


Cerebral Arterial Gas Embolism

D. Lindsie Cone, MD, UHM, FUHM

Air / Gas Embolism

Cerebral Arterial Gas Embolism



"Primary Training in Hyperbaric Medicine"
D. Lindsie Cone, MD

1

Cerebral Arterial Gas Embolism

...Definition

The introduction of gaseous emboli into the cerebral arterial circulation.

- In divers, it is not necessarily related to an excessive tissue gas burden.
- *Medicare: "Gas Embolism"
- *UHMS: "Air or Gas Embolism"

**Not specific to the brain/arterial circulation: supportive research is.*

2

Cerebral Arterial Gas Embolism

...Etiology

- Decompression
- Iatrogenic events
- Trauma

3

Cerebral Arterial Gas Embolism

...Mechanism: Decompression-Induced

*~ intratracheal pressures of 70-80mmHg
- Polack & Adams, 1932*

- Alveolar membrane failure
- A change in transpulmonary pressure of 70 – 80 mmHg is sufficient.
- Equivalent to an ascent from a depth of only ~1 meter of sea water.

4

Cerebral Arterial Gas Embolism

...Mechanism: Decompression-Induced

Shallowest recorded depth involving pulmonary barotrauma-induced CAGE as a clinical diagnosis

- P.J. Benton, et al. 1996 Avia Space Environ Med: 67(1)

- Helicopter escape training exercise
- Utilizing Short Term Air Supply System (STASS)

5

Cerebral Arterial Gas Embolism

...Iatrogenic events

<ul style="list-style-type: none"> • Cardiopulmonary bypass <ul style="list-style-type: none"> - Runaway pump head - Trapped ventricular gas - Bubble oxygenator - Line disconnection • Carotid endarterectomy • Laparoscopic surgery • Liver transplantation • Transthoracic needle bx 	<ul style="list-style-type: none"> • Central venous catheters <ul style="list-style-type: none"> - Insertion - Aspiration • Mechanical ventilation • Neurosurgery • Dental implant surgery • Nd:Yag laser surgery • Thoracotomy
---	--

6

Cerebral Arterial Gas Embolism

...Iatrogenic events - continued

- Balloon angioplasty
- Endometrium resection
- Closed wound suction kit
- Hysteroscopy
- Arthroscopy
- Transurethral prostate resection
- Hepatic protoenterostomy
- Peripheral venous catheter
- Knee arthrography
- Water jet dissector
- CPR
- Angioplasty
- Bronchoscopy
- Endoscopy
- Hip replacement
- Intra-aortic balloons
- Diagnostic monitoring
- Hemodialysis

7

Cerebral Arterial Gas Embolism

...Right to left shunting of bubbles

Patent foramen ovale

Thompson & Evans, 1930
Moorthy & Losasso, 1974
Verstappen et al, 1977
Gronert et al, 1979

