Therapeutic Hypothermia: HOW TO CHILL OUT Terrence Coulter MD, FCCM, FCCP Department of Pulmonary, Critical Care, and Sleep Medicine Ferrell-Duncan Clinic Medical Director, Department of Respiratory Therapy and Critical Care Cox Health System Springfield, MO

Disclosures • None

Therapeutic
Hypothermia

Targeted
Temperature
Management
(TTM)

Learning Objectives

- Evolution of therapeutic hypothermia
- Use in ACLS guidelines
- Methods of cooling
- Various temperature management
- What to watch clinically during hypothermia procedure
- Uses beyond cardiac arrest

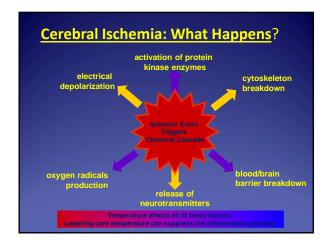


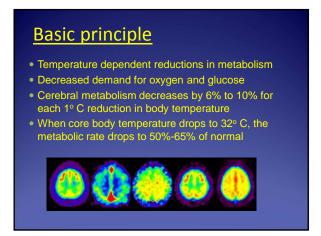
Cardiac Arrest

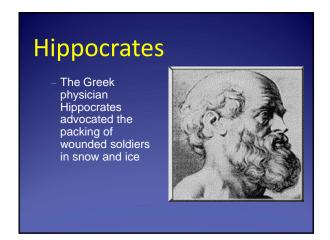
- 450,000 Americans/year
- 80% occur at home
- Rate of death is 90%
- In-hospital arrests have slightly better outcomes
 - Restoration of circulation in 44%
 - Survival to discharge in 17%

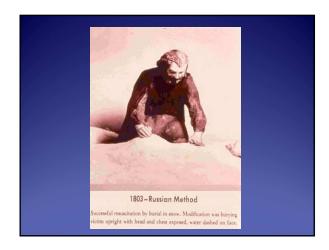
Young, G. Neurologic Prognosis after Cardiac Arrest. N Engl J Med 2009, 361:605-11.

"Chain of Survival" "Chain of Survival" "Chain of Survival" "Early Early Defibrillation Advanced Gare All of these advances have helped to achieve return of spontaneous circulation (ROSC) Prompt Early Early ACLS Access CPR Defib Care It still has not increased survivability or improved neurological outcomes

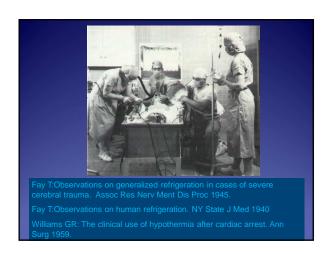






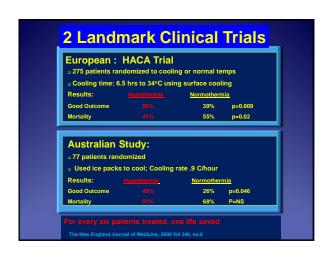






Shortcomings of initial trials Believed needed large reduction in temperature to achieve efficacy Most treated with deep hypothermia (<30° C) Core temperature difficult to maintain Duration of cooling ranged from 2-10 days Rewarming methods not reliable Intensive care facilities not widely available

Back again Positive studies in the early 80's Realized outcomes could be improved with mild to moderate hypothermia (31-35° C) Advancements in technology Evolution of critical care units Understanding and anticipating side effects of cooling



Box. Glasgow-Pittsburgh Cerebral Performance Categories* 1 Good Cerebral Performance HOME Conscious: Alert, able to work and lead a normal life. May have minor psychological or neurological deficits (mild dysphasia, nonincapacitating hemiparesis, or minor cranial nerve abnormalities). 2 [Moderate Cerebral Disability | REHAB | Conscious. Sufficient cerebral function for part-time work in sheltered environment or independent activities of daily life (dressing, traveling by public transportation, and preparing food). May have hemiplegia, seizures, ataxia, dsysarthria, dysphasia, or permanent memory or mental changes. 3 [Severe Cerebral Disability | LONG TERM FACILITY | Conscious. Dependent on others for daily support because of impaired brain function (in an institution or at home with exceptional family effort). At least limited cognition. Includes a wide range of cerebral abnormalities from ambulatory with severe memory disturbance or dementia precluding independent existence to paralytic and able to communicate only with eyes, as in the locked-in syndrome. 4 [Coma, Vegetative State] | LONG TERM FACILITY | Not conscious. Unaware of surroundings, no cognition. No verbal or psychological interactions with environment. 5 [Death] | SIX FEET UNDER | Certified brain dead or dead by traditional criteria. *Adapted with permission from Cummings et al. 13

