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DALL'EVIDENZA AI TRATTAMENTI FUTURI CENTRO CONGRESSI MAGAZZINI DEL COTONE PORTO ANTICO GENOVA

# Post Rianimazione BUNDLES DI TRATTAMENTO NEL BAMBINO





### Genova 14 Ottobre 2017



# Conflitti d'Interesse (CI)











# CI

8 Posti Letto 500 ricoveri x Anno NIMV -HFOV - NO 152 codici rossi/anno 15.000 accessi vs 100.000













CI

# Formazione Simulazione















# Post Rianimazione BUNDLES DI TRATTAMENTO NEL BAMBINO





Pre-Arrest	Cardiac Arrest	CPR	Post-Arrest stabilization
	No Flow	Low Flow	Low, Normal or High Flow
PROTECTION     Rapid Recognition     Call for Help	PRESERVATION  • Prompt CC • Defibrillate if VE	RESUSCITATION  • Push hard, Push Fast • Minimize interruption •Full recoil	•RESUSCITATION /REGENERATION • Temperature control • Blood pressure • Oxygen titration • 2 Cardiac Catheterization (PCI)
<ul> <li>Response Team</li> <li>Oxygen/Ventilation</li> </ul>	• Demormate in vi-	Assist ventilation?         Vasopressors?         Cooling?	<ul> <li>Glucose</li> <li>Ventilation (CO2)</li> <li>Seizure Control</li> <li>Goal Directed Care</li> </ul>



European Resuscitation Council and European Society of Intensive Care Medicine Guidelines for Post-resuscitation Care 2015 Section 5 of the European Resuscitation Council Guidelines for Resuscitation 2015<sup>\*</sup>

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# CoSTR= Consenso sulla Scienza con Raccomandazioni di Trattamento

- 1. Pico question su argomento
- 2. Revisione letteratura
- 3. Livello di evidenza
- 4. Consensus

of evidence

- 5. Forza della raccomandazione
- 6. Motivazione

Raccomandazione debole o forte



# **Expert Opinion**







## **RICERCA x FORMAZIONE x ATTUAZIONE**

# La "formula" della sopravvivenza





Case Report



Bambino di 5 anni annegato piscina comunale RCP prolungata sul posto dai bagnini e poi dal 118 Rosc dopo oltre 30 min Rcp ed adrenalina Con ritmo non defibrillabile Trasferito presso nostra terapia Intensiva Intubato e Ventilato



All' ammissione in PICU il bambino appare NON responsivo (poca sedazione) con I seguenti parametri vitali

✓ RR (assistita) 30 x min. ✓ O2 sat. 100 ✓ EtCO2 25 mm Hg ✓HR 140 ✓ BP 70/40 √T 33°C







# Trattamento post-rianimatorio

L'obiettivo è di ristabilire:

- Una funzione cerebrale normale
- Un ritmo cardiaco stabile
- Un' adeguata perfusione d' organo
- Un' adeguata qualità di vita



## Pediatric Post-Cardiac Arrest Syndrome

% patients	IHCA Non-cv	IHCA CV- postop	IHCA CV-nonsurg	IHCA PICU CV-postop	ОНСА
	NRCPR LO, 2011	NRCPR LO, 2011	NRCPR VN, 2006	HSC NdM, 2006	ROC DA, 2009
	N=1109	N=640	N=306	N=91	N=624
(%) Return of spontaneous circulation (ROSC)	53	67	53	82	10
(%) 24 hr survival	37	60	42	66	
(%) Survival to Hospital DC	23	37	28	25	6.4

Nadkarni, JAMA, 2006; De Mos, CCM, 2006; Ortmann, Circulation, 2011; Atkins, Circulation, 2009





Improving Survival and Neurological Function for Younger Age Groups After Out-of-Hospital Cardiac Arrest in Sweden:

### A 20 Year Comparison



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### Pediatric IHCA Survival to Hospital D/C Girotra, Circ Cardiovasc Qual Outcomes, 2013 45 40 35 30 Risk Adjusted 25 Survival (%) 20 15 10 5 0 200 2001 2002 2003 2004 2005 2006 2001 2008 2009

Survival trends *not* accompanied by higher rates of neurological disability among survivors over time (unadjusted *P for trend=0.32*)



✓ RR (assistita) 30 min. ✓O2 saturation 100% ✓EtCO2 25 mm Hg **√HR** 140 ✓ BP 70/40 ✓T 33°C







## **Post-ROSC oxygenation**

Multiple animal studies:

- ventilation with 100% oxygen during and following resuscitation contributes to free radical-mediated reperfusion injury to the brain.
- may be associated with more neurologic deficit than ventilation with room air, especially when high Pa02 is experienced in the first hour post-ROSC





## **Post-ROSC oxygenation**

What about neonatal studies?