Pulmonary oxygen toxicity

Butler & Hills, 1981

Vasodilators

Butler & Hills, 1985

Arteriovenous shunts

Spencer & Oyama, 1971

8

Cerebral Arterial Gas Embolism

...Right to left shunting of bubbles, continued

Bubble volume overload

Butler & Hills, 1985
Spencer & Oyama, 1971

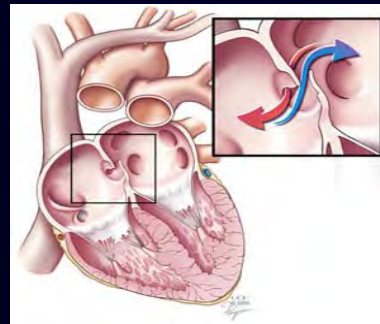
Elevated pulmonary artery pressures

Verstappen et al, 1977
Gronert et al, 1979

9

Cerebral Arterial Gas Embolism

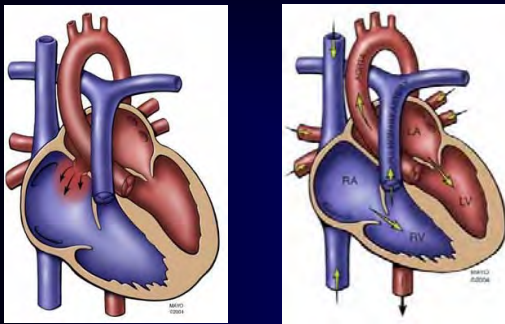
...Patent foramen ovale



10

Cerebral Arterial Gas Embolism

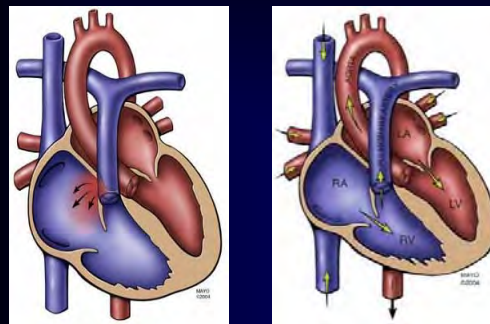
...Patent foramen ovale: septum secundum



11

Cerebral Arterial Gas Embolism

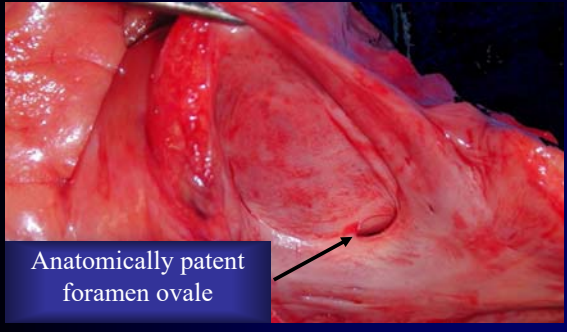
...Patent foramen ovale: septum primum



12

Cerebral Arterial Gas Embolism

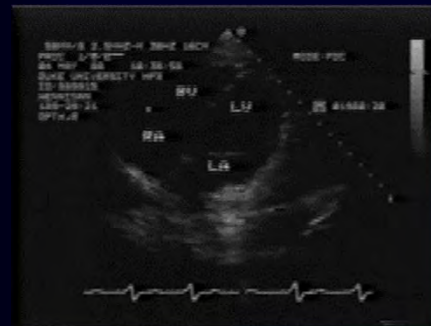
...Patent foramen ovale



13

Cerebral Arterial Gas Embolism

...Bubble Contrast Echocardiogram



14

Cerebral Arterial Gas Embolism

...TEE evidence of transpulmonary PAE

Massive VGE associated with open brain surgery in the sitting position.

"Air continued to traverse the left atrium per TEE for 15 minutes after all venous entrainment had ceased."

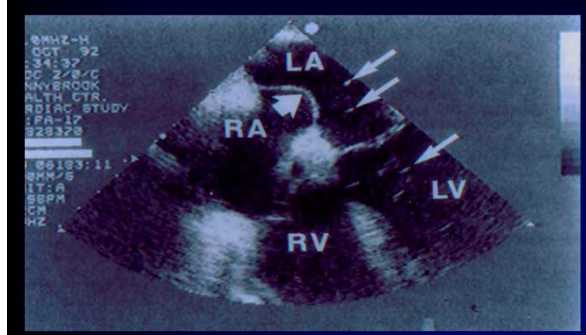
- Bedell EA, et al. 1994 Anaesthesiol: 80(4)

- PFO ruled out by negative findings on:
 - Multiple negative bubble contrast studies
 - Color flow doppler
 - Right-to-Left atrial septal bulging upon PEEP release

15

Cerebral Arterial Gas Embolism

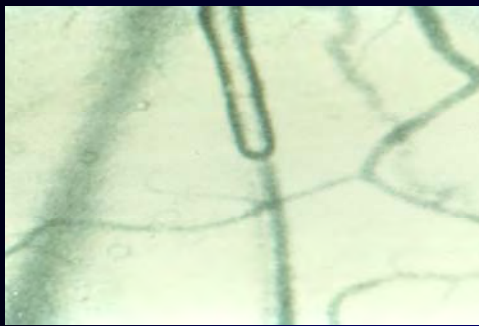
...Septal displacement on PEEP release



16

Cerebral Arterial Gas Embolism

...Vascular bubbles



17

Cerebral Arterial Gas Embolism

...Spontaneous redistribution

Evidence and mechanism of spontaneous redistribution

- Gorman DF 1986 UBR: 13(3): 317-335

- Bubble size less than 500 microns
 - Entrapment never occurred
 - Some evidence of vasoreactivity
- Bubble size greater than 5,000 microns
 - Entrapment and local circulatory arrest inevitable

