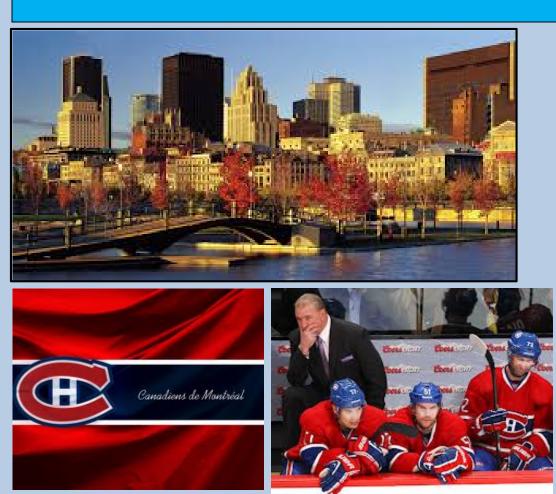
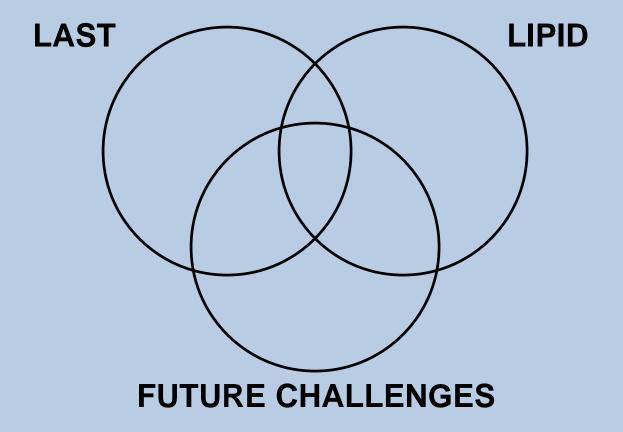
Treatment of Local Anesthetic Cardiac Toxicity



AAQ Montreal 16 Avril 2013 Guy Weinberg, MD Chicago, IL



Topics for Today



Classic Treatments for LAST Have <u>Miserable</u> Outcomes









Epidemiology

Mulroy M. Systemic Toxicity and Cardiotoxicity From Local Anesthetics: Incidence and Preventive Measures. *Regional Anesthesia and Pain Medicine, 2002; 27: 556–561*

INCIDENCE SYSTEMIC TOXIC REACTIONS

- ~7.5 20/10.000 PERIPHERAL NERVE BLOCKS
- ~1/10.000 EPIDURAL ANESTHESIA

maybe higher... <u>under-reporting</u> and misdiagnosis

maybe lower now ... <u>USGRA</u>

Cardiac arrest with PNB ~1/5000 (MH ~ 1/20,000-50,000 ~3 deaths/year).

The danger of experiential practice: Rare, uncommon events can be viewed as irrelevant We're all familiar with the refrain..... "That's never happened to me" As though, 'that never <u>could</u> happen to me'



Remember, inference, like hubris, is not our friend

block	drug	dose	timing	comment	Dr	result
fem	bup	200	Immed	.675%	A	Died
SQ	Bup/lid	220,450	30m	IHR	CA	Died
SQ	Bup/lid	70, 880	30m	Cocaine;	S	Survived
Klein's				phentermine		impaired
SQ	bup	100	<10m	wrist fracture	S	Died
SQ	bup	50	10m	lipoma	S	Died
SQ	bup	22	10m	Cn def	S	Recov
Klein's						
caudal	bup	25 (2/k)	Immed	infant	A	Recov
axillary	rop	200	immed	transart	А	Died

Complications Associated With Eye Blocks and Peripheral Nerve Blocks: An American Society of Anesthesiologists Closed Claims Analysis

Lorri A. Lee, M.D., Karen L. Posner, Ph.D., Frederick W. Cheney, M.D., Robert A. Caplan, M.D., and Karen B. Domino, M.D., M.P.H.

Claims with eye or peripheral nerve blocks performed perioperatively from 1980 through 2000 were analyzed. The injury patterns associated with peripheral nerve blocks and payment factors were analyzed.

Local anesthetic toxicity was associated with 7 of 19 claims with death or brain damage.

Reg Anesth Pain Med 2008;33:416-422

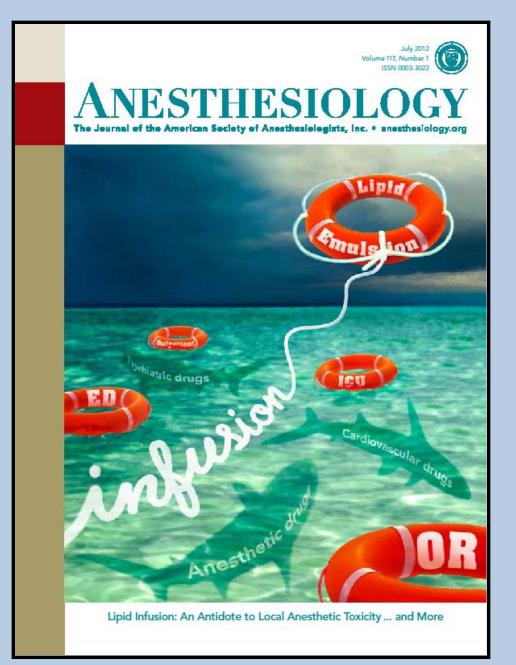
Emerging Patterns of Local Anesthetic Toxicity

- Non-anesthesia providers
- Office and off-site
- Atypical presentation
- Susceptible populations:
 - Elderly, metabolic disease
 - Very low cardiac output
 - Interval events: tourniquet, acidosis
- CNS: Extremely variable presentation
- Cardiac: variable and variable timing (late LAST)
 - Any arryhthmia;
 - Progressive bradycardia and hypotension

Cardiac History in Reports of Lipid Rescue from LA Toxicity = Susceptible Groups:

Rosenblatt: prior MI, RBBB and LAHB, angina Litz I: LABB, Stokes attacks, MVR, TVR Warren: S/P MI, RBBB, HTN Foxall: stable angina, QwV1-3, PAC Litz II: CAD, MVR Smith: S/P CABG, 2 stents

Note the unintended benefits of LipidRescue Resuscitation: More reports; clearer definition of clinical phenotype of LA toxicity. More insights into underlying mechanism and difference from lab models

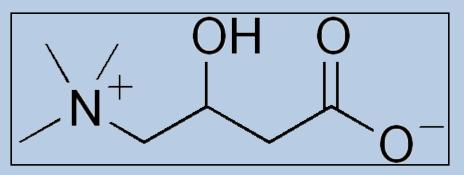


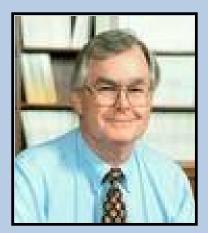
A Close Call in a Patient with a Metabolic Disease

20 yo patient with isovaleric acidemia: During SQ injection of bupivacaine, he develops arrhythmia with systolic BP 70, then complete heart block, then VT. <u>Bupivacaine dose= 22mg</u>. **Patient is severe carnitine deficiency**

> Weinberg, G.L., Laurito, C.E., Geldner, P., Pygon, B., Burton, B.J. Clin Anesthesia 1997

Carnitine





Charles Hoppel

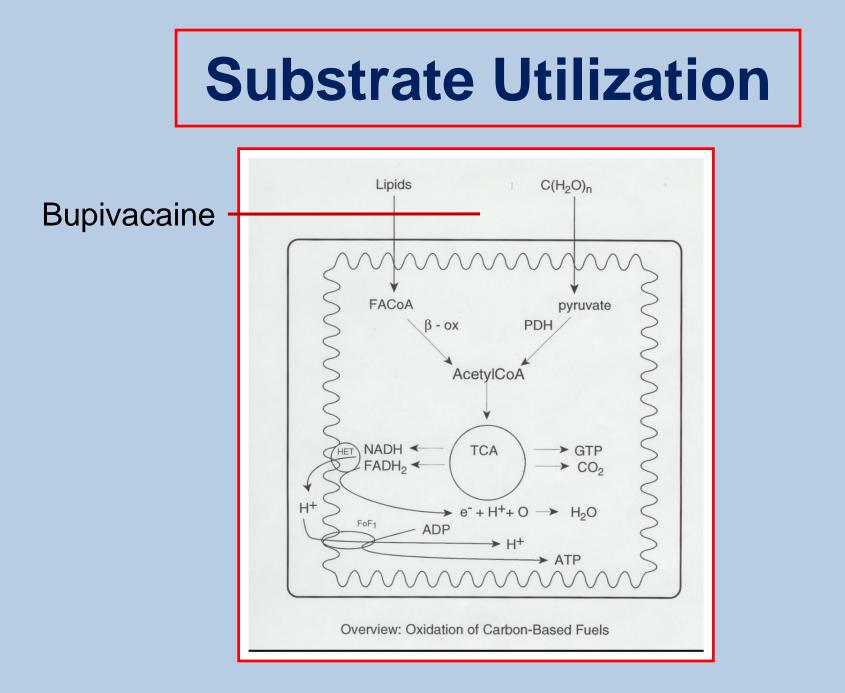
Biochemical Properties of Subsarcolemmal and Interfibrillar Mitochondria Isolated from Rat Cardiac Muscle*

