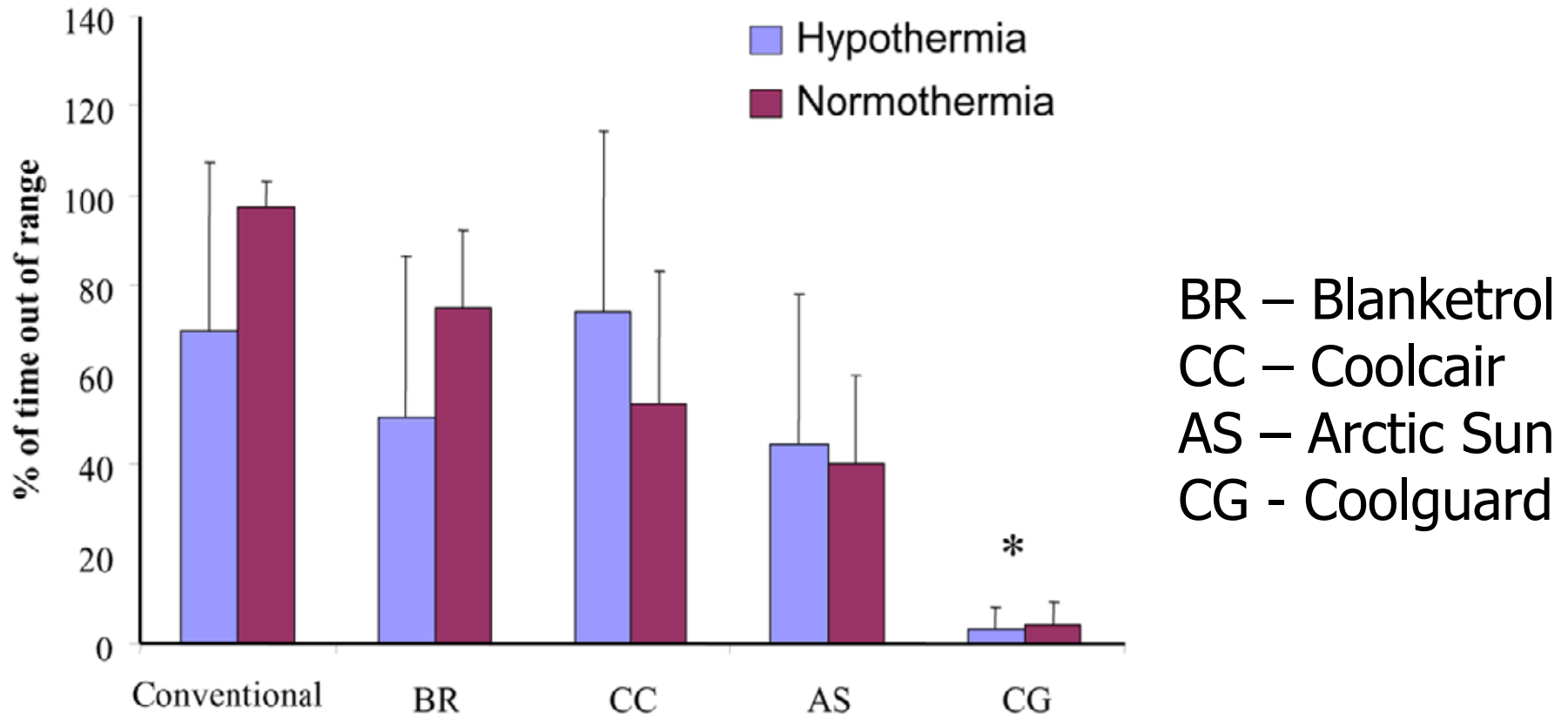


Intravascular cooling after OHCA

France, n = 40



Comparison of cooling methods: time out of range (n = 50)



Endovascular versus external cooling for OHCA

- ICEREA Study - Paris
 - 1^o outcome: CPC 1 or 2 at 28 days
 - Coolgard versus 'conventional cooling'
 - Finished recruiting Nov 2009 (400)
 - ClinicalTrials.gov NCT00392639

Duration of cooling?

- Optimal duration not defined
- Most centres using 24 h
- Optimal duration probably depends on:
 - time to ROSC
 - delay to target temperature
- Up to 72 h in some cases

Shivering

- More common during cooling and rewarming phases
- Sedation: e.g., propofol/opioid
- Neuromuscular blocker (NMB) bolus
- Occasionally, NMB infusion (Bath -12%)
 - Need for continuous EEG monitoring?

