

Pulmonary Problems at High Altitude

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I have nothing to disclose



Objectives

- To review the basic physiological changes that occur at high terrestrial altitude
- To describe the presentation and management of high altitude illnesses
- To discuss the exacerbation of sea level illnesses at high terrestrial altitude



Definitions

- Moderate Altitude: 5,000 - 8000 ft
- High Altitude: 8,000 - 14,000 ft
- Very High Altitude: 14,000 - 18,000 ft
- Extreme Altitude: 18,000 - 29028 ft

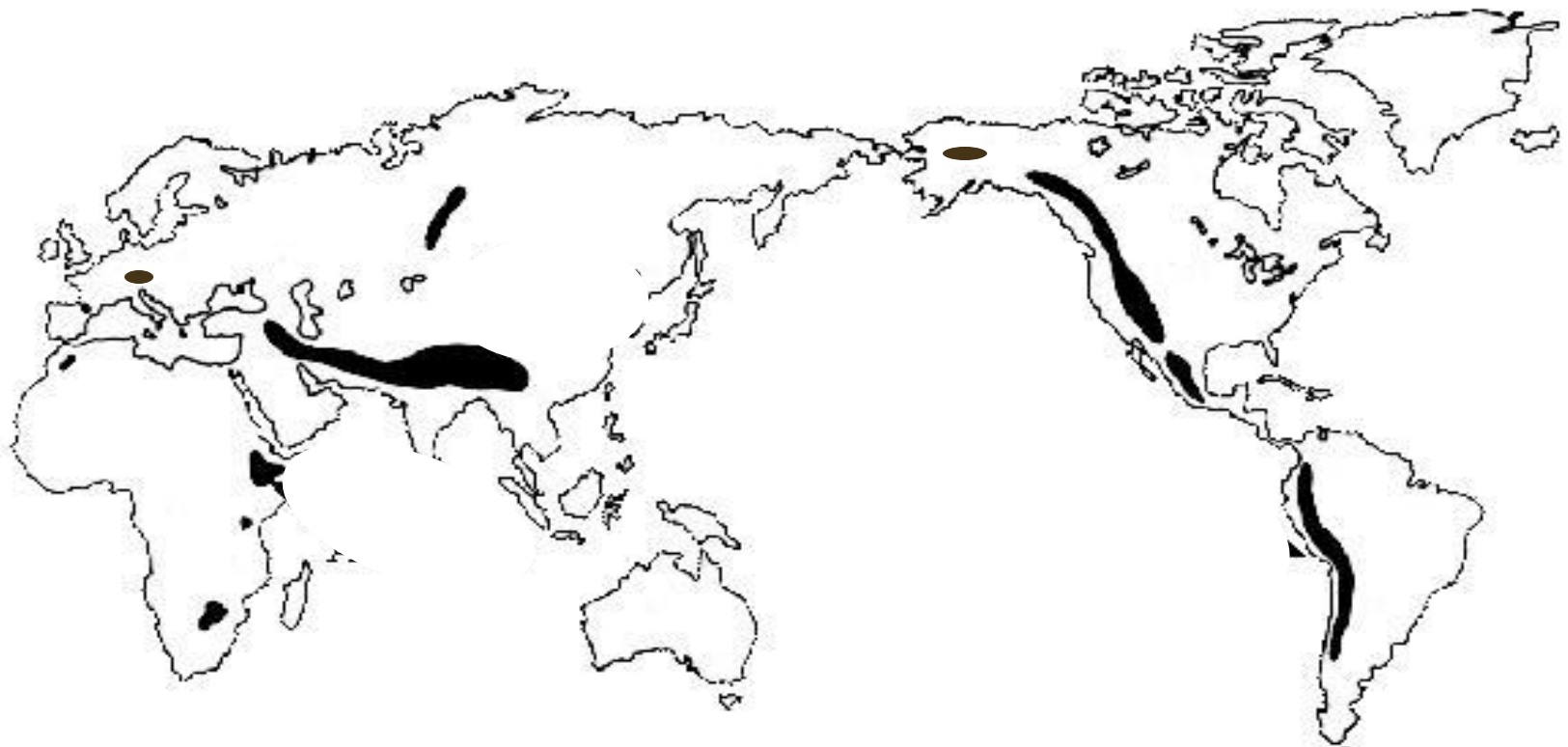




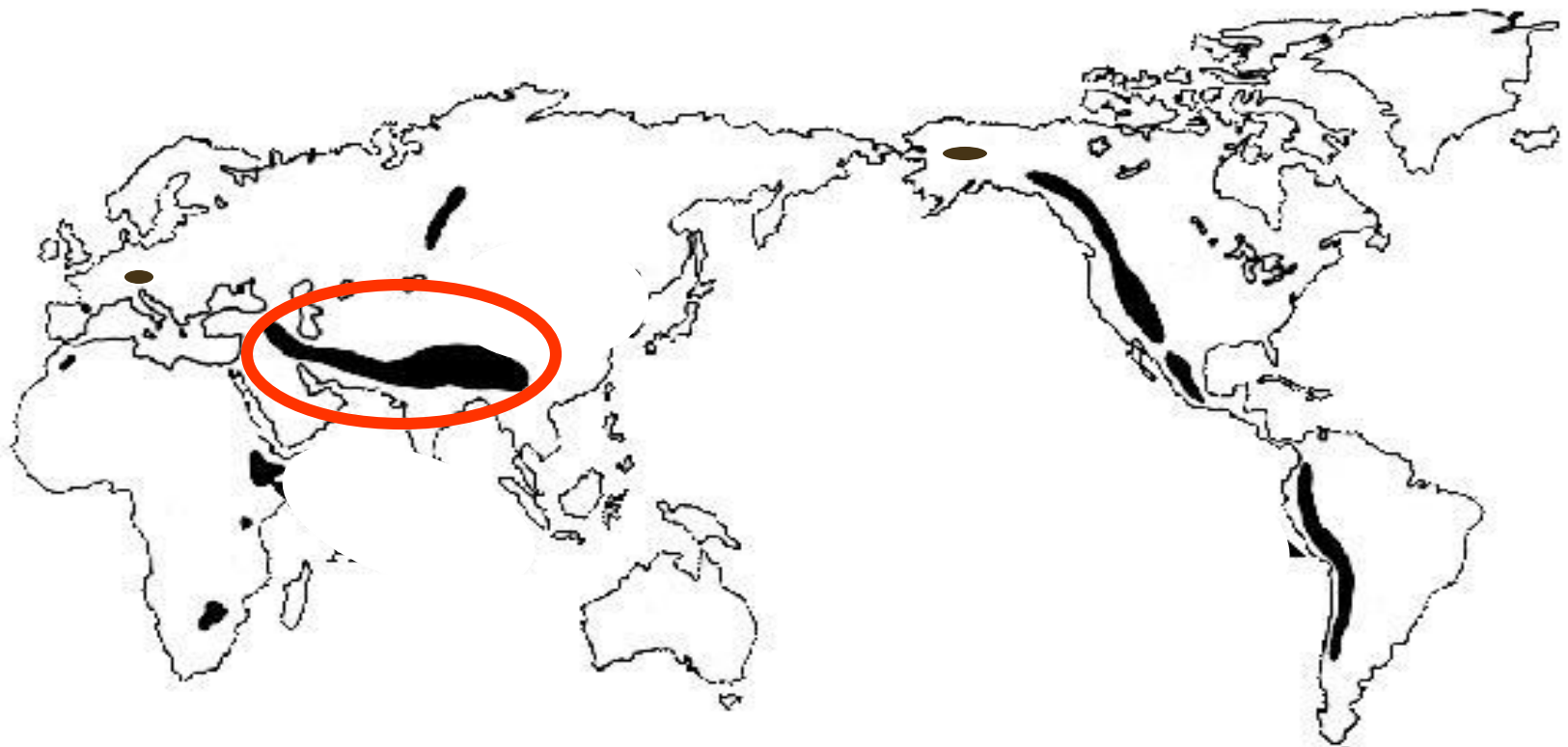
High Altitude Illnesses

- Serious Illnesses
 - High Altitude Pulmonary Edema