• Two meta-analyses of several randomized controlled trials contro

2004; Rabi, Resuscitation, 2007

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	Pa02>	300	Pa02 60	-300		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI
Del Castillo, 2012	10	19	59	145	100.0%	1.29 [0.81, 2.07]	<b>_</b>
Total (95% CI)		19		145	100.0%	1.29 [0.81, 2.07]	( )
Total events	10		59				$\checkmark$
Heterogeneity: Not ap	plicable						
Test for overall effect:	Z = 1.07	7 (P = 0)	.28)				Favours Pa02>300 Favours Pa02 60-300

	Pa02>	300	Pa02 60	-300		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI
Ferguson, 2012	91	207	478	1220	100.0%	1.12 [0.95, 1.33]	
Total (95% CI) Total events	91	207	478	1220	100.0%	1.12 [0.95, 1.33]	$\bigcirc$
Heterogeneity: Not ap Test for overall effect:	plicable Z = 1.34	(P = 0	).18)				0.01 0.1 1 10 100 Favours Pa02>300 Favours Pa02 60-300

	Pa02>	300	Pa02 60	-300		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI
Guerra, 2013	10	34	7	30	100.0%	1.26 [0.55, 2.90]	
Total (95% CI)		34		30	100.0%	1.26 [0.55, 2.90]	
Total events	10		7				
Heterogeneity: Not app	plicable						
Test for overall effect:	Z = 0.55	(P = 0)	.59)				Favours Pa02>300 Favours Pa02 60-300

	Pa02>	300	Pa02 60	-300		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI
Bennett, 2013	57	87	37	66	100.0%	1.17 [0.90, 1.52]	
		07			100.0%	1 17 (0 00 1 53)	
Total (95% CI)		87		66	100.0%	1.17 [0.90, 1.52]	
Total events	57		37				
Heterogeneity: Not app	plicable						
Test for overall effect:	Z = 1.16	$\delta (P = 0)$	.24)				Favours Pa02>300 Favours Pa02 60-300

Pediatric Post-CA Hyperoxia
<ul> <li>195 pediatric cardiac arrests: survive &gt; 6 h</li> <li>54% hyperoxia, 22% hypoxia</li> <li>No relationship with survival Statler, CCM 2013</li> </ul>
<ul> <li>74 ped CA: 51% hyperoxia; 14% hypoxia No relationship with survival Guerra-Wallace, PCCM 2013</li> </ul>
<ul> <li>PICANET: n= 1875 : 11% hyperoxia, 24% hypoxia</li> <li>Odds of death 1.25 (95% CI: 1.17-1.37)</li> </ul>
Ferguson, Circulation 2012



Post–Cardiac Arrest Oxygenation

 It may be reasonable for rescuers to target normoxemia after ROSC (Class IIb, LOE B-NR).

 Arterial oxyhemoglobin saturation of 100% may correspond to a PaO2 anywhere between 80 and approximately 500mmHg (target 94-99)

 may be reasonable (when the necessary equipment is available) for rescuers to wean oxygen. Ideally, oxygen is titrated to a value appropriate to the specific patient condition.





✓ RR (assistita) 30 min. ✓O2 saturation 96 % ✓ EtCO2 25 mm Hg **√HR** 140 ✓ BP 70/40 ✓T 33°C







## Hyperventilation post-ROSC

- Inadvertent hyperventilation is common
  - Proportion of CPR epochs with ventilation rates exceeding AHA guidelines (>10 bpm) was 63% (CI95 59–67%)
- Hyperventilation (increased RR or TV):
  - Hypocarbia shifts oxyhemoglobin dissociation curve
  - Reduce preload to heart
  - Cerebral vasoconstriction
- Systemic acidosis is common post-ROSC
  - usually metabolic not respiratory

McInnes, Resuscitation, 2011





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## Pediatric Post-arrest PaCO2 Management



Cerebrovascular reactivity to changes in arterial carbon dioxide tension is preserved in comatose patients post-ROSC Buunk Stroke 1997





### Post–Cardiac Arrest PaCO2

- It is reasonable for practitioners to target a PaCo2 after ROSC that is appropriate to the specific patient condition, and limit exposure to severe hypercapnia or hypocapnia
- (Class IIb, LOE C-LD).



- Monitor exhaled Co2 (ETCO2) especially during transport and diagnostic procedures.
- (Class IIa, LOE B).