(Received for publication, July 29, 1977)

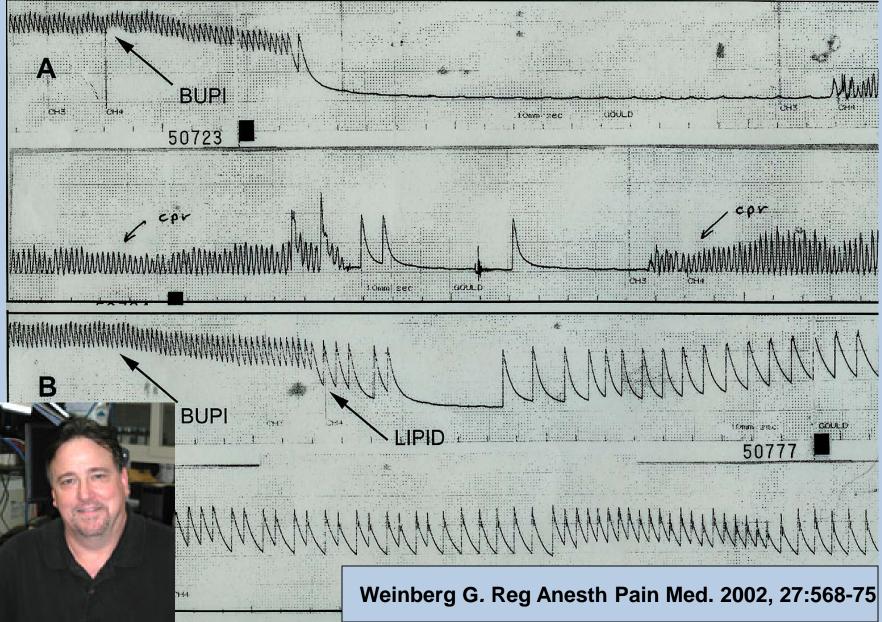
JUNE W. PALMER, BERNARD TANDLER, AND CHARLES L. HOPPELS

From the Veterans Administration Hospital and Departments of Pharmacology and Medicine, School of Medicine, and the Department of Oral Biology and Medicine, School of Dentistry, Case Western Reserve University, Cleveland, Ohio 44106

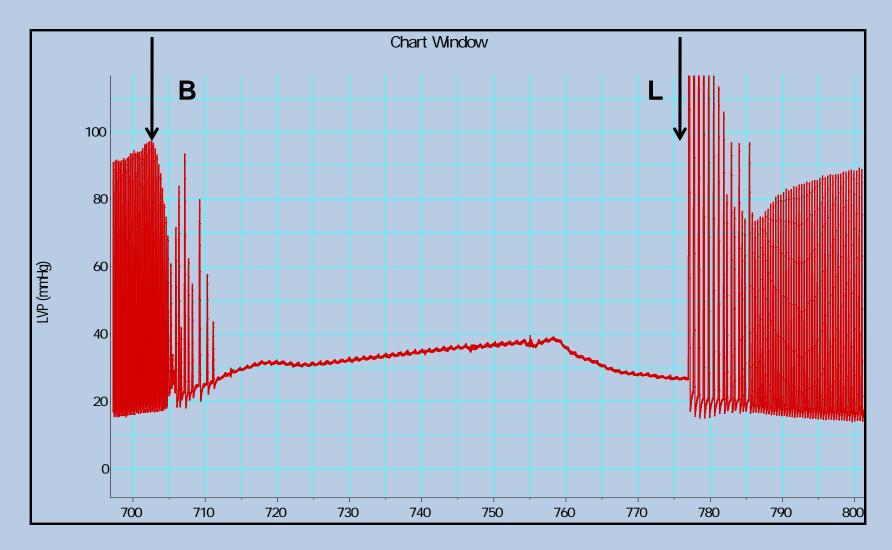
REQUIRED FOR FATTY ACID TRANSPORT INTO MITOS AND ATP SYNTHESIS FROM HEART'S PREFERED FUEL



Control vs. Lipid Rescue (15mg/kg)



Simplest Demonstration Possible



Translation of laboratory findings to the clinical setting

... the first successful use of Lipid

Anesthesiology 2006; 105:217-8

© 2006 American Society of Anesthesiologists, Inc. Lippincott Williams & Wilkins, Inc.

Successful Use of a 20% Lipid Emulsion to Resuscitate a Patient after a Presumed Bupivacaine-related Cardiac Arrest

Meg A. Rosenblatt, M.D.,* Mark Abel, M.D.,† Gregory W. Fischer, M.D.,† Chad J. Itzkovich, M.D.,‡ James B. Eisenkraft, M.D.§

INTERSCALENE BLOCK 20ml Bupivacaine 0.5% - 20ml Mepivacaine 1.5% Within 30 sec seizures (Propofol 50+100mg); 90 sec later CARDIAC ARREST... CPR,O2-ventilation,epinephrine,atropine,amiodarone,vasopressin, defibrillation... PULSELESS VT... Within 15 sec OF LIPID pulse/blood pressure detectable

YES!! Meg Rosenblatt (RAINER) saves the day



CASE REPORT Successful resuscitation of a patient with ropivacaine-induced asystole after axillary plexus block using lipid infusion*

R. J. Litz, M. Popp, S. N. Stehr and T. Koch

Anaesthesia, 2007, 62, pages 516–518

doi:10.1111/j.1365-2044.2007.05065.x

CASE REPORT Levobupivacaine-induced seizures and cardiovascular collapse treated with Intralipid[®]

G. Foxall,¹ R. McCahon,² J. Lamb,³ J. G. Hardman⁴ and N. M. Bedforth³

Anaesthesist 2007 · 56:449–453 DOI 10.1007/s00101-007-1147-3 Online publiziert: 3. Februar 2007 © Springer Medizin Verlag 2007

Redaktion K. Peter, München

C. Zimmer · K. Piepenbrink · G. Riest · J. Peters Klinik für Anästhesiologie und Intensivmedizin, Universitätsklinikum Essen, Essen

Kardio- und neurotoxische Nebenwirkungen nach akzidenteller intravasaler Bupivacainapplikation

Therapie mit Lidocain, Propofol und Lipidemulsion

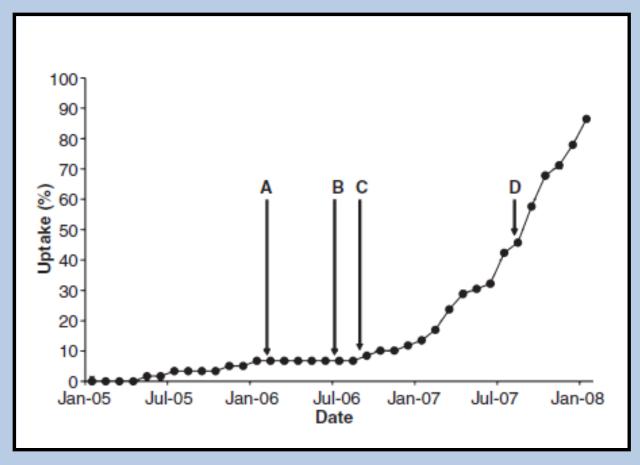
Anesthesiology 2007; 107:516-7

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Lipid Reversal of Central Nervous System Symptoms of Bupivacaine Toxicity

Andrew G. Spence, F.R.C.A., King Edward VII Memorial Hospital, Hamilton, Bermuda, aspence@transact.bm

ADOPTION OF LIPIDRESCUE IN LONDON HOSPITALS



Picard et al, Anaesthesia, 2009

LipidRescue[™] Resuscitation

... for drug toxicity

Click Here If You Need Help Right Now

Welcome

Background Treatment Overview Instructions (PDFs) Introducing LipidRescue to Your Facility Medical Literature I Local Anesthetic Toxicity Post Your Cases Sample LipidRescueTM Kit LipidRescueTM Experiments (Videos) Weinberg Bio Registry Proposal Ask Us An important horest for

all LipidRescuers

Professional Connections Places of interest <u>University of Illinois at</u> <u>Chicago Department</u> <u>of Anesthesiology</u> This has been my professional home for ao Welcome

LipidRescueTM resuscitation refers to the use of an intravascular vision of a lipid emulsion to treat severe, systemic drug toxicity or poisoning or was originally developed to treat local anesthetic toxicit' (a potentially fatal complication of regional anesthesia that can use even in other situations where patients receive local anesthetic mentions of or recently, LipidRescueTM has been shown in the performation of or recently, LipidRescueTM has been shown in the performation of or verdose caused by a wide array of other from total twesthetic) lipophilic agents. Initial support for this view way, totaled by a <u>most remarkable case</u> report where lipid emulsion infusion apparently saved a patient from overwhelming hepriorian bandose. Since then, evidence from both laboratory models and or endorts, indicates that LipidRescueTM can effectively in treat a wide valuety of non-local anesthetic overdoses, including reversal of both cardiovascular and central nervous system (CNS) signs and symptoms of toxicity.