What are the current S cardiac arrest? (2005 Evidence-based	1
>32-34	/VF Cardiac arrest 4 degrees 24 hours
American Heart Association AHA Recommendations	International Liaison Committee on Resuscitation ILCOR Recommendations
Class Ma: Unconscious adult patients with return of spontaneous circulation (ROSC) after out-of hospital cardiac arrest should be cooled to 32°C to 34°C (89.6°F to 93.2°F) for 12 to 24 hours when the initial rhythm was ventricular fibrillation (VF). Class Mb: Similar therapy may be beneficial for natients with non-VF arrest out of hospital or for	Level I evidence > 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 riythm was ventricular fibrillation (VF). > Such cooling may also be beneficial for other rhythms or in-hospital cardiac arrest.

Part 8: Post-Cardiac Arrest Care
2015 American Heart Association Guidelines Update for
Cardiopulmonary Resuscitation and Emergency Cardiovascular
Care
-Induce hypothermia for unconscious adult patients with return
of spontaneous circulation (ROSC) after out-of-hospital cardiac
arrest when the initial rhythm was ventricular fibrillation (VF) or
pulseless ventricular tachycardia (pVT) (class I, level of evidence:
B-R)
-Similar therapy may be beneficial for patients with non-VF/nonpVT (nonshockable) arrest out-of-hospital or with in-hospital
arrest (class I, level of evidence: C-EO)
-The temperature should be maintained between 329C and 369C
(class I, level of evidence: B-R)
-It is reasonable to maintain TTM for at least 24 hours (class IIa,

The 2015 American Heart Association (AHA) guidelines:

- Routine prehospital cooling of patients with ROSC with intravenous (IV) rapid infusion is not advised (class III: no benefit; level of evidence A)
- It is reasonable to prevent fever in comatose patients after TTM (class IIb, level of evidence C-LD)
- Hemodynamically stable patients with spontaneous mild hypothermia (>33°C) after resuscitation from cardiac arrest should not be actively rewarmed

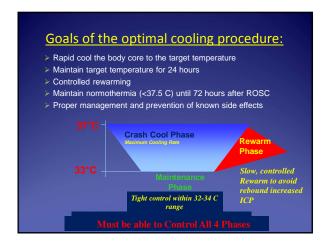
Inclusion criteria:



- Age greater than 18 years
- Intubated patients with treatment initiated within 6 hours after cardiac arrest (nonperfusing VT or VF)
- Patients able to maintain a systolic blood pressure above 90 mm Hg, with or without pressors, after cardiopulmonary resuscitation (CPR)
- Patients in a coma at the time of cooling

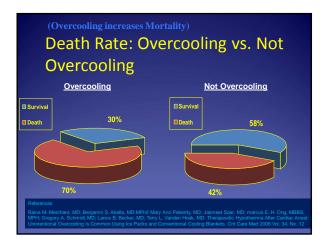
Exclusion criteria:

- The patient can follow verbal commands
- Recent major surgery within 14 days Possible risk for infection and bleeding
- Systemic infection/sepsis Small increase in risk of infection
- Coma from other causes (drug intoxication, preexisting coma prior to arrest)
- Known bleeding diathesis or with active ongoing bleeding -Hypothermia may impair the clotting system (however, patients may receive chemical thrombolysis, antiplatelet agents, or anticoagulants if deemed necessary in the treatment of the primary cardiac condition)
- In addition, hypothermia is inappropriate in patients with a valid do not resuscitate order (DNR).