✓ RR (assistita) 30 min.  $\checkmark$  O2 saturation 96% ✓ EtCO2 34 mm Hg **√HR** 140 ✓ BP 70/40 √T. 33°C







Post-Cardiac Arrest: Myocardial Dysfunction

## Animal models (VF)



- Myocardial stunning, with an onset within 30 min post-ROSC, and function returning to normal within 24-48 hours of the insult
- Systolic and diastolic dysfunction (biventricular)

Kern, Circulation, 1997

Kern, J Am Coll Card, 1996



# LV Systolic Function Following OHCA

- 58 patients admitted to CHOP PICU
- Echocardiogram within 24 hours of ROSC
- Decreased function vs Normal Function
- Only SVO2< 60 differentiated ECHO findings</li>

After controlling for VF and vasopressor scores, patients with decreased function had 13.7 times higher odds or death



Conlon, submitted



### Myocardial Function is Reversibly Depressed for 24 to 72 hours after Cardiac Arrest



Post-cardiac arrest myocardial dysfunction is responsive to inotropes and intravenous fluid



# **Post-CA Hypotension**

- Project IMPACT: 8736 patients
- Hypotension (ICU SBP<90 within 1 hr): 47%

### **Mortality**

- Hypotension 65%
- No hypotension37%
- Odds of death 2.7 (95% CI: 2.5-3.0)

Trzeciak, CCM 2009



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## Pediatric Post-Arrest I CU Hypotension

- PECARN: 383 children
- Hypotension (SBP<5%ile within 6 hr): 56%

### **Mortality**

- Hypotension 53%
- No hypotension 41%
- Odds of death: 1.71 (95% CI: 1.02, 2.89)

Topjian CCM, 2014



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Post–Cardiac Arrest Fluids and Inotropes

Pediatric Post-ROSC Hypotension Topjian, CCM, 2014

After controlling for patient and cardiopulmonary arrest characteristics, hypotension in the first 6 hours following ROSC was associated with:

Significantly increased odds of in-hospital mortality

adjusted odds ratio = 1.71; p = 0.042

Increased odds of unfavorable outcome

adjusted odds ratio = 1.83; *p* = 0.032.



## Post–Cardiac Arrest Fluids and Inotropes

- After ROSC, we recommend that parenteral fluids and/or inotropes or vasoactive drugs be used to maintain a systolic blood pressure greater than 5th percentile for age (Class I, LOE C-LD).
- When appropriate resources are available, continuous arterial pressure monitoring is recommended to identify and treat hypotension (Class I,LOE C-EO).





### Post–Cardiac Arrest Fluids and Inotropes

 It is reasonable to use an inodilator in a highly monitorated setting for tratment of myocardial dysfuction with increased systemic or pulmonary vascular resistence (Class IIa, LOE B).





## **BRAIN AUTOREGULATION**





## Post-resuscitation fluids and inotropic use Examples of TCD Profiles (n=18 patients)

Sundgreen, Stroke, 2001



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# **Boundles of Brain Management**



- Oxygen Titration
- BP Management
- Cerebral perfusion
- Temperature management
- Control of seizures
- Glucose control



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# All' ammissione in PICU il bambino appare NON responsivo (poca sedazione) con i seguenti parametri vitali

✓ RR (assisted) 30 ✓O2 saturation 96% ✓EtCO2 34 mm Hg **√HR** 120 ✓ BP 90/45 √T 33°C





### **Casistica Hypotermia post CA (GEMELLI)**

	ALL (n.33)	Survivors (n.21) <mark>63%</mark>	Dead (n.12) <mark>37%</mark>	р
Age (mths, median; IQR)	36 (3.55;177.5)	48	35	<mark>0.</mark> 846
Gend <mark>er (M/F)</mark>	1.35	1.33	1.4	0.946
CA e <mark>tiolog</mark> y (Tab. )	42.4 airway obstruction	42.8 airway obstruction	41.6 airway obstruction	0.1 <mark>6</mark> 2
IN v <mark>s OUT</mark> of hospital(%)	13/20	11/13 IN 10/20 OUT	2/13 IN 10/20 OUT	0.029
Earl <mark>y brain</mark> oedema score	2 (0.5-3.0)	1 (0; <b>2</b> .5)	3 (2.25;3)	0.013
Gap (b <mark>efore T</mark> H), median, hrs	2.5 (2.0;4.0)	2 (2 <mark>;3.5</mark> )	4 (2;5)	0.243
CPR leng <mark>ht – tim</mark> e to ROSC (min)	25 (17.5-35.0)	20 (10;35)	28.5 (25;45)	0.006
Blood pH 24hrs/48hrs	7.369/7.392	7.361/7.375	7.405/7.413	0.068 (24hrs) 0.042(48hrs)
PrismIII <sub>24</sub>	32 (25;45)	32 (22;39.8)	44.5 (31;48.3)	0.0019
VIS <sub>max</sub>	30 (21;37.5)	25 (15;30)	37.5 (31;43.75)	0.0020





Cardiac Arrest in Children

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## **Temperature Management**

 For infants and children remaining comatose after OHCA, it is reasonable either to maintain 5 days of continuous normothermia (36°C to 37.5°C) or to maintain 2 days of initial continuous hypothermia (32°C to 34°C) followed by 3 days of continuous normothermia (Class IIa, LOE B-R)

For infants and children
 remaining comatose after IHCA
 there is insufficient evidence



to recommend cooling over normothermia.