Magnesium

(e.g., 5 g infused over 5 h)

- Increases rate of surface cooling
 - Zweifler RM. Stroke 2004; 35: 2331-4
- Reduces shivering threshold
 - Wadhwa A. BJA 2005; 94: 756-62.
- Neuroprotective when combined with hypothermia
 - Zhu H. Brain Research 2004; 1014: 53-60.
- Anti-arrhythmic

Clinical practice protocols

Resuscitation 80 (2009) 418–424



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ACTA ANAESTHESIOLOGICA SCANDINAVICA
doi: 10.1111/j.1399-6576.2008.01881.x

Scandinavian Clinical practice guidelines for therapeutic hypothermia and post-resuscitation care after cardiac arrest

M. CASTRÉN¹, T. SILFVAST², S. RUBERTSSON³, M. NISKANEN⁴, F. VALSSON⁵, M. WANSCHER⁶ and K. SUNDE⁷ (Task Force on Scandinavian Therapeutic Hypothermia Guidelines, Clinical Practice Committee Scandinavian Society of Anaesthesiology and Intensive care Medicine)

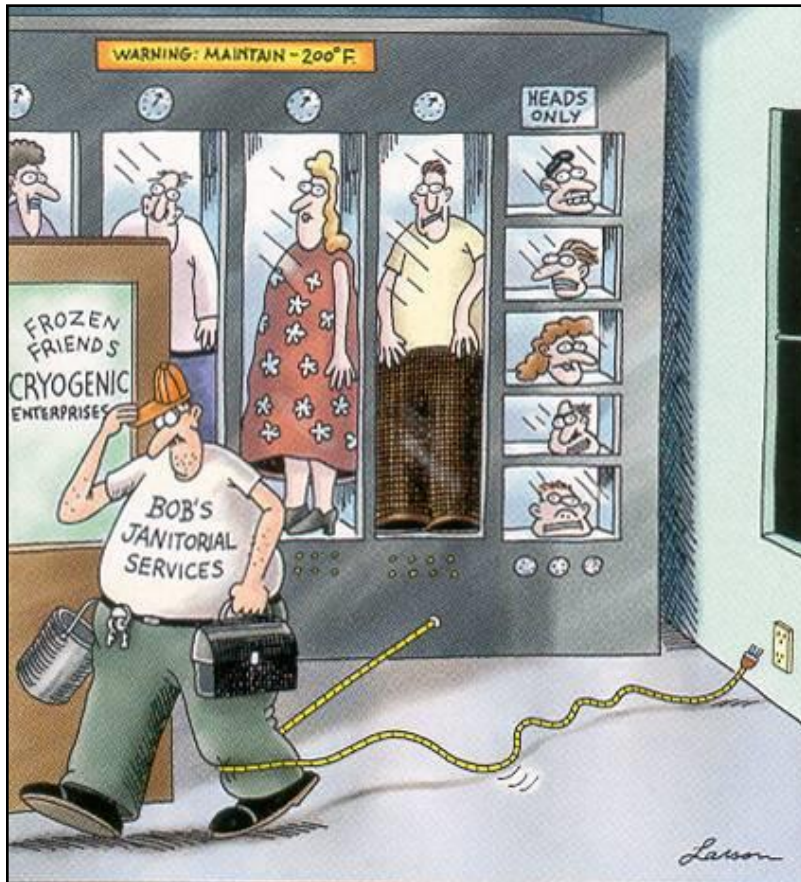
www.med.upenn.edu/resuscitation/hypothermia

Current practice in Bath

- Patients remaining comatose after cardiac arrest are cooled to 32-34°C once decision made to admit to ITU
- 2 l ice-cold crystalloid over 30 min
- Maintain hypothermia with intravascular cooling for 24 h
- Rewarm at 0.25°C h⁻¹
- Leave cooling catheter for 72 h (36.5°C)