 - High Altitude Cerebral Edema
- Disabling Illnesses
 - Acute Mountain Sickness
 - Chronic Mountain Sickness
- Exacerbation of Sea Level Illnesses
 - COPD
 - Restrictive or interstitial lung disease
 - Pulmonary Hypertension





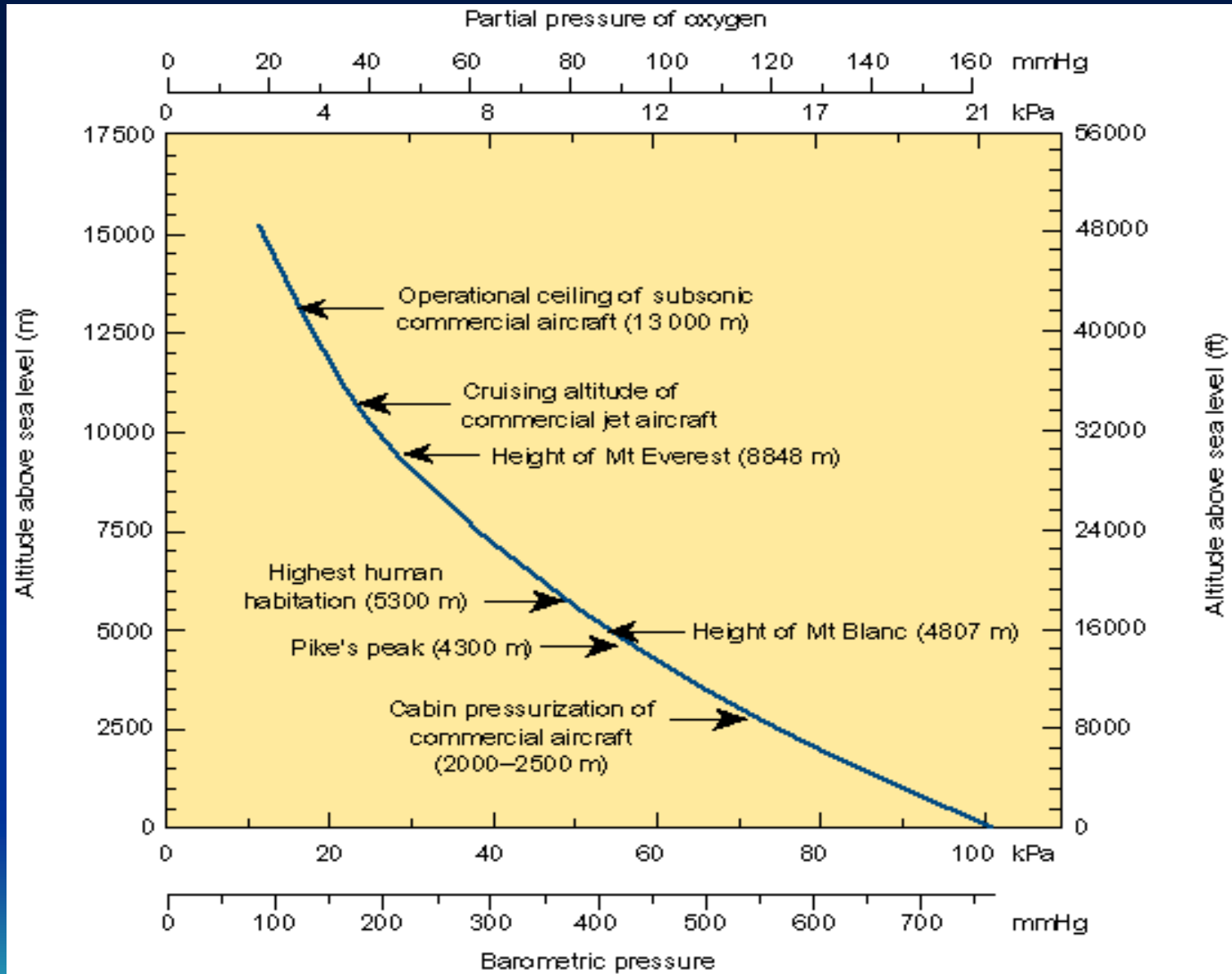
Areas of the world over 3,000 meters elevation (Frisancho 1993:224)