18

Cerebral Arterial Gas Embolism

...Spontaneous redistribution, continued

Evidence and mechanism of spontaneous redistribution

- Gorman DF 1986 UBR; 13(3): 317-335

- Bubble size between 500 – 5,000 microns
 - Entrapment common but redistribution occurred within 3 minutes
 - Entrapment occurred most frequently in arterioles between 50 – 200 microns diameter.
 - Redistribution occurred only during the hypertensive period
 - Re-embolism occurred in 12% of observations

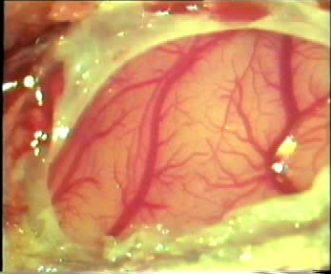
19

Cerebral Arterial Gas Embolism

...Spontaneous redistribution

Cerebral Arterial Gas in Rabbit Model

- Helps and Gorman



20

Cerebral Arterial Gas Embolism

...Decompression induced CAGE

Clinical correlations to cerebral arterial gas

- Helps SC, Gorman DF 1990 Stroke; 21: 94-99

- Five percent died
 - Permanent occlusion of pial vessels
- Thirty-five percent suffered sustained neurological injury
 - Temporary occlusion of pial vessels
- Sixty percent experienced spontaneous recovery of neurological function
 - Free transit of pial vessels

21

Cerebral Arterial Gas Embolism

...Decompression induced CAGE

Leukopenic animal model compared to controls

- Help SC, Gorman DF 1991 Stroke; 22: 351-354

It appears that decreases in both cerebral blood flow and brain function seen after embolism require the presence of leukocytes.

22

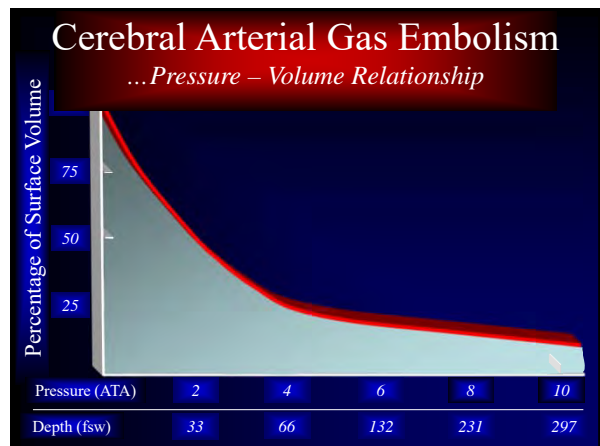
Cerebral Arterial Gas Embolism

...Decompression induced CAGE

Pressure and geometric proportions of bubbles

The danger of decompression-induced arterial gas embolism is greatest as the diver nears the surface.

23



24

Pressure Effects on Bubbles

Depth in Feet	Pressure in ATA	Relative Volume (%)	Relative Diameter (%)	Relative Surface Area (%)
0	1	100	100	100
33	2	50	79.4	63
66	3	33.3	69.4	48
99	4	25	63	39.7
132	5	20	58.6	34.3
165	6	16.7	55	30.3

25

Cerebral Arterial Gas Embolism

...Cause of death in divers

<i>Exhaustion, embolism, panic</i>	205
<i>Air embolism</i>	123
<i>Cave diving</i>	113
<i>Out of air at depth</i>	60
<i>Cardiovascular event</i>	48
<i>High waves or surf</i>	37
<i>Entrapped in kelp or weeds</i>	26
<i>Lost under ice</i>	22

- NUADC 1976-1985

26

Cerebral Arterial Gas Embolism

...Clinical significance

“Clinically significant cases of iatrogenic CAGE are under-diagnosed, under-treated, and under-reported”

- Tovar EA, et al. Ann. Thorac. Surg. 60, 1995

27

Cerebral Arterial Gas Embolism

...Neuro-protection

Stroke and cognitive decline – previously considered infrequent, and a small price to pay for life saving/function improving surgery.

Recent systematic analysis: these complications are common; more than half such patients so disabled at discharge.

Newman MF, et al. NEJM 344:2001

28

Cerebral Arterial Gas Embolism

...CPB as a source of embolization

Bubbles, per se, are injurious: it doesn't take a large volume to injure the brain.