Difficult to achieve target temperature 14% never achieved target and 70% of patients required additional cooling methods (the HACA study)* Difficult to maintain target temperature— overshoot 363% of chance of overcooling when using surface cooling methods** HACA, New Eng J Med, 2002 **Rana M. Merchani, MD. Benjamin S. Aballa, MD MPHJ Mery, Ann Peberdy, MD. Jaemeet Soar, MD; marcus E. H. Ong, MBBS, MPH, Grappy A. Schmidt, MD, Lance B. Backer, MD, Terry L. Vanden Hosk, MD. Therapeude Hypothemia Alter Cardiac Arnast Unintentional Overcooling is Common Using Ice Packs and Conventional Cooling Blankers. Crit Care Med 2006 Vol. 34, No. 12 **Mayer, Neuroscit Care, 2005



Limitations of Surface Cooling Difficult to achieve target temperature 14% never achieved target and 70% of patients required additional cooling methods (the HACA study)* Difficult to maintain target temperature— overshoot a 63% of chance of overcooling when using surface cooling methods** Uncontrolled rewarming— rebound ICP and hyperthermia Hindered access to patient (i.e during PCI procedure) Very nurse "Labor" intensive: often 2:1 ratio (nurses/pt) Skin monitoring and care required Increased shivering and paralytic usage **** HACA, New Erg J Med, 2002 "Raran M. Merchart, MD, Benjamin S. Abella, MD MPHI Mary Ann Peberdy, MD; Jasmeet Sox, MD; marcus E. H. Ong. MBBS, MPH, Gegop; A. Schnidt, MD; Larce B. Becker, MD; Terry L. Vanden Hoek, MD: Therapeudic Hypothemia After Cardiac Arrest: Unitereficial Overcoding is Common Using Ice Placks and Conventional Cooling Blankets. Crit Care Med 2000 Md. 34, No. 12 "Mayer, Namerri Care. 2005

Cooling the Patient's Skin Skin is a temperature receptor for brain Brain/Hypothalamus tries to maintain homeothermia Therefore, cold applied to skin sends "cold" stress message to hypothalamus: Triggers shivering → generates heat Causes vasoconstriction→stops heat loss May cause thermal damage

Shivering is the Enemy

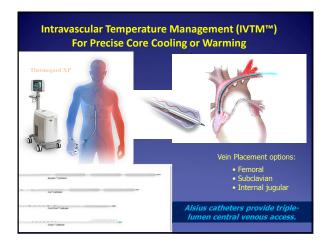
- Increased oxygen consumption
- Metabolic rate up to 600%
- Fights the cooling process
- Increasing Heart rate (HR), Respiratory rates (RR), Intracranial pressure (ICP)
- Result in hypoxemia, myocardial ischemia
- Use of high doses of sedation and paralytics
 - Hypotension ischemia
 - Cardiac depression AMI
 - Prolong on ventilator ARDS and Ventilator Associated Pneumonia
 - Reduced muscle tone is further associated with risk for skin breakdown

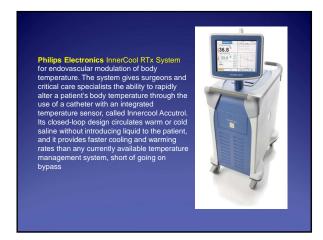
Cold IV Fluid

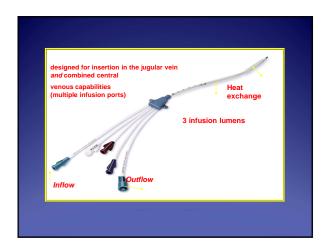
- Good way to get jump started
- Can be initiated pre-hospital, at ED or during transportation
- 30-40 cc/kg of 4°C NS
- Difficult to reach target temperature
- Difficult to maintain target temperature for 24 hours
- Limitation on the fluid quantities due to heart condition

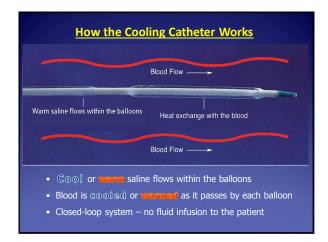
Kliegel A, et al: cold infusion alone are effective for induction of therapeutic hypothermia but do not keep patients cool