This site was established in 2006 to serve as a source of information on LipidRescue¹³⁴ methodology and related issues. The intent was to provide a venue for the robust exchange of ideas on topics including the mechanisms, epidemiology, diagnosis, presentation, prevention and treatment of lifethreatening local anesthetic overdose and other types of severe cardiac and CNS toxicity.

Content includes the aforementioned aspects of local anesthetic toxicity plus links to related educational sites, typical experiments from my Aust Published...... A Comprehensive Review of Lipid Resuscitation REVIEW

Lipid emulsions in the treatment of acute poisoning: a systematic review of human and animal studies

 $CHLO\acute{E} JAMATY^{1}, BENOIT BAILEY^{2}, ALEXANDRE LAROCQUE^{3}, ERIC NOTEBAERT^{1}, KARINE SANOGO^{1}, and JEAN-MARC CHAUNY^{1}$

¹Department of Emergency Medicine, Hôpital du Sacré-Coeur de Montréal, Montréal, Quebec, Canada ²Division of Emergency Medicine, Department of Pediatrics, CHU Sainte-Justine, Montréal, Quebec, Canada ³Department of Emergency Medicine, Centre Hospitalier de l'Université de Montréal, Université de Montréal, Montréal, Québec, Canada

"In summary, there is convincing anecdotal and experimental evidence that IFE is effective in treating toxicity caused by local anesthetics".

"...Current evidence suggests that <u>IFE should be</u> <u>administered as soon as a diagnosis</u> of local anesthetic toxicity is established,"



American Society of Regional Anesthesia and Pain Medicine

Practice Advisory on Treatment of Local Anesthetic Systemic Toxicity

For Patients Experiencing Signs or Symptoms of Local Anesthetic Systemic Toxicity (LAST)

- Get Help
- Initial Focus
 - o Airway management: ventilate with 100% oxygen
 - o Seizure suppression: benzodiazepines are preferred
 - o Basic and Advanced Cardiac Life Support (BLS/ACLS) may require prolonged effort
 - Intuse 20% Lipid Emulsion (values in parenthesis are for a 70 kg patient)
 - o Bolus 1.5 mL/kg (lean body mass) intravenously over Mmin (~100 mL)
 - o Continuous infusion at 0.25 mL/kg/min (~18 mL/min; adjust by roller clamp)
 - o Repeat bolus once or twice for persistent cardiovascular collapse
 - o Double the infusion rate to 0.5 mL/kg per minute if blood pressure remains low
 - o Continue infusion for at least 10 mins after attaining inculatory stability
 - o Recommended upper limit: approximately 10 mL kg lipid emulsion over the first 30 mins
- · Avoid vasopressin, calcium channel blockers, p-blockers, or local anesthetic
- · Alert the nearest facility having cardiopulmonary bypass capability
- · Avoid propofol in patients having signs of cardiovascular instability
- · Post LAST events at www.lipidrescue.org and report use of lipid to www.lipidregistry.org

Part 12: Cardiac Arrest in Special Situations 2010 American Heart Association Guidelines for Cardiopulmonary Resuscitation and Emergency Cardiovascular Care

Terry L. Vanden Hoek, Chair; Laurie J. Morrison; Michael Shuster; Michael Donnino; Elizabeth Sinz; Eric J. Lavonas; Farida M. Jeejeebhoy; Andrea Gabrielli

This section of the 2010 AHA Guidelines for CPR and ECC addresses cardiac arrest in situations that require special treatments or procedures beyond those provided during basic life support (BLS) and advanced cardiovas-cular life support (ACLS). We have included 15 specific cardiac arrest situations. The first several sections discuss cardiac arrest associated with internal physiological or metabolic conditions, such as asthma (12.1), anaphylaxis (12.2), pregnancy (12.3), morbid obesity (12.4), pulmonary embolism (PE) (12.5), and electrolyte imbalance (12.6).

The next several sections relate to resuscitation and treatment of cardiac arrest associated with external or environmentally related circumstances, such as ingestion of toxic substances (12.7), trauma (12.8), accidental hypothermia (12.9), avalanche (12.10), drowning (12.11), and electric shock/lightning strikes (12.12).

The last 3 sections review management of cardiac arrest that may occur during special situations affecting the heart, including percutaneous coronary intervention (PCI) (12.13), cardiac tamponade (12.14), and cardiac surgery (12.15).

Part 12.1: Cardiac Arrest Associated With Asthma

Asthma is responsible for more than 2 million visits to the

Pathophysiology

The pathophysiology of asthma consists of 3 key abnormalities:

- Bronchoconstriction
- · Airway inflammation
- · Mucous plugging

Complications of severe asthma, such as tension pneumothorax, lobar atelectasis, pneumonia, and pulmonary edema, can contribute to fatalities. Severe asthma exacerbations are commonly associated with hypercarbia and acidemia, hypotension due to decreased venous return, and depressed mental status, but the most common cause of death is asphyxia. Cardiac causes of death are less common.⁴

Clinical Aspects of Severe Asthma

Wheezing is a common physical finding, although the severity of wheezing does not correlate with the degree of airway obstruction. The absence of wheezing may indicate critical airway obstruction, whereas increased wheezing may indicate a positive response to bronchodilator therapy.

Oxygen saturation (SaO₂) levels may not reflect progressive alveolar hypoventilation, particularly if oxygen is being administered. Note that SaO₂ may fall initially during therapy because β_2 -agonists produce both bronchodilation and vasodilation and

RECOMMENDATIONS:

- Oxygenation/ Ventilation!CPR
- Seizure Control (BDZ preferred) •20% LIPID 1.5ML/KG BOLUS •INFUSION 0.25-0.5ML/KG/MIN X 30min •REPEAT BOLUS x 2 IF NEEDED •NO Propofol, CC or ß Blockers, LAs Ready Cardiopulmonary Bypass * Don't give up – bupivacaine is protective

Emil Mayer on LAST, 1924

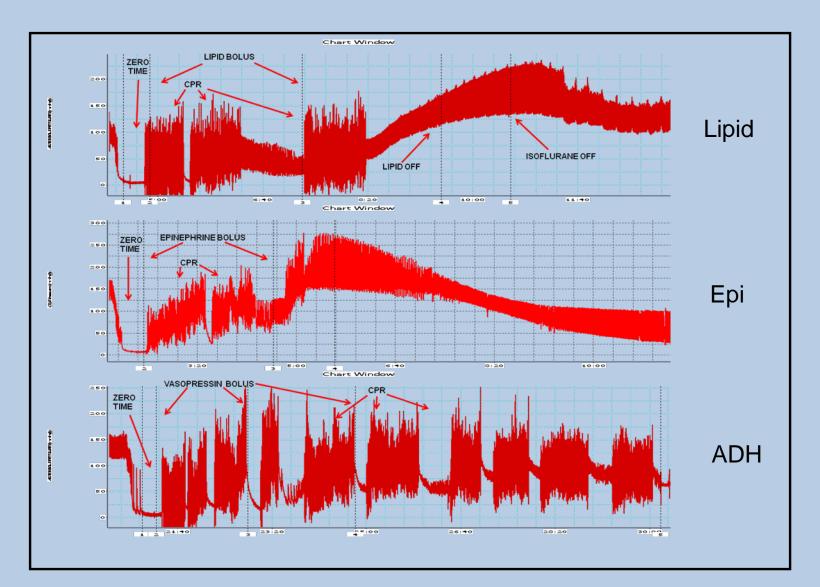
1. "Such accidents are more frequent than is commonly supposed"

2. "Unexplained differences in susceptibility exist"

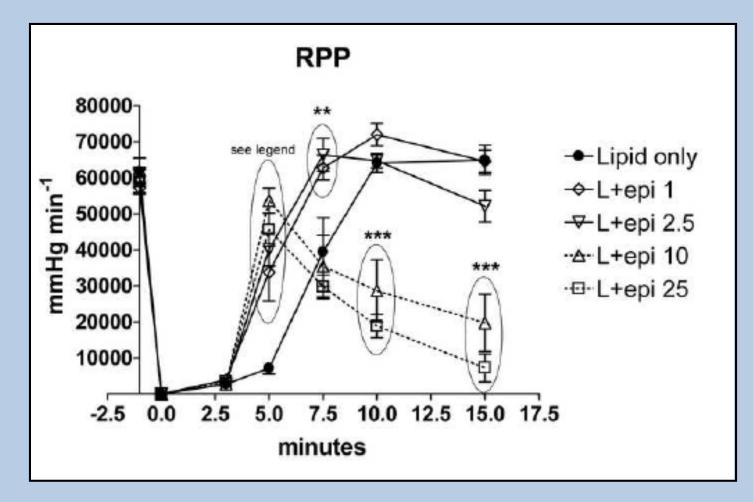
3. "A proportion of cases are avoidable mistakes"

4. "In treatment, the first place must be assigned to artificial respiration, perhaps with cardiac massage"5. The committee advises strongly against the routine use of morphin and epinephrin in the treatment of accidents"