Off-pump CABG does result in a reduction in micro-emboli and cerebral hypoperfusion.

- Diegeler A, et al. Ann. Thor. Surg; 69, 2000

However, there is no difference in the degree of cognitive decline at 12 months between on-and-off-pump procedures

- Van Dijk D, et al. JAMA; 287 (11), 2002

29

Cerebral Arterial Gas Embolism

...CPB as a source of embolization

“Perhaps the optimal approach to minimizing morbidity is to find effective neuro-protective agents”

Mark DB Newman MF JAMA : 287 (11) 2002

Most MEE's are probably gaseous

Stygall J, et al. Stroke; 31, 2000

- A double opportunity for hyperbaric medicine?
 - ~ treatment of a larger volume of patients
 - ~ preoperative neuro-protection

30

Cerebral Arterial Gas Embolism

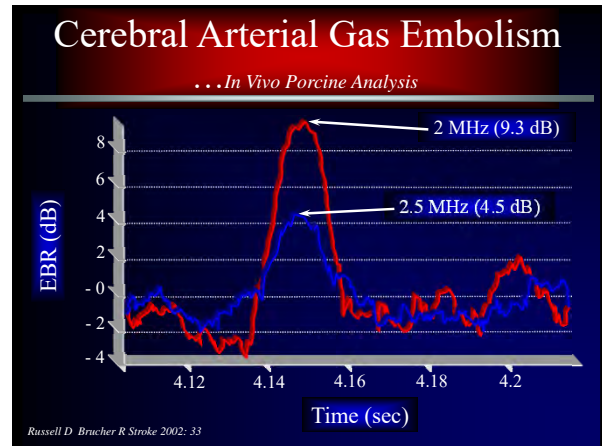
...Mechanical heart valves

"Most MEE's in mechanical heart valve patients are gaseous"

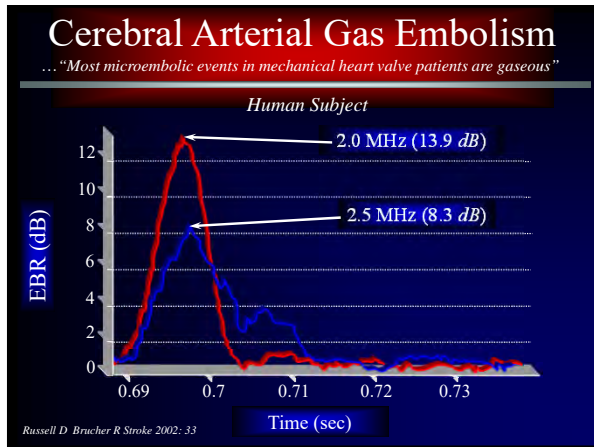
- Russell D. Brucher R Stroke 33, 2002

- 105 gas bubbles injected into a pig's cerebral circulation – 99 (94%) confirmed as gaseous, the balance as uncertain gaseous...none solid
- Right MCA microembolus in MHV patient identified as gaseous
- 433 of 514 emboli (84.2%) in 15 MHV pts. classed as gaseous, 74 (14.4%) as solid, and 7(1.4%) uncertain solid or gas

31



32



33

Cerebral Arterial Gas Embolism

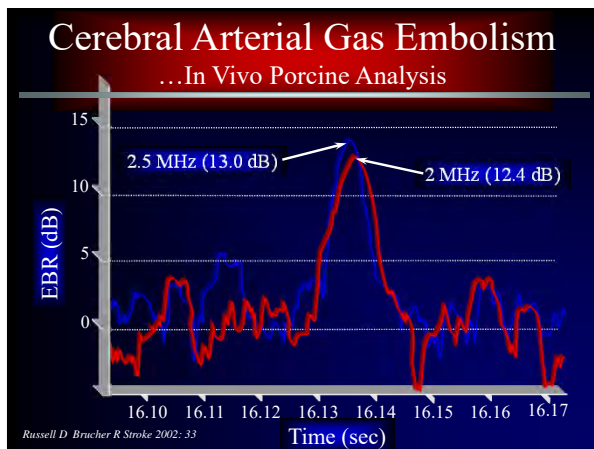
...Mechanical heart valves

"Most MEE's in carotid surgery pts. are solid"