risk for

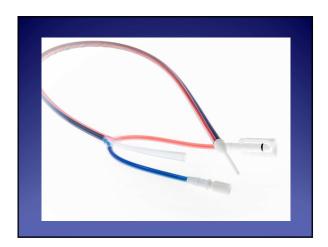




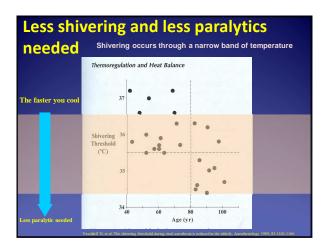


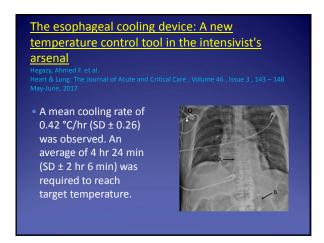


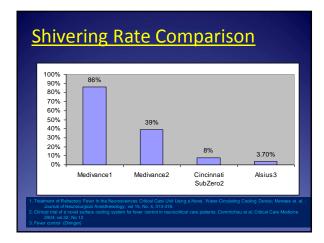




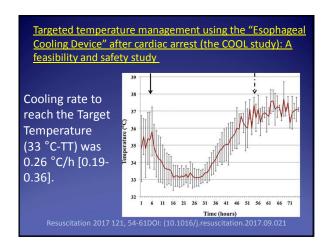








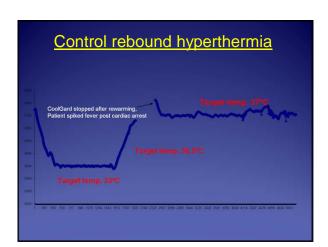
-Compatible with multiple chiller brands -Reduced risk of skin complications -Reduced risk of needle sticks -Reduced risk of CLABSI -Reduced risk of blood clots (DVT and PE) -Unobtrusive and maintains easy access to patient -Non-sterile placement procedure -Reduced shivering compared to surface devices -Rapid placement via single provider

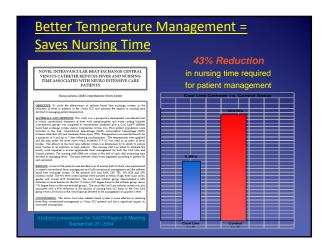


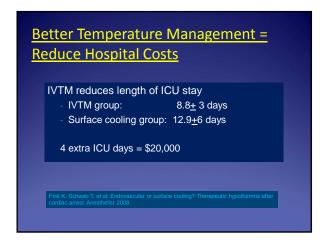
Targeted temperature management using the "Esophageal Cooling Device" after cardiac arrest (the COOL study): A feasibility and safety study

• Temperature deviation outside the TT during TTM-maintenance was 0.10 °C [0.03–0.20]. Time with deviation >1 °C was 0 h. Rewarming rate was 0.20 °C/h [0.18–0.22].

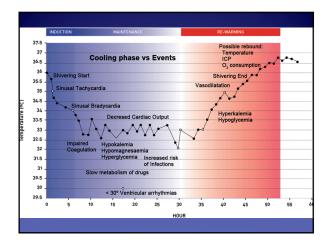
Resuscitation 2017 121, 54-61DOI: (10.1016/j.resuscitation.2017.09.021



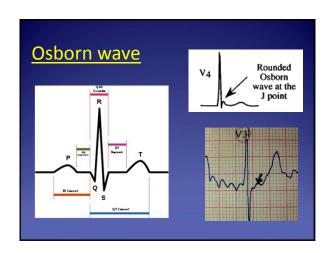




Potential complications of Therapeutic Hypothermia



What to Watch For Clinically During Therapeutic Hypothermia Hypothermia induction-maintenance phase Avoid shivering Increases the metabolic demands of the brain, foxygen consumption and decreases rate of cooling. Sedating and chemical paralyzing patient prior to cooling. Avoid hypovolemia Hypothermia-induced diuresis Avoid arrhythmia Bradycardia is common, transient, at early phase of cooling induction Do not let temperature go less than 32°C – lead to ventricular fibrillation



What to Watch For Clinically During Therapeutic **Hypothermia**

- Avoid hyperglycemia
 Hypothermia ↓ Insulin sensitivity and secretion
 - Tight glucose regulation with high dose of insulin to maintain at normal level
 - Finger stick is not reliable, draw blood from CVC or A line

Altered clearance of medications

- Clearance of various drugs
 Known drugs: muscle paralyzers, propofol, fentanyl, phenytoin, pentobarbital, verapamil, propanol and volatile anesthetics

Prevent electrolyte disorders
- Potassium shifts intra cellular, monitoring and correct
Potassium level