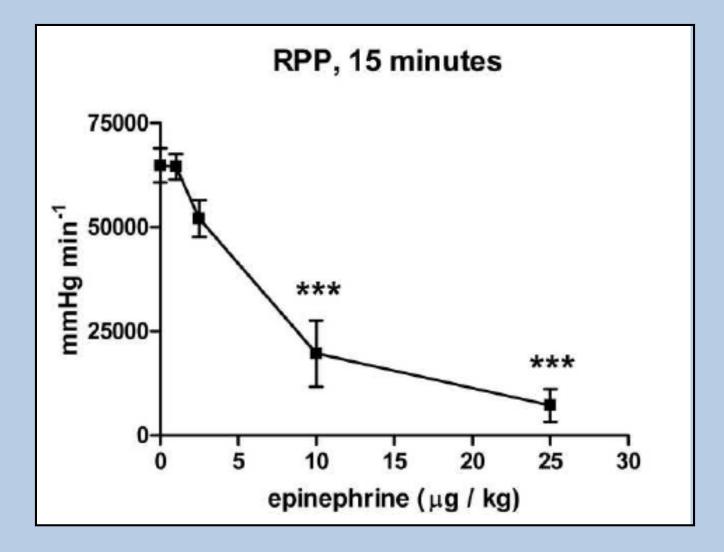
Lipid vs Pressors



Lipid + Epinephrine: Paradox of Initial Recovery



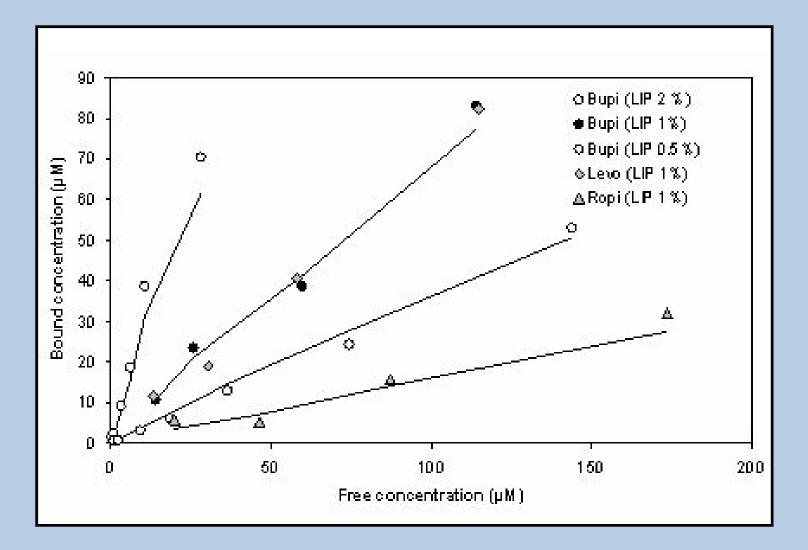
Epinephrine Impairs Outcome



Things you should know about ILE

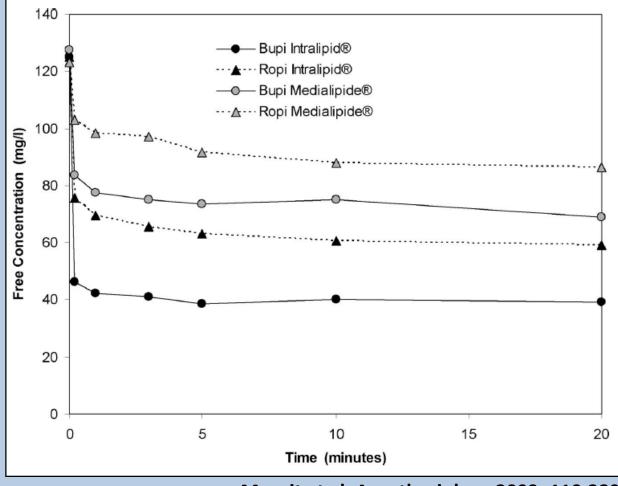
- Need 20%, start with bolus then follow
- Dose to ideal body weight (esp in US)
- Infusion rate is not precise (roller clamp OK)
- Don't forget to turn it off (12mL/kg/30min)
- Works with ALL local anesthetics
- Early is better
- Safe scant evidence of adverse events
- Less pressor therapy (NO vasopressin)
- Severe CAD can impair efficacy of ILE

SO, HOW DOES IT WORK?



Mazoit et al ASA '08 in vitro lipid LA binding

Rapid binding of local anesthetics by lipid emulsions

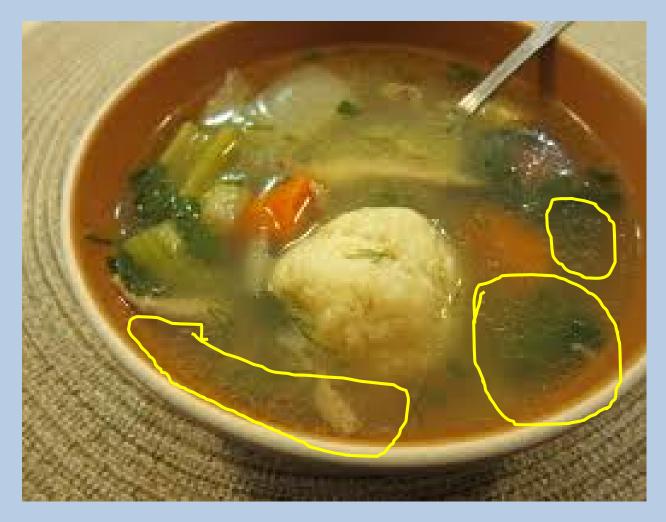


Mazoit et al. Anesthesiology 2009; 110:380.

This is *in vitro* experiment shows very rapid drop of 'free' (nonlipid-bound) local anesthetic concentraton in 1% solutions of Intralipid or Medialipid over time. The starting concentrations of both local anesthetics is 125mg/L. Intralipid binds more anesthetic than Medialipid and each lipid binds more bupivacaine than ropivacaine. ³³



Typical initial lipid resuscitation bolus: 100mL of 20% = 20g fat



Total fat in a single serving of chicken soup: 27.2g

But, there is more to lipid than a sink

Phosphorylation of GSK-3β Mediates Intralipid-induced Cardioprotection against Ischemia/Reperfusion Injury

Slamak Rahman, M.D.,* Jingyuan Ll, M.D., Ph.D.,† Jean Chrisostome Bopassa, Ph.D.,† Soban Umar, M.D., Ph.D.,† Andrea lorga, B.Sc.,‡ Parisa Partownavid, M.D.,§ Mansouren Eghbal, Ph.D. Clinical Toxicology (2011), **49**, 729–733 Copyright © 2011 Informa Healthcare USA, Inc. ISSN: 1556-3650 print / 1556-9519 online DOI: 10.3109/15563650.2011.613399



RESEARCH ARTICLE

Fatty acids antagonize bupivacaine - induced I_{Na} blockade

ALLAN R. MOTTRAM¹, CARMEN R. VALDIVIA², and JONATHAN C. MAKIELSKI²

¹University of Wisconsin, Division of Emergency Medicine, 600 Highland Ave, F2/204 CSC, MC 3280, Madison, United States ²University of Wisconsin, Department of Medicine, 600 Highland Ave, Madison, United States

Drug Redistribution: Rethinking the "Sink"

Anaesthesia 2012, 67, 600-605

doi:10.1111/j.1365-2044.2012.07056.x

Original Article

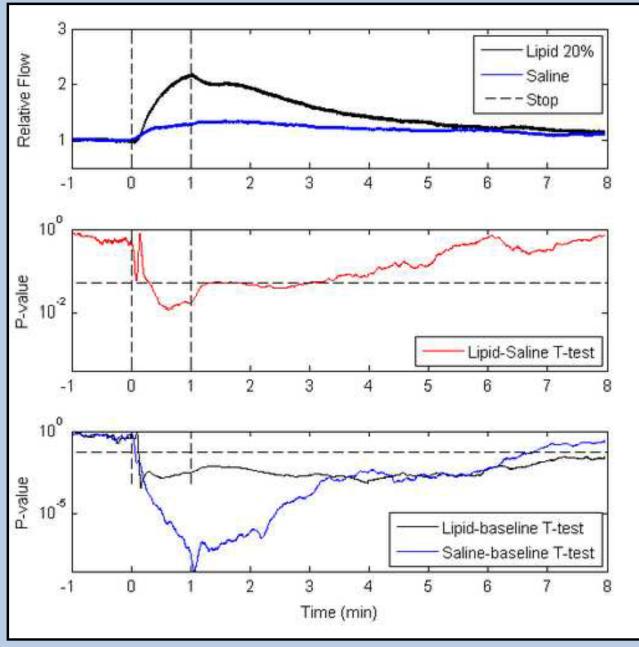
Effect of intravenous lipid emulsion on bupivacaine plasma concentration in humans