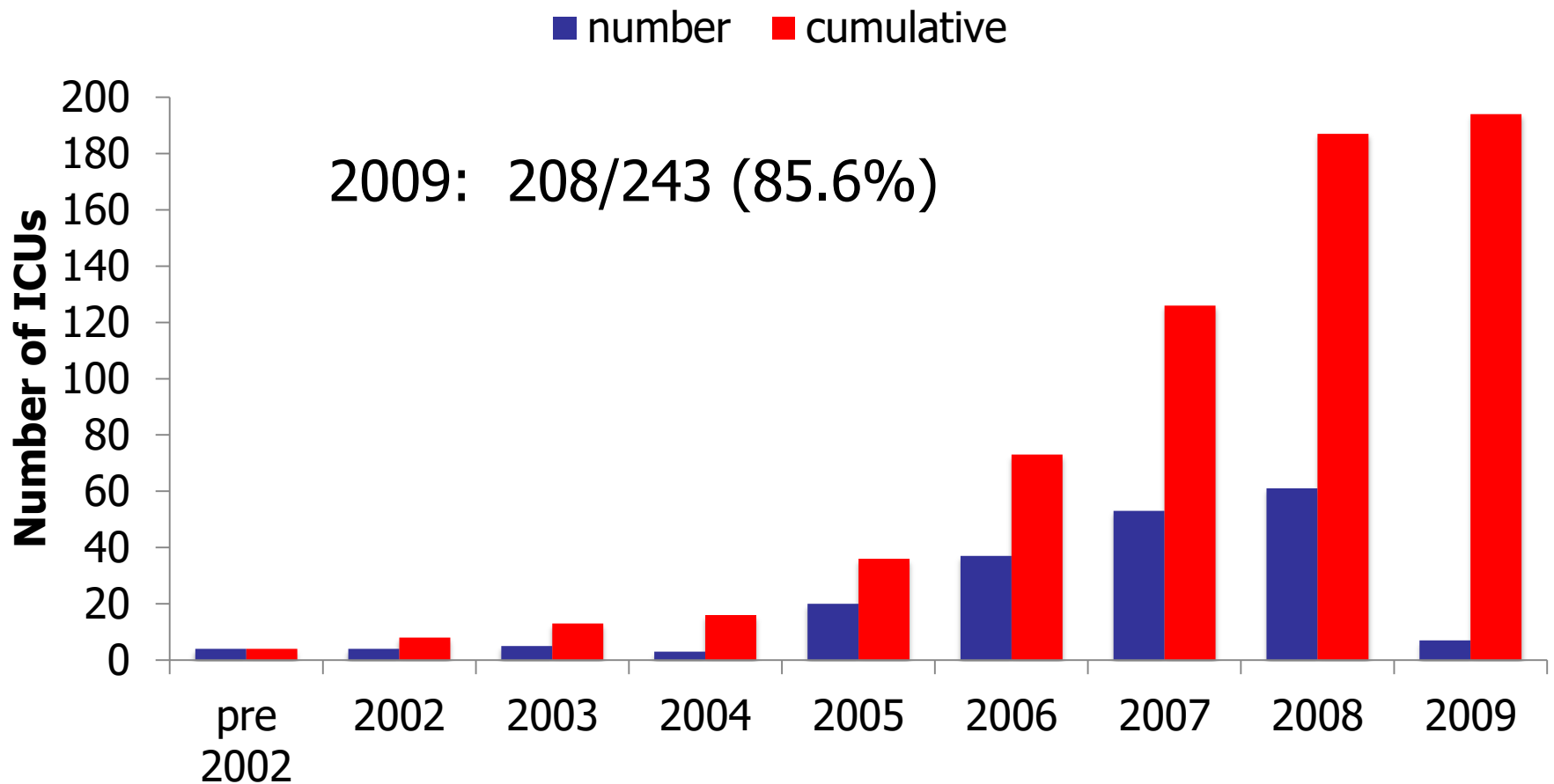
Therapeutic Hypothermia:

Physiological effects / complications

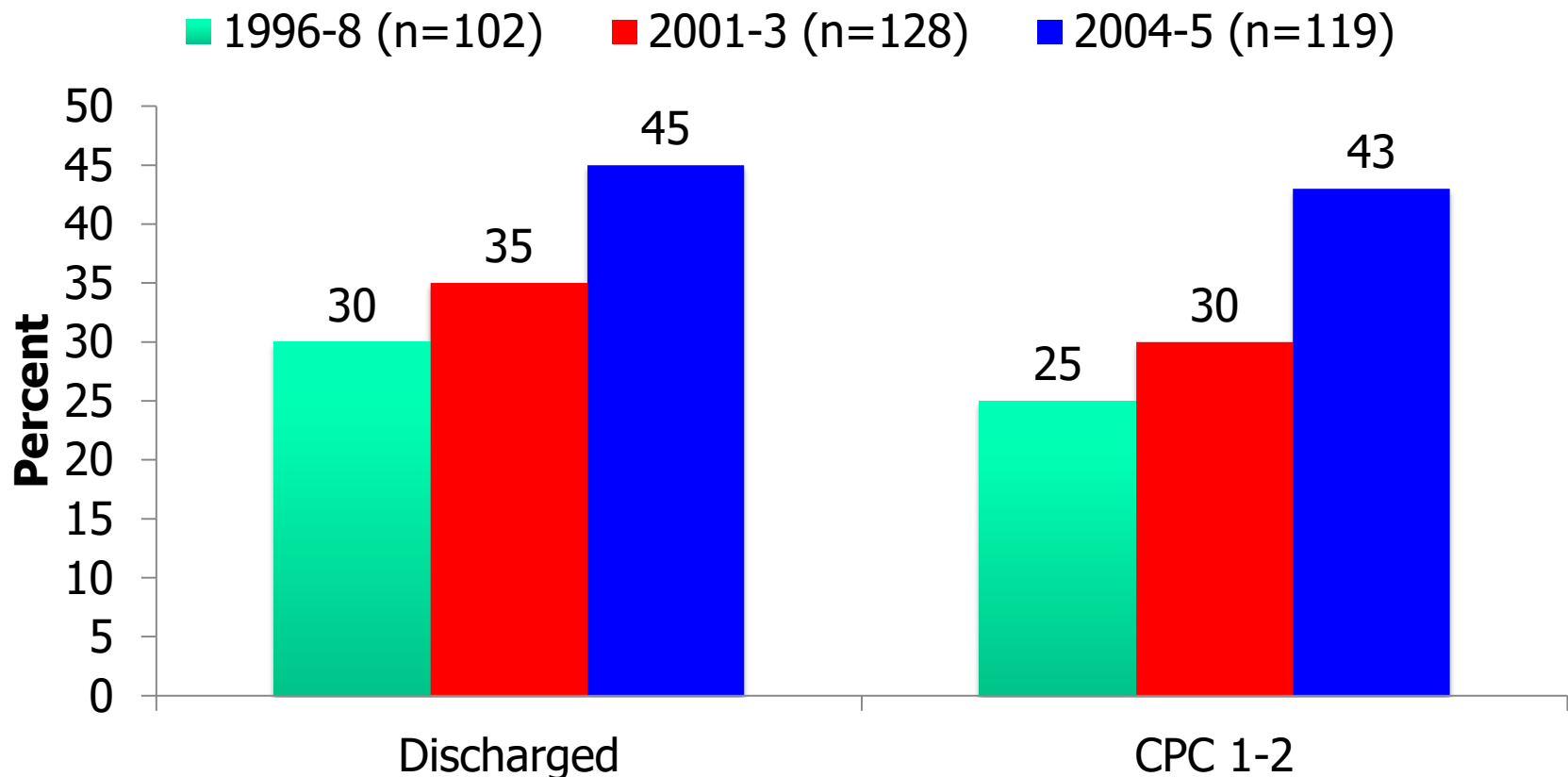


- Shivering
- Vasoconstriction
- Bradycardia
- Infection, coagulopathy
- Diuresis - hypovolaemia
- $\downarrow K^+$, $\downarrow Mg^+$, $\downarrow Ca^{2+}$
- $\downarrow$ Insulin sensitivity
- Impaired GI absorption

Implementation of hypothermia in UK ICUs (by year)



Survival after admission to ICU for OHCA – Oslo, Norway (All rhythms)



Reliable clinical indicators of bad neurological outcome



"I admire your persistence doctor.
But face it. You've lost this patient."

- Absent pupil **and** corneal reflexes at 3 days after cardiac arrest
- Absent or extensor motor reflexes at 3 days
- Myoclonic status epilepticus at any time
- Influence of hypothermia

Prognostication after cardiac arrest and hypothermia

Outcome at 3-6 months (n = 109)	CPC 1-2	CPC 3-5
Patients n (%)	25 (23)	84 (77)
ROSC > 25 min	6 (24)	46 (55)
≥ 1 brainstem reflex absent*	2 (8)	45 (54)
Motor response worse than flexion	4 (16)	64 (76)
Early myoclonus	1 (4)	35 (42)
Epileptiform activity on first EEG	3 (12)	36 (43)
Unreactive EEG background	0	56 (67)
Bilaterally absent N20 on SSEP	0	33 (46)

*pupillary, oculoccephalic, corneal

Rossetti AO. Ann Neurol, in press

Regional Systems of Care for Out-of-Hospital Cardiac Arrest

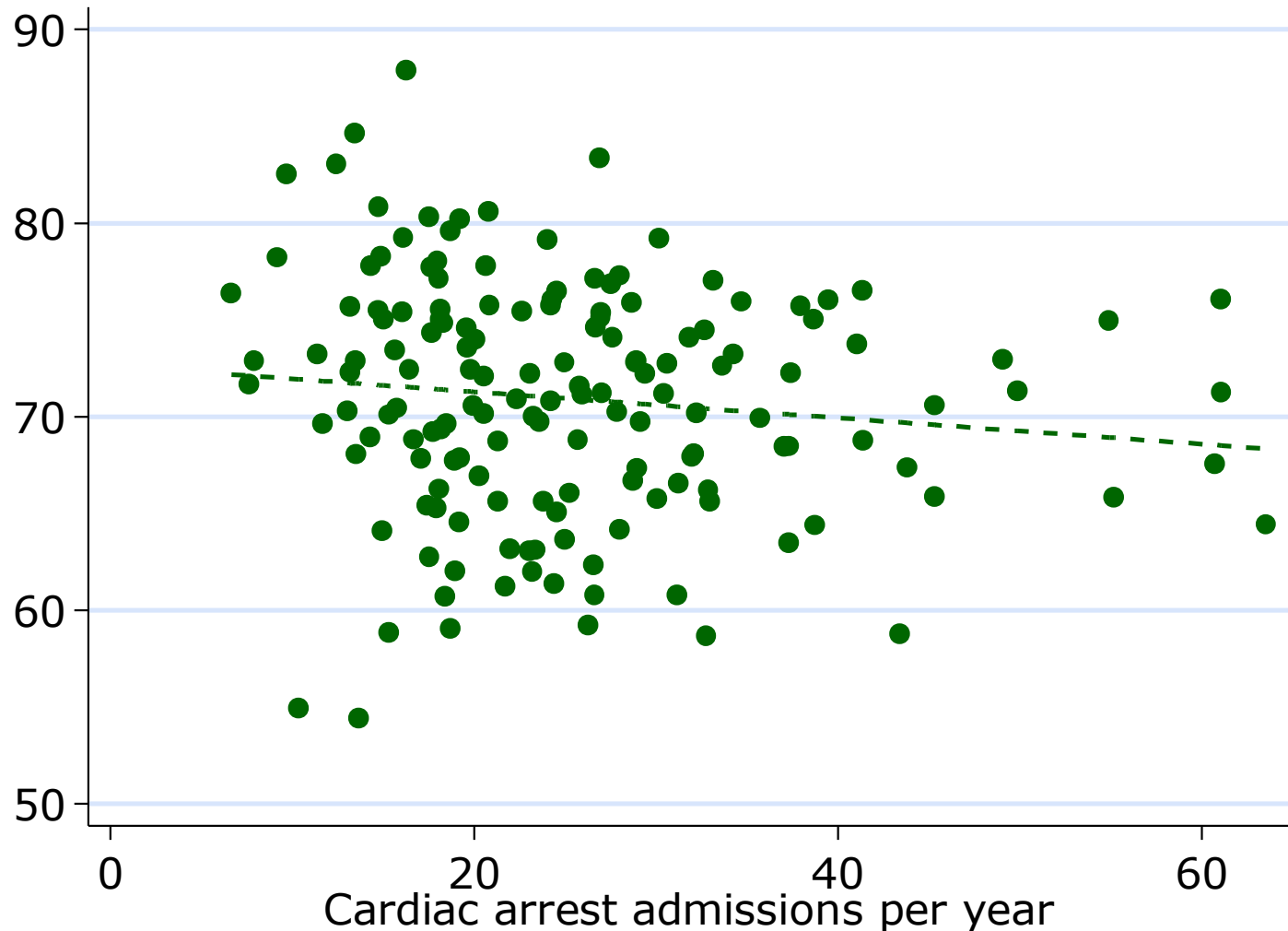
A Policy Statement From the American Heart Association