Ventilator management

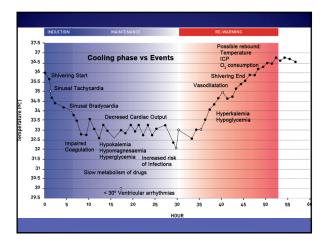
- When core body temperature drops to 32° C, the metabolic rate drops to 50%-65% of normal
- Oxygen consumption and CO2 production will decrease by the same percentage
- If vent settings left unchanged, then will lead to hyperventilation (resp alkalosis) which will cause cerebral vasoconstriction
- Need to adjust vent accordingly



Ventilator management

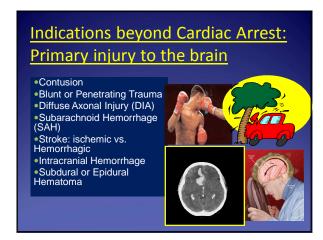
- Arterial line done ASAP
- Goal is normal pH
- Do not temperature correct ABG
- If can't get good finger pulse ox, try forehead
- Incidence of pneumonia is 30-50%
 - Usually due to aspiration at time of
 - Immunosuppressive effects of hypothermia
 - If suspected, give prophylactic antibiotics

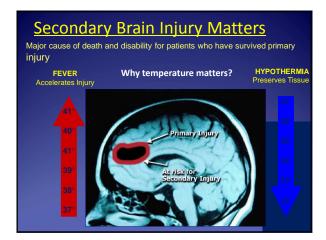




What to Watch For Clinically During Therapeutic Hypothermia	
Rewarming phase	
Slow and controlled rewarming (< 0.5°C/ hr)	
Avoid rebound hyperthermia and ↑ICP	
Keep normothermia at least 48 hours	
Avoid hyperkalemia	
Potassium shifts extra cellular	
 Potassium supplements should be stopped prior to rewarming 	
Avoid hypotension and hypovolemia due to vasodilation	
	Therapeutic Hypothermia Rewarming phase Slow and controlled rewarming (< 0.5°C/ hr) Avoid rebound hyperthermia and ↑ICP Keep normothermia at least 48 hours Avoid hyperkalemia •Potassium shifts extra cellular •Potassium supplements should be stopped prior to rewarming Avoid hypotension and hypovolemia due to

72 hour re-evaluation • Extubate • Continue supportive care • Withdrawal of life support





North American Brain Injury Study: Hypothermia Multicenter study of post-TBI hypothermia sponsored by the National Institutes of Health Hypothermia was induced to 33 degrees C buy 8 hours after insult and maintained to 48 hours Mortality was 28% with hypothermia and 27% with normothermia; poor outcomes equal at 57% Lack of Effect of Induction of Hypothermia after Acute Brain Injury Gyl. Citick M.D. Emmy R. Mine, Ph.D. R. M. Sung C. Cho, Ph.D., Herrey S. Lenn, Ph.D. Stephen McChauley, Ph.D. Kenneth R. Smith, Jr. M.D. J. Plan Murelaux, M.D. Ph.D. Franklin G. Wagner, Jr. M.D. Donald W. Merlen, M.D. M.D. Brain M.D. Rendel M. Chemps, M.D., and Michael Schwertz, M.D. N. Engl. J. Med. 2001; 344:566-563 February 22, 2001

TH in traumatic brain injury

- Studies not as positive as those with cardiac arrest
- Why?
 - Time course of neuronal death is different
 - Peak levels of neuron-specific enolase (serum biomarker of neuronal death) occurs days after cardiac arrest, whereas they peak hours after traumatic brain injury

Berger RP, et al. Serum biomarkers after traumatic and hypoxemic brain injuries: insight into the

NABIS: Hypothermia IIR

- The primary hypothesis of NABIS:H IIR is as follows: Induction of hypothermia after severe traumatic brain injury to reach 33.0°C within 4h of injury and maintained for 48h in patients aged 16–45 years will result in an increased number of patients with good outcomes at 6 months after injury compared to patients randomized to management at normothermia.
- This trial did not confirm the utility of hypothermia as a primary neuroprotective strategy in patients with severe traumatic brain injury.

Clifton GL, et al "Very early hypothermia induction in patients with severe brain injury (the Nationa Acute Brain Injury Study: Hypothermia III): a randomised trial" Lancet Neurol 2010; DOI:10.1016/S1474-4422(10)70300-8.

Clinical trials assessing the effects of hypothermia on neurological outcome in patients with traumatic brain injury and intracranial hypertension.