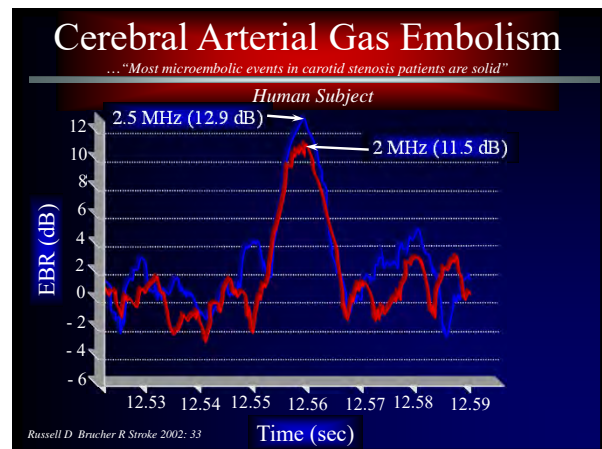
- Russell D. Brucher R Stroke 33, 2002

- 159 plastic microspheres injected into a pigs cerebral Circulation - 152 confirmed as solid (96%), the balance as uncertain solid ...none as gaseous
- Right MCA microembolus in a carotid stenosis patient identified as solid
- (94%) of 52 emboli in 17 carotid stenosis patients classified as solid, balance as uncertain...none as gaseous

34



35



36

Cerebral Arterial Gas Embolism

...Incidence of decompression illness

US divers only, 1997

- Divers Alert Network, 1999 Duke University Medical Center

• Type I DCS	222
• Type II DCS	598
• CAGE	68
• No breakdown	84
Total	972

37

Cerebral Arterial Gas Embolism

...Incidence during submarine escape training

	<u># of Cases</u>	<u>CAGE</u>	<u>Deaths</u>
<i>RN</i>	212,000	91	4
<i>USN</i>	250,000	44	8

Presentation in 91 cases of CAGE

• Coma	35%
• Stupor	23%
• Focal or hemi paresis	20%
• Somatic sensory changes	12%
• Visual disturbances	9%
• Collapse, without coma	8%

- Ah-See AK, Proc. VI Int. Cong. HBO 1977

38

Cerebral Arterial Gas Embolism

...Relapse In Decompression-Induced CAGE

While prompt recompression produces immediate relief in most cases, approximately 20% will later develop focal neurological deficits.

- Hallenbeck J, 1977

One third of those immediately recompressed relapsed within 20 minutes to several hours.

- Pearson RR, 1982

In 89 cases recompressed with prompt clearing of deficits, 27 (30%) suffered a relapse, or new signs/symptoms during latter stages of treatment.

- Ah-See AK, 1977

39

Cerebral Arterial Gas Embolism

...Neurological Response to Decreased Blood Flow

40

Cerebral Arterial Gas Embolism

...Comparison Of Tables 6A & 6; USN experience

Retrospective review of 2 yrs. prior to (6A) and 2 yrs. after (6) revision 3 of the USN Diving Manual.

<u>Pre-Revision (30)</u>	<u>Post-Revision (13)</u>
<ul style="list-style-type: none"> • 53% (16) treatment on TT 6A • 75% complete relief within 30 minutes @ 6 ATA • 88% complete relief prior to ascent to 1 ATA • 10% (3) with treatment delay >1hr. • 10% recurrence 	<ul style="list-style-type: none"> • 100% treatment on TT 6 • 54% complete relief during 1st O₂ cycle • 85% complete relief prior to ascent to 1 ATA • 23% (3) with treatment delay >1hr. • 0% recurrence

Howarth C, et al. UHM 24 (S147), 1997

41

Cerebral Arterial Gas Embolism

...Treatment: USN Table 6

Compression rate 25 ft/min
 Decompression rate 1 ft/min
 Total Elapsed time 285 min
 (with all extensions 485 min)

42

Cerebral Arterial Gas Embolism

...Adjunctive Therapies - Positioning

Modified Trendelenburg no longer recommended

-Dukta AJ, UHMS Workshop # 41, 1990

- Considered likely to compound neurological pathology, by increasing venous return and decreasing cerebral perfusion
- Maintain patient in the supine, or coma, position
- Advise not to strain or valsalva maneuvers

43

Cerebral Arterial Gas Embolism

...Adjunctive Therapies – Surface Oxygen

100% oxygen may relieve some symptoms and may reduce likelihood of others developing

- Corrects any hypoxemia
- Enhances inert gas elimination
- Reduces size of embolic gas
- Prophylaxes against DCS
- Treats established DCS

44

Cerebral Arterial Gas Embolism

...Adjunctive Therapies – Anti-platelet Drugs

There are more arguments against its use (aggravates inner ear or spinal cord hemorrhage; increases risk of dysbaric osteonecrosis)

- Limited evidence that agents such as aspirin modify platelet action upon decompression
- Aspirin “prophylaxis” in professional drivers!