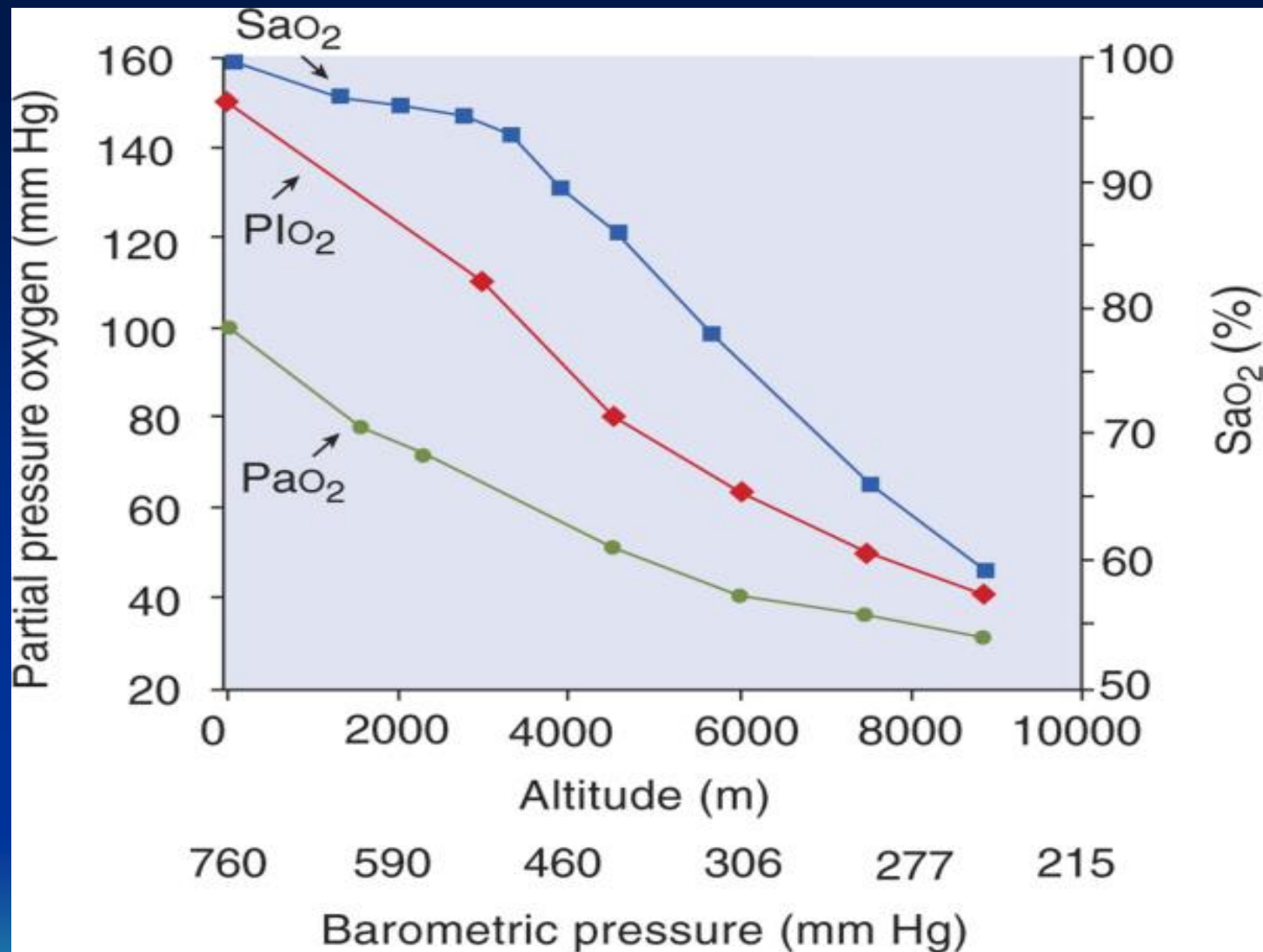


Areas of the world over 3,000 meters elevation (Frisancho 1993:224)