Fever Above >38 C: Subarachnoid Hemorrhage: 73%	
Subarachnoid Hemorrhage: 73%	
Intracerebral Hemorrhage: 91%	
Traumatic Brain Injury: 68%	
Ischemic Stroke: 61%	
Fever Is Harmful	
Temperature elevations after brain ischemia worsen the	e resulting degree of injury
Fever associated with increased mortality and worse or	utcome
1 degree increase raises risk of poor outcome by 2.2 til	mes
 Mortality rate of stroke patients studied at 3 months wa 	s 1% in normothermic

	butor to excessive length
Fever Level	Increased # days in ICU
Medium (38.5-39.5C)	5.5 days
Low (to 38.5C)	1.7 days
ringer M, et al. Crit Care Med 2004 Vo	1.32, No7

IVTM for fever in	1 ICU
Threatment of fever in the neutral policy intensive care unit with a catcheter-based beat exchange system that the second of the	Prospective, randomized, controlled study of 296 Neuro ICU patients(TBI, SAH, CI, ICH) Compared intravascular cooling to existing best fever management methods
The control of the co	Result: 64% reduction in the incidence of fever with IVTM 25% reduction in use of
When it is all which the first is all and the first	antipyreticsNo safety issue!

Hypothermia in Acute Spinal Cord Injury Football Spina Injuries When New York Acute Spinal Repute When New York Acute Spinal When New York

Hypothermia in multisystem trauma

- 30%-50% of trauma mortality occurs before hospital
- Of those who die after arriving at the hospital, 70%-80% die within 24-48 hours
- Exsanguination is the most common cause of death in the first 48 hours after admission
- Once controlled, usually significant ischemic insult
- Some clinical trials underway, but by extrapolation, would hope this would improve outcomes

Clinical Applications Where Intravascular Temperature Management Has Been Used for

Cooling

Therapeutic hypothermia after cardiac arrest Fever control in Neuro/Surgical ICU

ICP (intracranial pressure) management

Therapeutic hypothermia for brain trauma and stroke

Acute liver failure

Adjunct with interventional procedure Malignant hyperthermia

Heat stroke

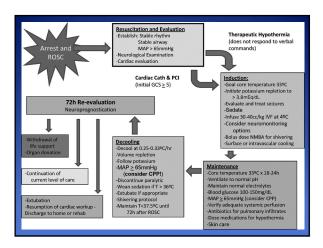
Spinal cord injury

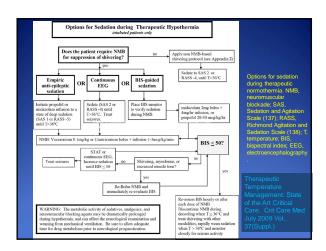
Spinal surgery

Adjunct with hemicraniectomy

Status epilepticus

Clinical Applications Where Intravascular Temperature Management Has Been Used for Warming Trauma victims Accidental hypothermia Burn surgery and intensive care Cardiac surgery OPCAB (off-pump coronary artery bypass) Post-bypass pump (prevention of after-drop) LVAD (left ventricular assist device) Transplant Thoracic aneurysm surgery Maintain viable donor organs for transplantation





System	Physiologic Effect	Nursing Actions
Systemic	Decreased metabolic demands	
	Decreased CO ₂ production	W 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1
Neurologic	Decreased O ₂ consumption Decreased cerebral metabolic	Monitor pulse oximetry and PaO2
reurologie	demands	
	Decreased intracranial pressure	
Cardiovascular	Decreased level of consciousness Tachycardia (observed during	
Cardiovascular	induction)	
	Bradycardia	None, unless symptomatic
	Hypertension	Wean vasopressors, administer analgesics
	Hypotension	and sedation if appropriate Consider fluid administration and
	riypotension	vasopressors
	Prolonged PR, QRS, and QT	vasopicasors
	intervals	
	Dysrhythmias (observed ≤32°C)	Prevent overcooling
	Decreased cardiac output Increased central venous pressure	None, unless hypotensive or symptomatic
	Increased mixed venous	
	oxygenation values	
Gastrointestinal	Decreased motility	Feeding may be delayed until the
Genitourinary	Diuresis	rewarming phase Monitor urine output; replace fluids as
ountourning.	2.141.4010	needed
Endocrine	Insulin resistance	Administer insulin to maintain glucose
		within a prescribed range
Immune	Suppression of white blood cells	Institute ventilator-associated pneumonia
		bundles, elevated head of bed, take measures to prevent infection