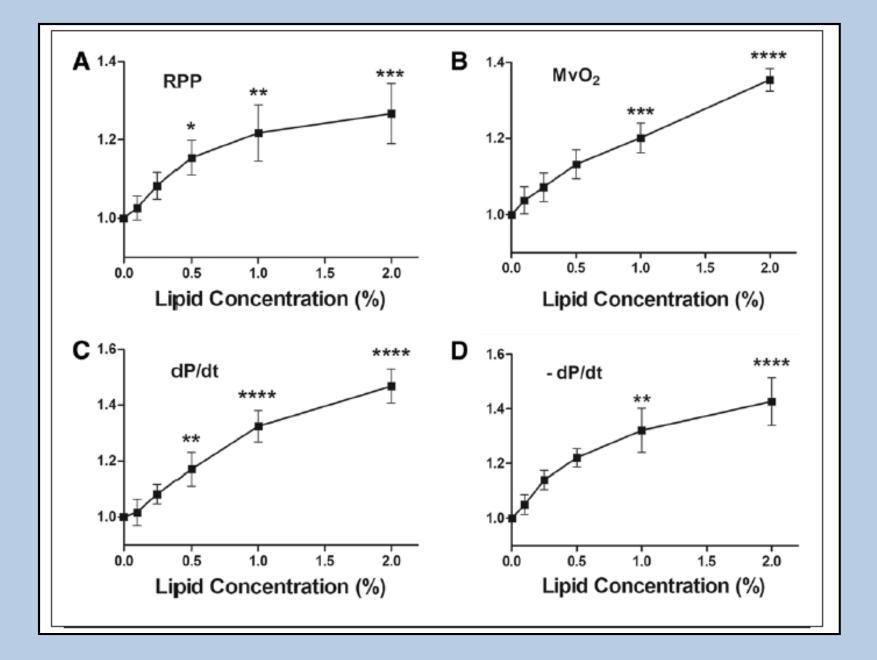
E. Litonius,¹ P. Tarkkila,² P. J. Neuvonen³ and P. H. Rosenberg⁴

 Specialty Registrar, 4 Professor, Anaesthesiology and Intensive Care Medicine, Helsinki University Central Hospital and University of Helsinki, Helsinki, Finland
Chief Physician, Anaesthesiology and Intensive Care Medicine, Töölö Hospital, Helsinki, Finland
Professor, Department of Clinical Pharmacology, University of Helsinki and HUSLAB, Helsinki, Finland

"Intravenous lipid emulsion infusion reduced the context-sensitive half-life of total plasma bupivacaine from 45 min to 25 min; p = 0.01. We observed no significant adverse effects of lipid emulsion. In conclusion, lipid emulsion may slightly increase the rate of bupivacaine tissue distribution. No 'lipid sink' effect was observed with the non-toxic dose of bupivacaine used".

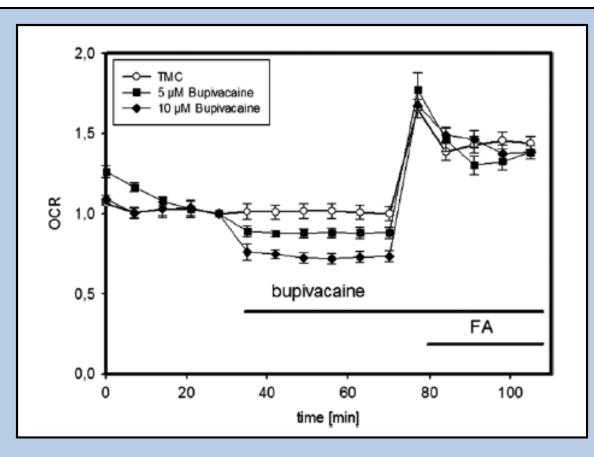
DIRECT CARDIOVASCULAR BENEFIT

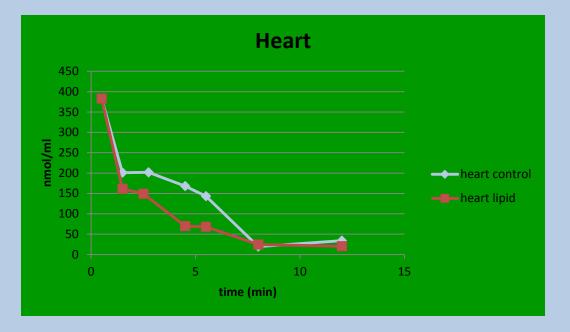


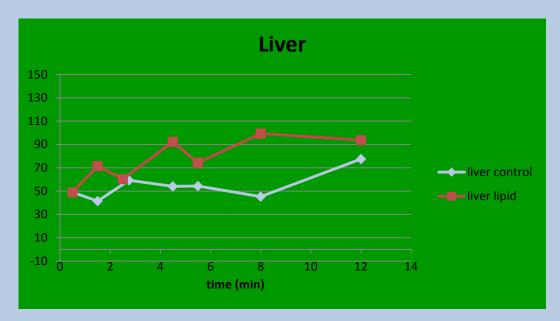


Myocardial Accumulation of Bupivacaine and Ropivacaine Is Associated with Reversible Effects on Mitochondria and Reduced Myocardial Function

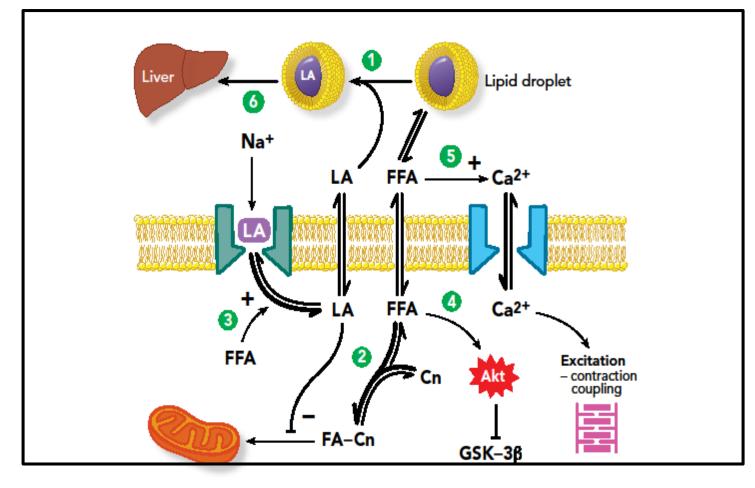
Nicole Hiller, MSc,* Peter Mirtschink, MD,† Christine Merkel, MD,* Lilla Knels, MD,‡ Reinhard Oertel, PhD,§ Torsten Christ, MD,|| Andreas Deussen, MD,¶ Thea Koch, MD,* and Sebastian N. Stehr, MD, DESA*

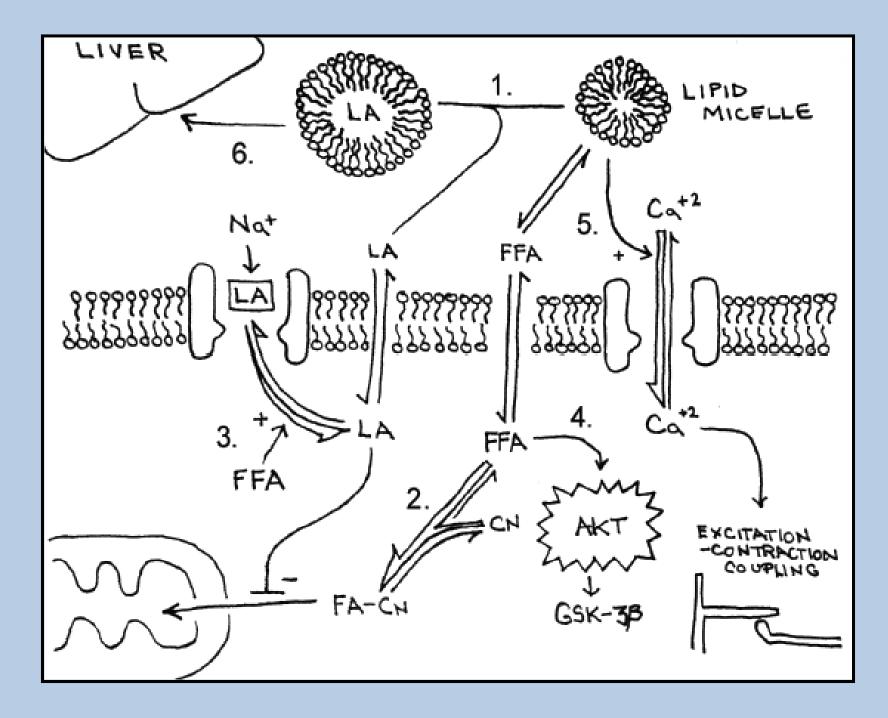




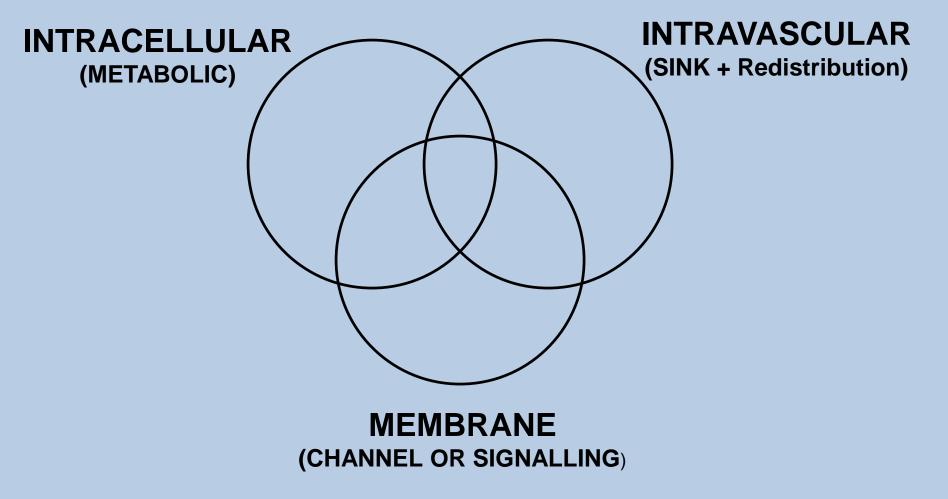


Mechanisms of LipidRescue -More than a Sink





Mechanism(s) Summary



SO, WHAT'S NEW?