## **Temperature Management**

 Continuous measurement of temperature is recommended (Class I, LOE B-NR)

 Fever (temperature 38°C or higher) should be aggressively treated after ROSC (Class I, LOE B-NR)





# **Questions about hypothermia**

Who should be cooled?
 When to cool?



Slide modified from T. van den Hoek, MD, U Chicago

Dopo 12 ore dal ricovero in PICU negli ultimi 20 minuti si sono verificati ripetuti sbalzi della Pa, FC e Sat O2. Si sospetta la presenza di crisi convulsive senza la presenza di movimenti tonico-clonico associati





Electroencephalographic monitoring during hypothermia after pediatric cardiac arrest Abend Neurology 2009

- 19 children had continuous EEG monitoring during TTM
- Electrographic seizures occurred in 47%, and 32% of all patients developed status epilepticus
- Seizures were nonconvulsive in 67% and electrographically generalized in 78%
- Seizures commenced during the late hypothermic or rewarming periods









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	MORTALITY <u>OR (95% CI)</u>	↑ PCPC <u>OR (95% CI)</u>
No Seizures	ref	ref
Electrographic	1.3 (0.3, 5.1)	1.2 (0.4, 3.9)
Seizures	p=0.74	P=0.77
Electrographic	5.1 (1.4, 18)	17.3 (3.7, 80)
Status Epilepticus	p=0.01	P<0.001



## **Post-ROSC Seizure Control and Prevention**

- Aggressive anticonvulsant treatment (and prophylaxis) makes sense
  - to date no data exists to suggest that doing so alters outcome post-ROSC (in adults or children)
- Choice of treatment approach is personal and not directly evidence-based
- Caution needs to be paid to the hemodynamic consequences of drug choice for seizure control



 Continuous EEG monitoring should be considered post-ROSC, regardless of whether or not the patient is going to require paralytics



# **EEG Monitoring**



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Figure. Phases of post-cardiac arrest syndrome.

Neumar, Circulation, 2006

### Priorities post-ROSC?

 Prevent on-going primary injury (eg. Sepsis); (multi-system)

•Minimize (secondary) post-ROSC injury; (multi-system)

Prognostication

•Allow for family centered care







Gold, Resuscitation, 2013





I Genitori chiedono di avere un colloquio e capire quali sono i rischi neurologici del bambino. L' infermiera riferisce che fra di loro hanno discusso riguardo all' intesinvità delle cure



### **Predictive Factors After Cardiac Arrest**



- EEGs performed within the first 7 days after pediatric cardiac arrest may be considered in prognosticating neurologic outcome at the time of hospital discharge (Class IIb, LOE C-LD) but should not be used as the sole criterion.
- The reliability of any 1 variable for prognostication in children after cardiac arrest has not been established. Practitioners should consider multiple factors when predicting outcomes in infants and children who achieve ROSC after cardiac arrest (Class I, LOE C-LD).





Figure. Phases of post-cardiac arrest syndrome.



### Allow for family centered care







# Sistemi Complessi





# **Formula-One for Survival!**



# **Pit Stop Mentality**

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# Domains of competencies





## Data Driven Debriefing in the Clinical Setting

- Adults
  - Improved ROSC from 44 to 59% (Edelson, 2008)





# Improving future performance

It is reasonable for in-hospital systems of care to implement performance-focused debriefing of rescuers after IHCA in both adults and children (Class IIa, LOE C-LD).





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# Riassunto

- La sindrome post-arresto è complessa
- La qualità del trattamento post-rianimatorio influenza l'outcome finale
- Monitoraggio adeguato, trasferimento in sicurezza e ottimizzazione continua delle funzioni d'organo
- Difficile formulare una prognosi







Presidente del Congresso: Gina Ancora

Presidente AMIETIP: Luca Tortorolo

Congresso Nazionale Gioved 23 novembre

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Mos Presidents Marca De Luna Angelo Antgot Lata McKapia





# Coast to Coast Arrivederci a Rimini