- PCI 24 h, hypothermia, IABP, EEG and prognostication
 - Bobrow BJ. Curr Opin Crit Care 2009;15:221-7
- Transport interval after ROSC does not influence survival
 - Spaite DW. Ann Emerg Med 2009
- Outcome better with > 50 ICU cases/yr
 - Carr BG. Resuscitation 2009;80:30-4

Nichol G. Circulation 2010;121:709-29

Volume – outcome relationship?

ICNARC data





Post cardiac arrest patient in ICU

Summary

- 30% survive to discharge in UK ITUs
- Post cardiac arrest syndrome
- Heart – early revascularisation, circulatory support
- Brain – hypothermia, glucose control?
- Prognostication - problematic



National Cardiac Arrest Audit (NCAA)

- Secure, web-based data entry
- Validation, on data entry and centrally
- Online comparative reporting
- Initial scope- Individual receiving chest compressions and/or defibrillation from the hospital based resuscitation team (or equivalent)

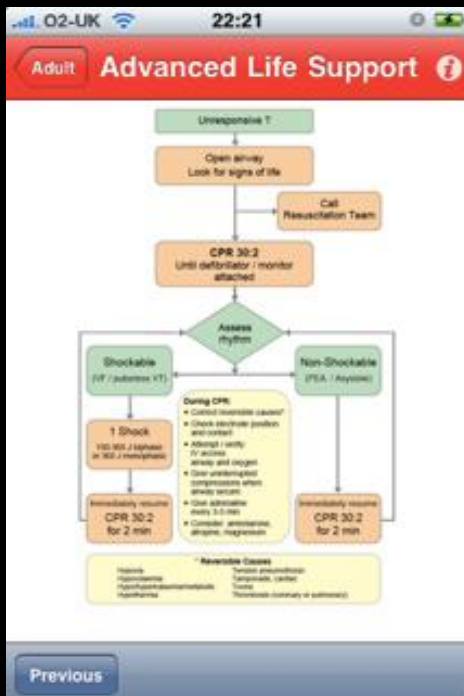
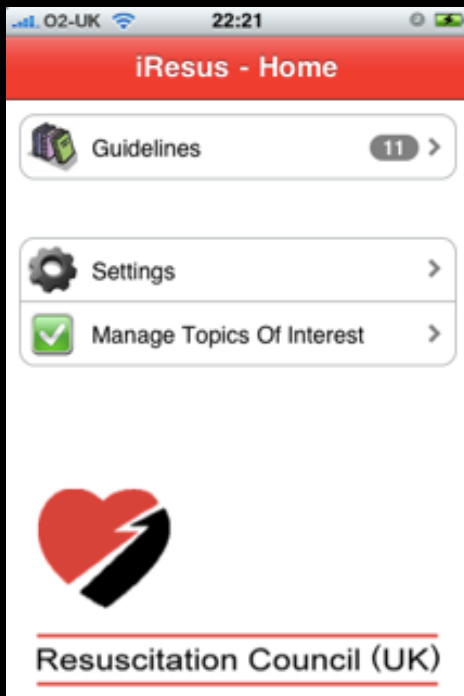