ORIGINAL ARTICLE

ASRA Checklist Improves Trainee Performance During a Simulated Episode of Local Anesthetic Systemic Toxicity

Joseph M. Neal, MD,* Robert L. Hsiung, MD,* Michael F. Mulroy, MD,* Brian B. Halpern, RN,† Alison D. Dragnich, MD,* and April E. Slee, MSc‡



Pediatric Anesthesia ISSN 1155-5645

ORIGINAL ARTICLE

Comparison of epinephrine vs lipid rescue to treat severe local anesthetic toxicity – an experimental study in piglets

Jacqueline Mauch^{1,2}, Olga Martin Jurado³, Nelly Spielmann¹, Regula Bettschart-Wolfensberger³ & Markus Weiss¹

1 Department of Anesthesia, University Children's Hospital Zurich, Zurich, Switzerland

2 Department of Anesthesia and Perioperative Medicine, Kantonsspital Aarau, Aarau, Switzerland

3 Section Anesthesiology, Equine Department, Vetsuisse Faculty University of Zurich, Zurich, Switzerland

Checklist for Treatment of Local Anesthetic Systemic Toxicity

The Pharmacologic Treatment of Local Anesthetic Systemic Toxicity (LAST) is Different from Other Cardiac Arrest Scenarios

- Get Help
- Initial Focus
 - □ Airway management: ventilate with 100% oxygen
 - □ Seizure suppression: benzodiazepines are preferred; AVOID propofol in patients having signs of cardiovascular instability
 - □ Alert the nearest facility having cardiopulmonary bypass capability
- Management of Cardiac Arrhythmias
 - Basic and Advanced Cardiac Life Support (ACLS) will require adjustment of medications and perhaps prolonged effort
 - □ AVOID vasopressin, calcium channel blockers, beta blockers, or local anesthetic
 - □ REDUCE individual epinephrine doses to <1 mcg/kg
- Lipid Emulsion (20%) Therapy (values in parenthesis are for 70kg patient)
 - □ Bolus 1.5 mL/kg (lean body mass) intravenously over 1 minute (~100mL)
 - Continuous infusion 0.25 mL/kg/min (~18 mL/min; adjust by roller clamp)
 - □ Repeat bolus once or twice for persistent cardiovascular collapse
 - Double the infusion rate to 0.5 mL/kg/min if blood pressure remains low
 - □ Continue infusion for at least10 minutes after attaining circulatory stability
 - Recommended upper limit: Approximately 10 mL/kg lipid emulsion over the first 30 minutes
- □ Post LAST events at www.lipidrescue.org and report use of lipid to www.lipidregistry.org



Pediatric Anesthesia ISSN 1155-5645

ORIGINAL ARTICLE Pediatric Anesthesia 21 (2011) 1103–1108

Comparison of epinephrine vs lipid rescue to treat severe local anesthetic toxicity – an experimental study in piglets

Jacqueline Mauch^{1,2}, Olga Martin Jurado³, Nelly Spielmann¹, Regula Bettschart-Wolfensberger³ & Markus Weiss¹

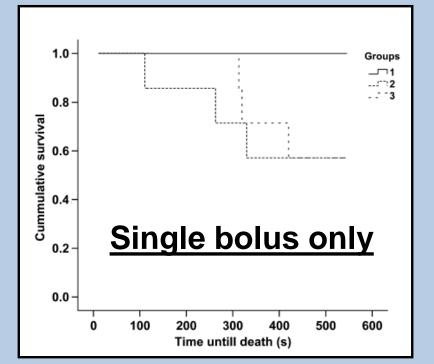
1 Department of Anesthesia, University Children's Hospital Zurich, Zurich, Switzerland

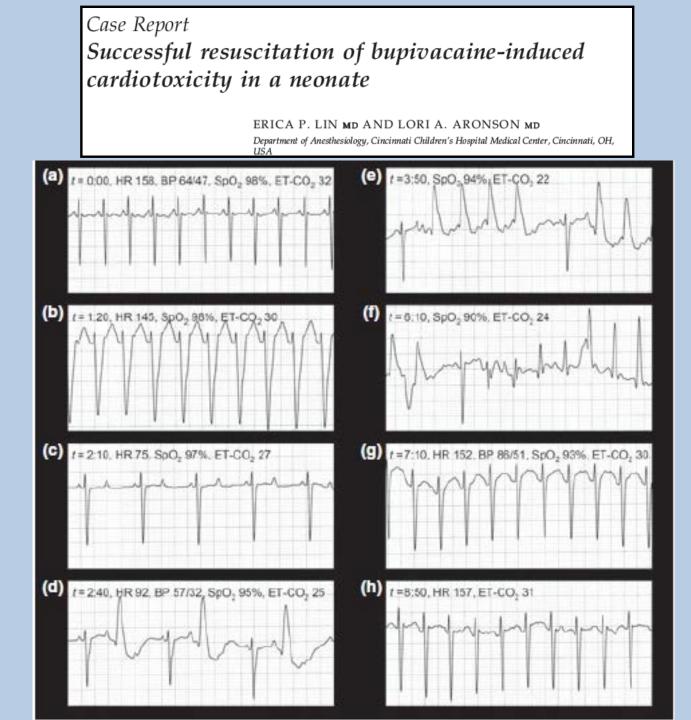
2 Department of Anesthesia and Perioperative Medicine, Kantonsspital Aarau, Aarau, Switzerland

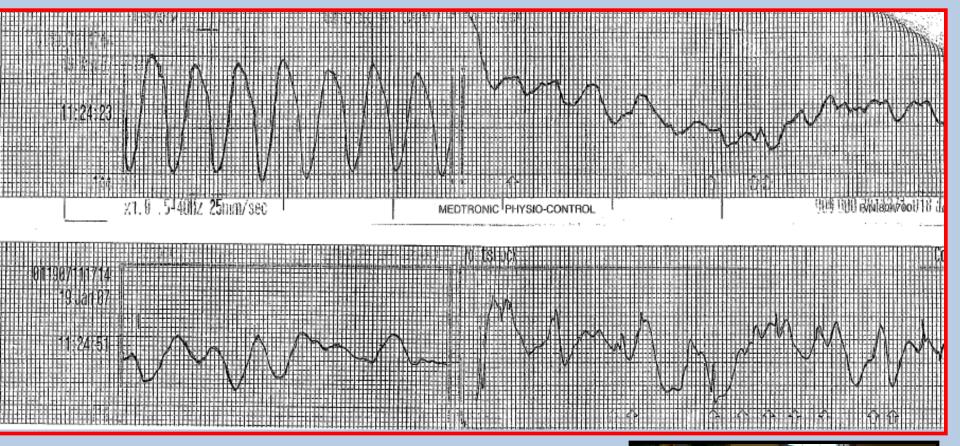
3 Section Anesthesiology, Equine Department, Vetsuisse Faculty University of Zurich, Zurich, Switzerland

Administering epinephrine resulted in an overshoot in MAP over baseline by factor 1.5–2.5 and in heart rate by factor 1.4–1.9. With epinephrine administration, selflimiting tachy-arrhythmias were observed, sometimes transiently compromising blood pressure.

For the treatment of severe hemodynamic compromise owing to bupivacaine intoxication in piglets, firstline rescue with epinephrine was more effective than Intralipid







Archie Sirianni saved the day



>90 Minutes Total CPR

			CXR	LI Capno Device							
Time	Vital Signs	EKG Bhythm	Interventions Shock, Drugs, IV, Intubation	Response/ Comments	CPR: (yes/no)	Time	Vital Signs	EKG Rhythm	Interventions Shock, Drugs, IV, Intubation Procedures	Response/ Comments	CPR: (yes/no)
1115		Initial PEA Rhythm:	Procedures Aturan 2	DiOConner	Y	1221	94	mitial PO'74 Rhythm:	epi - Epigtt		y
1118,	[]	VT	Eper Defit 200	Disharma/	4	1222	Pulse Street	CP1 POTU	acar 385	20	y.
1120		Vī	Dyib 300 Epc. Def 360		4	122	\$ 75	- and y	Epi ABG		ý
11224	1440	VT	Ept: Def 360		4	1229	42 ECP	L	Epi.		4
1124	54/28		Defib 360.		y	12.34			Epi r'		- Ye
1125		VT VT	Del 360.	mia 300 Bd	T	1235	p Pul	x \$	Pube CPR		4
1127	130	VF	Epi Defito 360		Ty		12/5)	leve stt.		Y Q
1120		VF .	Evil Mars Igna /16	hi hi hikd	4	123		da)	Epi:		
1130		VF	Epic Mag Igm/16	a moubut	yt	1343	01/14	Nor-		pladspush	y-
1132	140	WT.	Epi		14	1240			Vent TRZO 100 Door Wide	¹ 600, ⁰	
133,	10000 -		Dilib 300	cle Choice Physician C	14				PEast Arrest Clature, Circle	Obstan of	·

1min after 100mL lipid, ROSC, CPR stopped, QRS narrows. Patient recovers, neurologically intact.