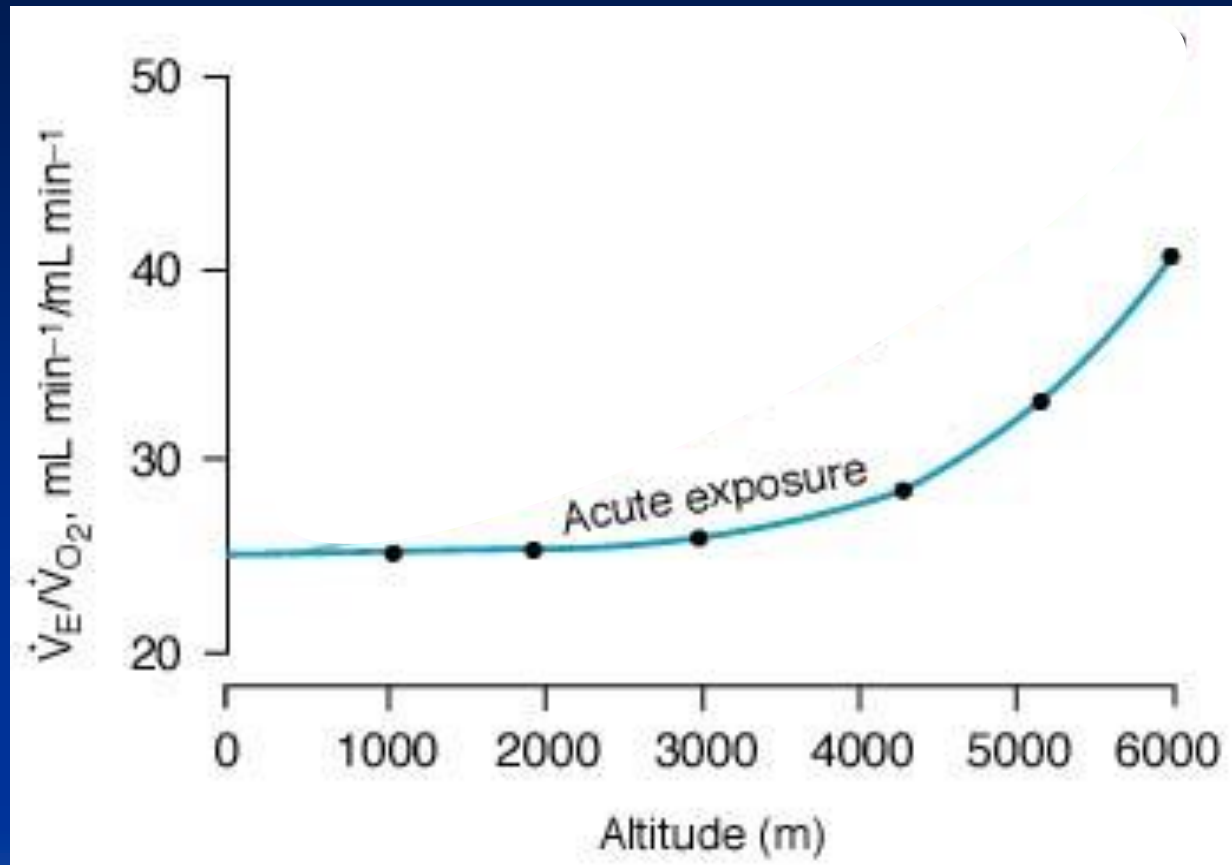
Basic Physiological Changes at High Altitude