Small, modular dataset

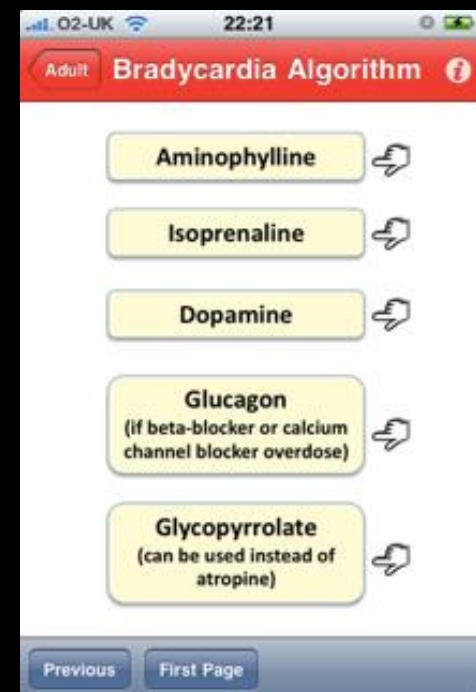
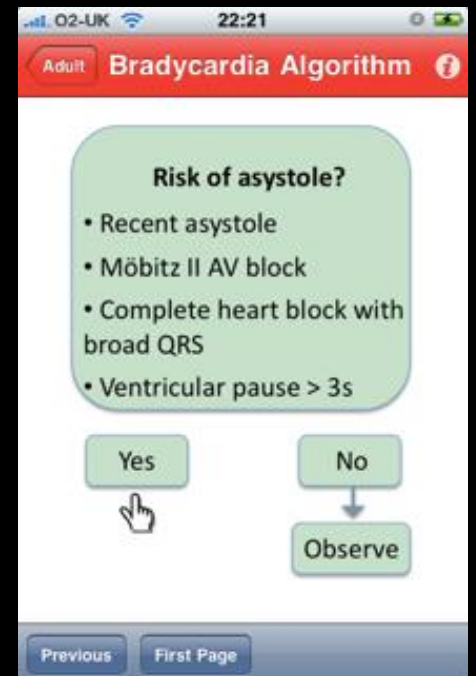
To take part, or to find out more about this audit, contact the NCAA team on ncaa@icnarc.org