45

Cerebral Arterial Gas Embolism

...Adjunctive Therapies – Fluid Resuscitation

Frequently indicated, particularly in acute setting

-Moon RE, Sheffield PJ, Aviat. Space Environ. Med. 68, 1997

- Hemo-concentration
- Platelet and PMNL accumulation/clumping
- Endothelial injury
- Glucose-free agents, unless hypoglycemia present

46

Cerebral Arterial Gas Embolism

...Adjunctive Therapies – Lidocaine

“There is sufficient evidence to justify lidocaine in clinical settings where DCI is highly likely”

- Mitchell SJ, 2001

- Neuroprotection thought to be afforded through several pathways:
 - modulates leukocyte activity
 - anti-inflammatory properties
 - reduces cerebral metabolism
 - decelerates ischemic ion shifts
 - reduces intracranial pressure

Mitchell SJ, et al. Ann. Thoracic Surg. 67, 1999 Francis J, SPUMS Journal, 32 (2), 2002

47

Cerebral Arterial Gas Embolism

...Adjunctive Therapies – Steroids

Data supporting reduction of edema and modification of inflammatory responses comes from traumatic and vascular brain injuries, not DCI

- Moon RE, SPUMS Journal, 2000

- Widespread acceptance in neuronal cases despite little convincing evidence

48

Cerebral Arterial Gas Embolism

...Adjunctive Therapies – NSAIDS

Data supporting reduction of edema and modification of inflammatory responses comes from traumatic and vascular brain injuries, not DCI

- Moon RE. SPUMS Journal, 2000

Large (180pts) RCT found that Tenoxicam, 20mg daily x 7, significantly reduced number of tx's necessary; no difference in outcomes.

- Bennett MH, et al. UHM ASM #29,2002

49

Cerebral Arterial Gas Embolism

... The spontaneously "recovered" diver

- Relapse is common, and unpredictable
- Suppression of evolution of concurrent DCS
- Antagonism of leukocyte-mediated I-R injury
- Protection/treatment of simultaneous embolism elsewhere
- Limiting/resolving cytotoxic and vasogenic edema
- Protection against cerebral infarcts

- Clarke D, et al. Aviat. Space Environ Med; 73, 2002

50

Cerebral Arterial Gas Embolism

...Delayed treatment

Excellent Clinical Outcome In Delayed Hyperbaric Treatment Of CAGE

- 23 hours - L. Lau, 1991
- 24 hours - J. Mader, 1979
- 30 hours - C. Armon, 1991
- 42 hours - E. Massey, 1990
- 52 hours - C. Wherrett, 2002
- 60 hours - H. Bitterman, 1993

51

Cerebral Arterial Gas Embolism

... Cerebral Infarcts

CT's of untreated cerebral arterial gas embolism have demonstrated areas of brain infarct, even in patients with apparent complete clinical recovery

Kearns PT, West J. Med.;140:615-617, 1984
Warren LP, Am J Radiol.;151:1003-1008, 1988

52

Cerebral Arterial Gas Embolism

...Treatment outcomes

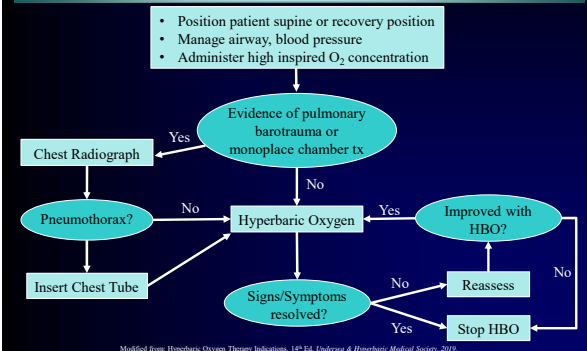
Outcome	HBO	NO HBO
Full Recovery	346 (84.24%)	74 (24.7%)
Residual	45 (10.9%)	63 (21.9%)
Death	20 (4.9%)	151 (52.4%)
Total	411 (100%)	288 (100%)