Toxicologists Now Aware of Lipid Therapy

ARTICLE IN PRESS

TOXICOLOGY/CASE REPORT

Use of <u>Lipid Emulsion</u> in the Resuscitation of a Patient With Prolonged Cardiovascular Collapse After <u>Overdose of Bupropion</u> and Lamotrigine

Archie J. Sirianni, MD Kevin C. Osterhoudt, MD Diane P. Calello, MD Allison A. Muller, PharmD Marie R. Waterhouse, MD Michael B. Goodkin, MD Guy L. Weinberg, MD Fred M. Henretig, MD From the Department of Anesthesiology (Sirianni) and Division of Cardiology (Goodkin), Riddle Memorial Hospital, Media, PA; the Department of Pediatrics, University of Pennsylvania School of Medicine and The Children's Hospital of Philadelphia, Philadelphia, PA (Osterhoudt, Calello, Muller, Waterhouse, Henretig); the Section of Clinical Toxicology, Division of Emergency Medicine, and the Poison Control Center, The Children's Hospital of Philadelphia, Philadelphia, PA (Osterhoudt, Calello, Muller, Henretig); and the Department of Anesthesiology, University of Illinois College of Medicine at Chicago, and Jessie Brown VA Medical Center, Chicago, IL (Weinberg).

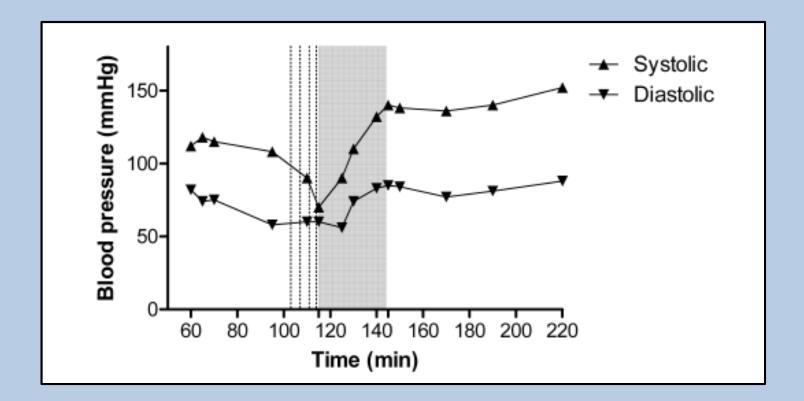
Animal studies show efficacy of intravenous lipid emulsion in the treatment of severe cardiotoxicity associated with local anesthetics, clomipramine, and verapamil, possibly by trapping such lipophilic drugs in an expanded plasma lipid compartment ("lipid sink"). Recent case reports describe lipid infusion for the successful treatment of refractory cardiac arrest caused by parenteral administration of local anesthetics, but clinical evidence has been lacking for lipid's antidotal efficacy on toxicity caused by ingested medications. A 17-year-old girl developed seizure activity and cardiovascular collapse after

CASE REPORT

Open Access

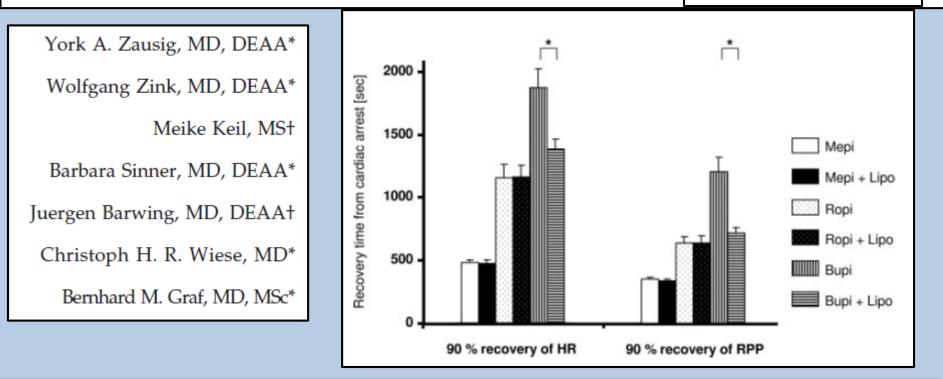
Case report: successful lipid resuscitation in multidrug overdose with predominant tricyclic antidepressant toxidrome

Martyn Harvey^{1*} and Grant Cave²



Lipid Emulsion Improves Recovery from Bupivacaine-Induced Cardiac Arrest, but Not from Ropivacaine- or Mepivacaine-Induced Cardiac Arrest

(Anesth Analg 2009;109:1323-6)



CONCLUSIONS: These data show that the effects of lipid infusion on LA-induced cardiac arrest are strongly dependent on the administered LAs itself. We conclude that lipophilicity of LAs has a marked impact on the efficacy of lipid infusions to treat cardiac arrest induced by these drugs.

Case Report

Intractable cardiac arrest due to lidocaine toxicity successfully resuscitated with lipid emulsion* (Crit Care Med 2011; 39:872–874)

Stephanie K. Dix, MD; Gregg F. Rosner, MD; Monica Nayar, PharmD; Julian J. Harris, MD; Maya E. Guglin, MD; Jeffery R. Winterfield, MD; Zhiling Xiong, MD, PhD; Gilbert H. Mudge, Jr., MD

Use of Intravenous Lipid Emulsion to Reverse Central Nervous System Toxicity of an Iatrogenic Local Anesthetic Overdose in a Patient on Peritoneal Dialysis

Ann Pharmacother 2012;46:e37.

D Bruce Lange, Daniel Schwartz, Gerald DaRoza, and Robert Gair

fore, at approximately 30 minutes from the initial signs of lidocaine toxicity, a 20% lipid emulsion (Intralipid, Fresenius Kabi) was administered as a 1.5-mL/kg intravenous bolus (100 mL) over approximately 10 minutes. Within approximately 5 minutes (ie, after 50 mL of lipid was infused), the patient became more alert and exhibited improved muscle function. In approximately 10 minutes (after 100 mL), the patient was able to speak coherently and



Anaesthesia Journal of the Association of Anaesthetists of Great Britain and Ireland

Anaesthesia, 2011

doi:10.1111/j.1365-2044.2011.06895.x

CASE REPORT Treatment of cocaine overdose with lipid emulsion

R. Jakkala-Saibaba,¹ P. G. Morgan² and G. L. Morton²

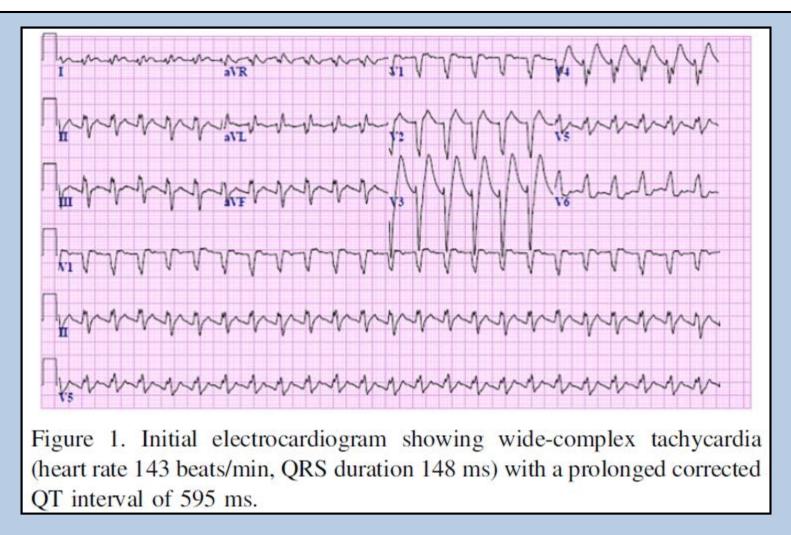
1 Specialist Registrar, 2 Consultant, John Hammond Department of Anaesthesia, East Surrey Hospital, Redhill, UK

Summary

We describe the management and recovery of a 28-year-old man following a history of overdose by nasal inhalation of cocaine. The patient was presented in a comatose state suffering from seizures and marked cardiovascularly instability. Intravenous lipid emulsion was administered following initial resuscitation and tracheal intubation, as a means of treating persistent cardiac arrhythmias and profound hypotension. Following lipid emulsion therapy, the patient's life-threatening cardiovascular parameters rapidly improved and he recovered well without any side effects, thus being discharged within 2 days.