Supported by: Resuscitation Council (UK) and ICNARC





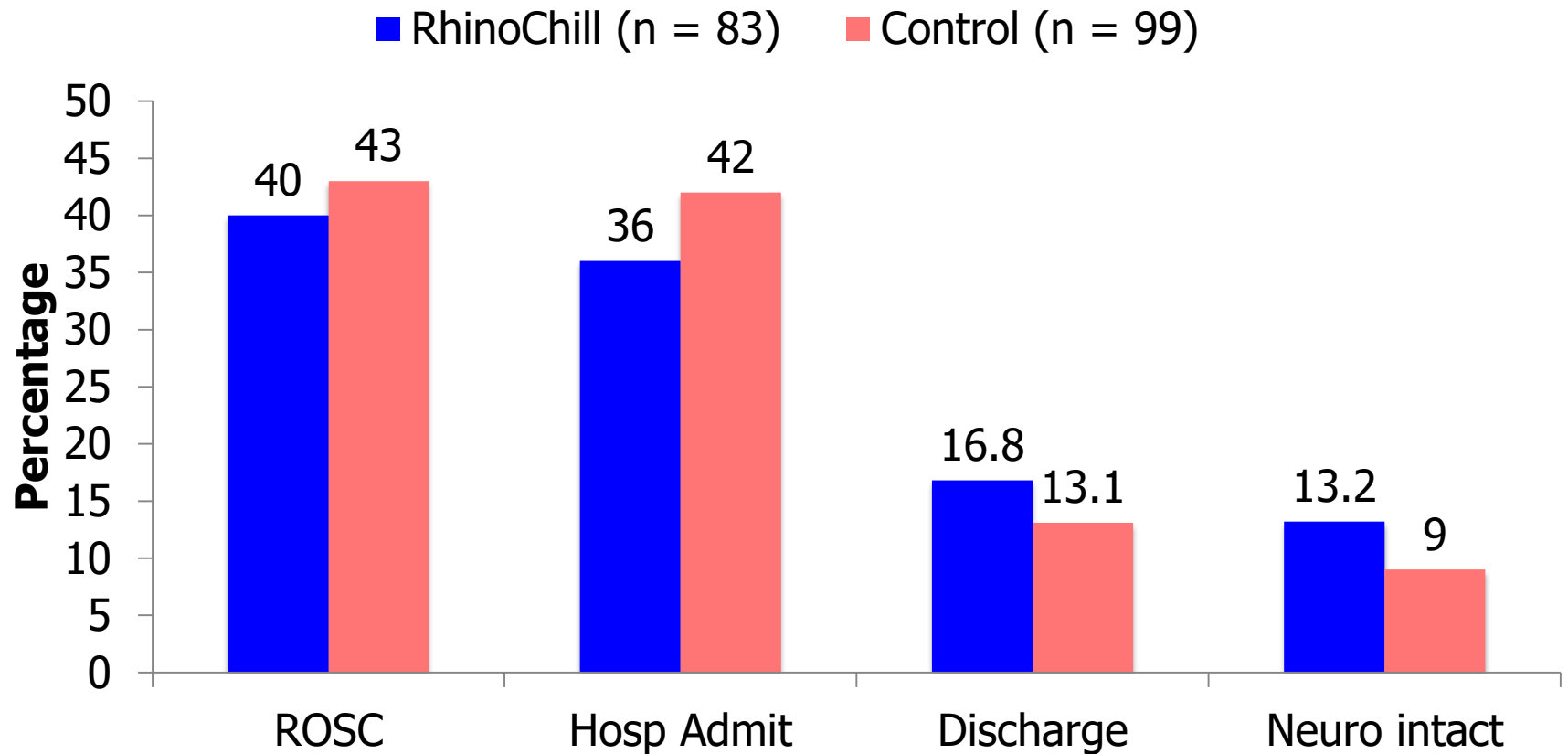
iResus



The PRINCE Trial (Pre-ROSC Intra-Nasal Cooling Effectiveness)

- Intra-arrest cooling
 - RhinoChill (n = 83)
 - Control (n = 99)
- Tympanic temp of 34°C reached 3 h sooner with RhinoChill

The PRINCE Trial

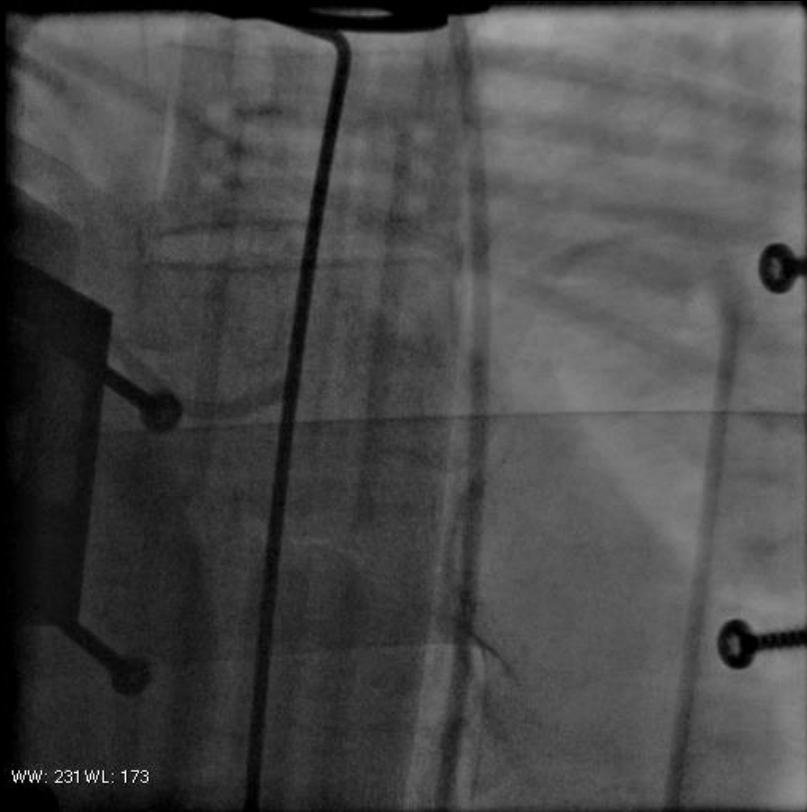


Mechanical CPR to facilitate PCI during cardiac arrest



- LUCAS in 43 patients over 5-year period
- 17 (40%) survived the procedure
- 11 (26%) discharged neurologically good