Dutka AJ, UHMS Critical Review, 1991

53

Cerebral Arterial Gas Embolism

...Treatment Flowchart



Modified from: Hyperbaric Oxygen Therapy Indications, 14th Ed. Undersea & Hyperbaric Medical Society, 2008

54

REFERENCES

- Ah-See AK: **Review of arterial air embolism in submarine escape.** Proceed. Sixth Int. Congress Hyperbaric Medicine. 1977; Aberdeen University Press 349-351.
- Armon C, et al.: **Hyperbaric treatment of cerebral air embolism sustained during an open-heart surgical procedure.** May Clinic Proceedings 1991;66:565-571
- Bedell EA, et al.: **Paradoxical air embolism during venous air embolism: transesophageal echocardiographic evidence of transpulmonary air passage.** Anesthesiology 1994;80(4):947-950
- Bennett MH, et al.: **Adjunctive treatment of decompression illness with a non-steroidal anti-inflammatory drug (Tenoxicam) reduces compression requirement (Abstract).** Undersea Hyper Med 2002;29(2)S:29.
- Benton PJ, et al.: **Arterial gas embolism following a 1-meter ascent during helicopter escape training: a case report.** Aviation, Space and Environmental Medicine 1996;67(1):63-64
- Bitterman H, et al.: **Delayed hyperbaric treatment of cerebral air embolism.** Israel Journal Medical Science 1993;29(1):22-26
- Butler BD, et al.: **Effect of Excessive Oxygen Upon the Capability of the Lungs to Filter Gas Emboli.** Proceedings: 7th Symposium on Underwater Physiology 1981. Undersea & Hyperbaric Medical Society.
- Butler BD, et al.: **Transpulmonary passage of venous air emboli.** J. Applied Physiology 1985;59(2):543-547.
- Clarke D, et al.: **Pulmonary barotrauma-induced cerebral arterial gas embolism with spontaneous recovery: commentary on the rationale for therapeutic compression.** Aviat Space Environ Med 2002;73(2):139-146.
- Diegeler A, et al.: **Neuromonitoring and neurocognitive outcome in off-pump versus conventional coronary bypass operation.** Ann Thorac Surg 2000;69:1162-1166.
- Divers Alert Network: **1999 Report on decompression illness and diving fatalities.** DAN Box 2823, Duke University Medical Center, Durham, North Carolina.
- Dutka AJ: **Air or gas embolism.** In Hyperbaric Oxygen Therapy: A Critical Review. Eds. EM Camporesi and AC Barker 1991:1-10.
- Dutka AJ: **Diving Accident Management Proceedings.** 41st UHMS Workshop 1990. Undersea and Hyperbaric Medical Society.
- Francis J: **A randomised prospective trial of lignocaine in the management of acute neurological decompression illness – an update.** SPUMS J 2002;32(2):97-105.
- Gorman DF, et al.: **Cerebral Vasoreactivity and Arterial Gas Embolism.** UBR 1986;13(3):317-335.
- Gronert GA et al: **Paradoxical Air Embolism From a Patent Foramen Ovale.** Anesthesiology 1979;50:548-549.
- Hallenbeck JM, et al.: **Impaired microvascular perfusion and secondary deterioration in dysbaric cerebral air embolism.** 13th Ums Workshop on Arterial Air Embolism; May 1977, Toronto, Canada
- Helps SC, et al.: **Air Embolism of the Brain in Rabbits Pretreated With Mechlorethamine.** Stroke, 1991; 22:351-354.
- Helps SC, et al.: **The effect of gas emboli on rabbit cerebral blood flow.** Stroke, 1990;21(1):94-99.
- Hills BA: **Pulmonary Barotrauma: A Possible Role For Surfactant in Opposing the Entry of Air Into the Circulation.** SPUMS Journal 1993;23(2):59-64.
- Howsare C, et al.: **Comparison of USNTT6 vs. TT6A for the treatment of AGE in the US Navy. A retrospective study. (Abstract)** Undersea and Hyperbaric Medicine 1997; 24(Supplement):147
- Kearns PJ, et al.: **Homonymous hemianopia due to cerebral air embolism from central venous catheters.** The Western Journal of Medicine 1984;140:615-617
- Lau K, et al.: **Systemic air embolism: a complication of ventilator therapy in hyaline membrane disease.** Clinical Radiology 1991;43:16-18
- Leitch D, et al.: **Cerebral arterial air embolism: 1. Is there benefit in beginning HBO treatment at 6 bar?** Undersea Biomed. Research 1984;11(3):221-235.
- Mader JT, et al.: **Delayed hyperbaric treatment of cerebral air embolism.** Archives of Neurology 1979;36:504-505
- Mark DB, et al.: **Protecting the brain in coronary artery bypass graft surgery (Editorial).** JAMA 2002;287(11):1448-1450.
- Masey EW, et al.: **Hyperbaric Oxygen Therapy of Iatrogenic Air Embolism.** J. Hyperbaric Medicine 1990;5(1):15-21.
- Mitchell SJ: **Lidocaine in the treatment of decompression illness: a review of the literature.** Undersea Hyper Med 2001;28(3):165-174.