Usefulness of Intravenous Lipid Emulsion for Cardiac Toxicity from Cocaine Overdose

Natasha Purai Arora, MD^a, William Allen Berk, MD^b, Cynthia Kurke Aaron, MD^{c,d}, and Kim Allan Williams, MD^{e,*}



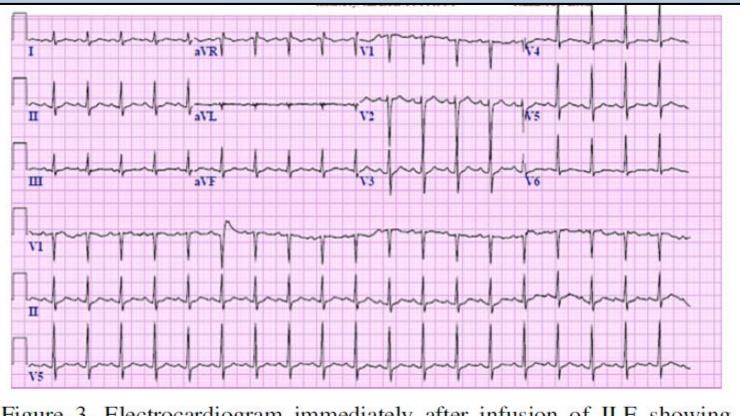


Figure 3. Electrocardiogram immediately after infusion of ILE showing regular sinus rhythm (heart rate 118 beats/min) with normal QRS (82 ms) and corrected QT (412 ms) intervals.

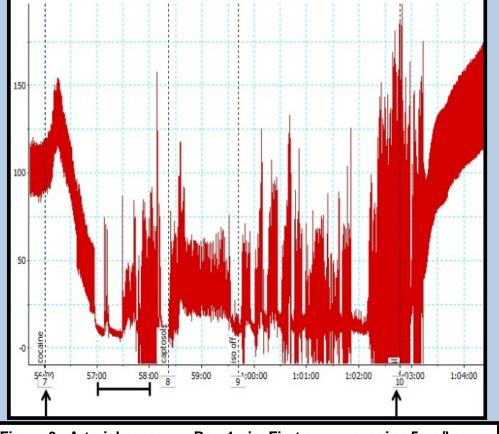
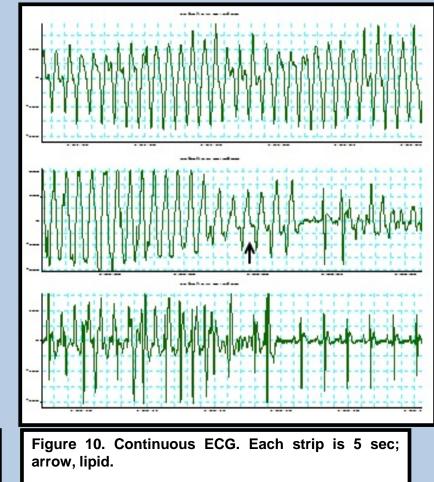
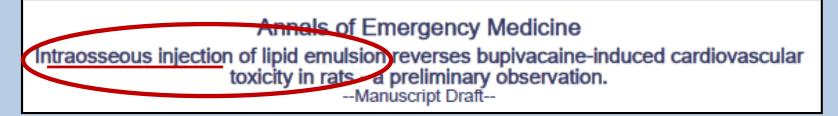
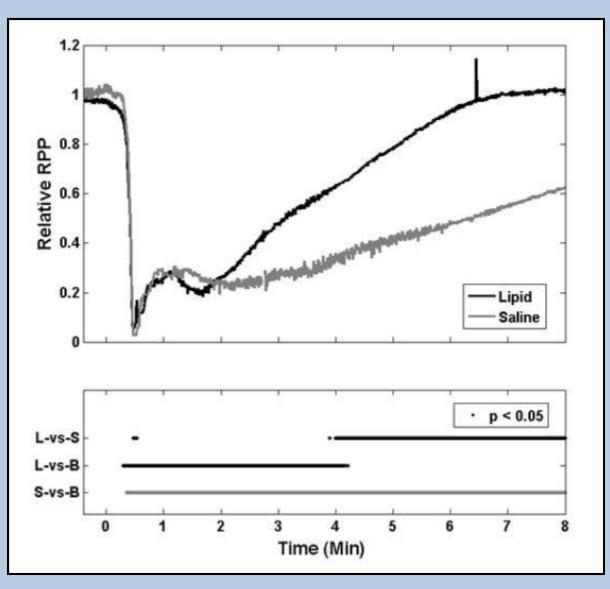


Figure 9. Arterial pressure. Bar ,1min. First arrow, cocaine 5mg/kg; second arrow, lipid.





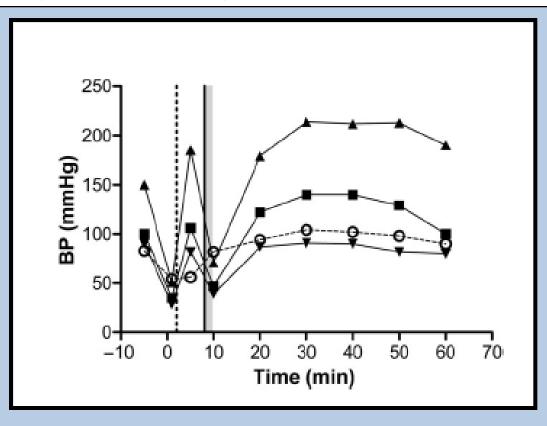


TOXICOLOGY



Successful resuscitation from bupivacaine-induced cardiovascular collapse with intravenous lipid emulsion following femoral nerve block in an <u>emergency department</u>

Martyn Harvey,¹ Grant Cave,² Giles Chanwai¹ and Tonia Nicholson¹ ¹Waikato Hospital, Hamilton and ²Hutt Hospital, Lower Hutt, New Zealand



Kensley Kirby, 5-Year-Old Atlanta Girl, Dies From Lethal Dose Of Local Anesthetic



Lipo doctor accused in patient's death gets 30-day suspension

By VANESSA HO, SEATTLEPI.COM STAFF Updated 12:57 p.m., Thursday, September 29, 2011

A Bellevue surgeon accused in the death of a liposuction patient has been suspended for 30 days, the state Department of Health said Thursday.

The Medical Quality Assuarance Commission had charged Marco Sobrino with unprofessional conduct last year, after his patient died following liposuction of her abdomen and upper arms.

After the three-hour procedure, Sobrino left the clinic, without ensuring his patient was in stable condition, investigators said. They said he left Javellana with no nurse, only a medical assistant, and without discharge instructions or a follow-up phone number.

After the procedure, Javellana vomited and was drowsy. She was told an adult should pick her up and stay with her for 24 hours, but not the reason. When no one came to pick up Javellana, Sono Bello staff put her in a cab alone, with no idea of where she was going, the state said. Javellana died in a hotel room the next day. The King County Medical Examiner ruled she died from 'acute lidocaine intoxication'.

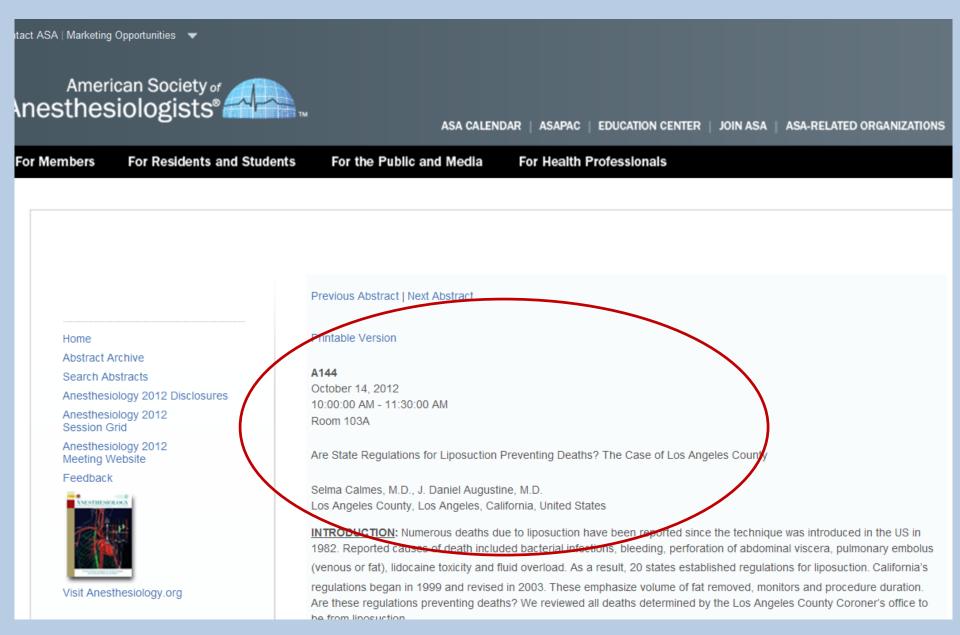
Investigators: Autopsy reveals overdose caused cosmetic surgery death



by LINDA BYRON / KING 5 News



Are New Laws or Regulations Required To Improve Safety in 'Cosmetic Surgery'?



Conclusions

- LAST occurs despite our best efforts
- Preparation makes a difference
- Lipid infusion makes a difference
- Education makes a difference
- Let our colleagues know about LAST