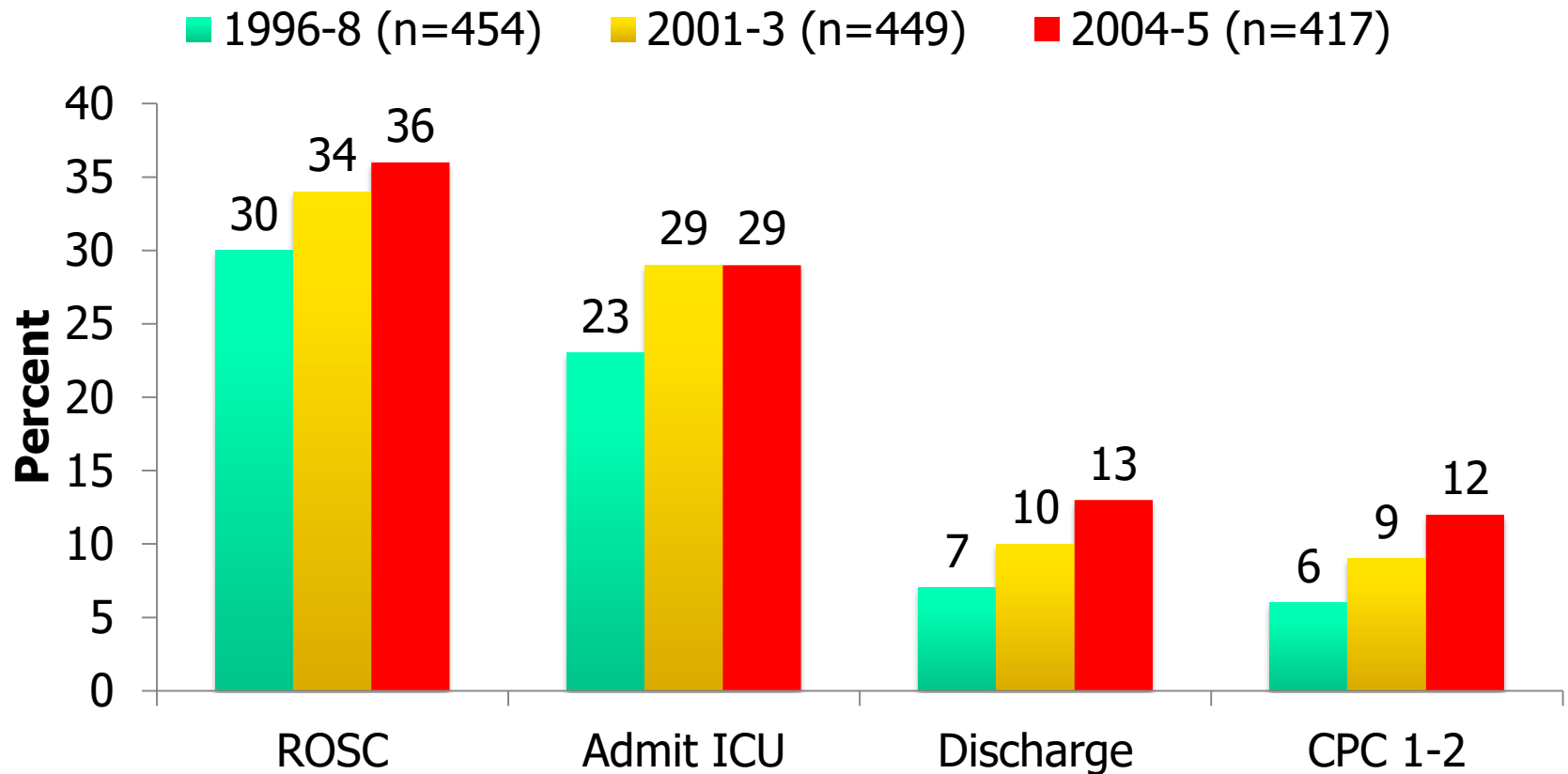
Mechanical CPR to facilitate PCI during cardiac arrest



Courtesy of Sunde K, Oslo



Survival after out-of-hospital cardiac arrest – Oslo, Norway



Hypothermia:

Mechanism of action?

- Suppression of free radicals
- Blocking pathological protease cascades
- Suppression of apoptosis (48 h)
- Suppression of pro-inflammatory cytokines (5 days)
 - No? Callaway CW. Crit Care Med 2008;36:2607-12

External techniques: overcooling

- Retrospective review of 32 cases
- Surface cooled to target of 32-34°C
 - 20/32 (63%) < 32°C
 - 9/32 (28%) < 31°C
 - 4/32 (13%) < 30°C
- Rebound hyperthermia (>38°C) at 12-18 h after rewarm in 7/32 (22%)

Implementation of standardised post-resuscitation care after OHCA

Sunde K. Resuscitation 2007;73:29-39	Control (n=58) 1996 - 8
Reperfusion n (%)	2 (3)
Hypothermia	0
Inotropes	29 (50)
Balloon Pump	0
Insulin	4 (7)
Survival (CPC 1-2)	15 (26)

Implementation of standardised post-resuscitation care after OHCA

Sunde K. Resuscitation 2007;73:29-39	Control (n=58) 1996 - 8	Standard (n=61) 2003 - 5
Reperfusion n (%)	2 (3)	30 (49)
Hypothermia	0	40 (66)
Inotropes	29 (50)	43 (80)
Balloon Pump	0	8 (15)
Insulin	4 (7)	27 (44)
Survival (CPC 1-2)	15 (26)	34 (56)

ICU Volume and outcome

ICU cases/year	OR survival	95% CI	p value
<20	1.00	-	-
20-34	0.78	0.55 – 1.11	0.16
35-50	0.71	0.45 – 1.11	0.13
> 50	0.62	0.45 – 0.86	0.01

Carr BG. Resuscitation 2009; 80: 30-34.

Oxygenation: Treatment recommendation



Standards for the
Management of Patients
After Cardiac Arrest

STANDARDS AND GUIDELINES

Adjust the FiO_2 to
achieve an arterial
oxygenation 94-98%