- Mitchell SJ, *et al*: **Cerebral protection by lidocaine during cardiac operations.** Ann Thorac Surg 1999;67:1117-1124.
- Moon RE: **Adjunctive therapy in decompression illness: Present and future.** SPUMS 2002;30(2):99-112.
- Moon RE, *et al*: **Guidelines for treatment of decompression illness.** Aviat., Space & Environ Med 1997;68(3):234-243.
- Moorthy SS, *et al*: **Patency of the Foramen Ovale in the Critically Ill Patient.** Anesthesiology 1974;41(4):405-407.
- Newman MF, *et al*: **Longitudinal assessment of neurocognitive function after coronary-artery bypass surgery.** New England J Med 2001;344:395-402.
- Oyama Y, *et al*: **Cardiopulmonary effects of intravenous gas embolism; with special reference to fate of intravascular gas bubbles.** Japanese Circ J 1971;35(12):1541-1549.
- Pearson RR, *et al*: **Delayed cerebral edema complicating cerebral arterial gas embolism: case histories.** Undersea Biomed Research 1982;9(4):283-296.
- Polak B, *et al*: **Traumatic air embolism in submarine escape training.** US Naval Medical Bulletin 1932;30:165-177
- Stoney WS, *et al*: **Air embolism and other accidents using pump oxygenators.** Annals Thoracic Surgery. 1980;29(4):336-340.
- Stonier JC: **A study in prechamber treatment of cerebral air embolism patients by a first provider at Santa Catalina island.** Undersea Biomedical Research 1985;12:ABST. 58
- Stygall J, *et al*: **Cerebral microembolism detected by transcranial doppler during cardiac procedures.** Stroke 2000;31:2508-2510.
- Thompson T, *et al*: **Paradoxical Embolism.** Quarterly Journal of Medicine,1930;23:135-152.
- Tovar EA, *et al*: **Postoperative management of cerebral air embolism: gas physiology for surgeons.** Annals of Thoracic Surgery 1995;60:1138-1142
- **US Navy Diving Manual, 1999, Revision 4.** Published by Direction of Commander, Naval Sea Systems Command, Washington, DC 20362.
- Van Diik D, *et al*: **Cognitive outcome after off-pump and on-pump coronary artery bypass graft surgery.** JAMA 2002;287(11):1405-1412.
- Verstappen FTJ, *et al*: **Effects of Pulmonary Gas Embolism on Circulation and Respiration in the Dog.** European Journal of Physiology 1977;370:67-70.
- Warren LP, *et al*: **Neuroimaging of scuba diving injuries to the CNS.** AJR 1988;151:933-938:1003-1008
- Wherrett CG, *et al*: **Cerebral arterial gas embolism following diagnostic bronchoscopy: delayed treatment with hyperbaric oxygen.** Can J Anesth 2002;49(1):96-99.

Additional Reading

- Edmonds C, Lowry C, Pennefather J, Walker R: **Diving and Subaquatic Medicine, Fourth Edition, 2002.** Oxford University Press.
- Bove AA: **Diving Medicine, Third Edition, 1997.** W. B. Saunders Company.