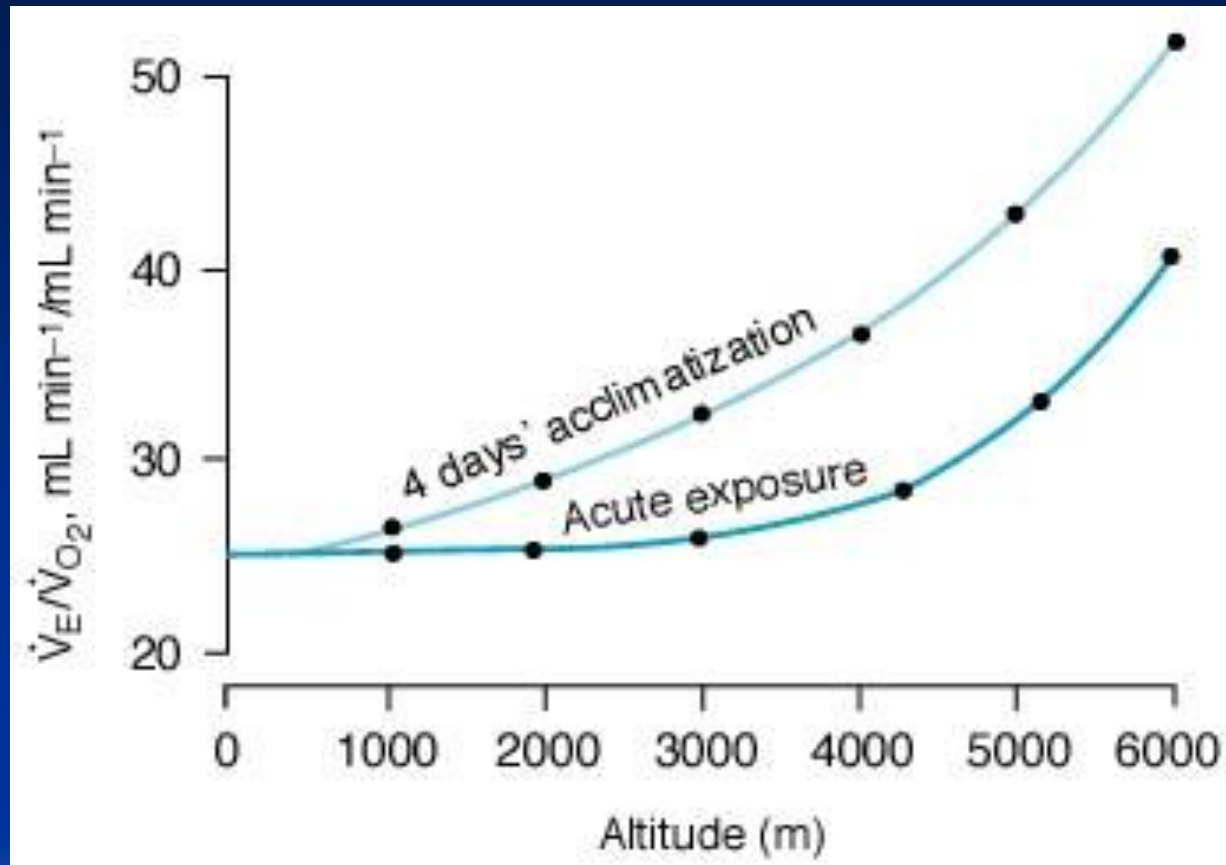




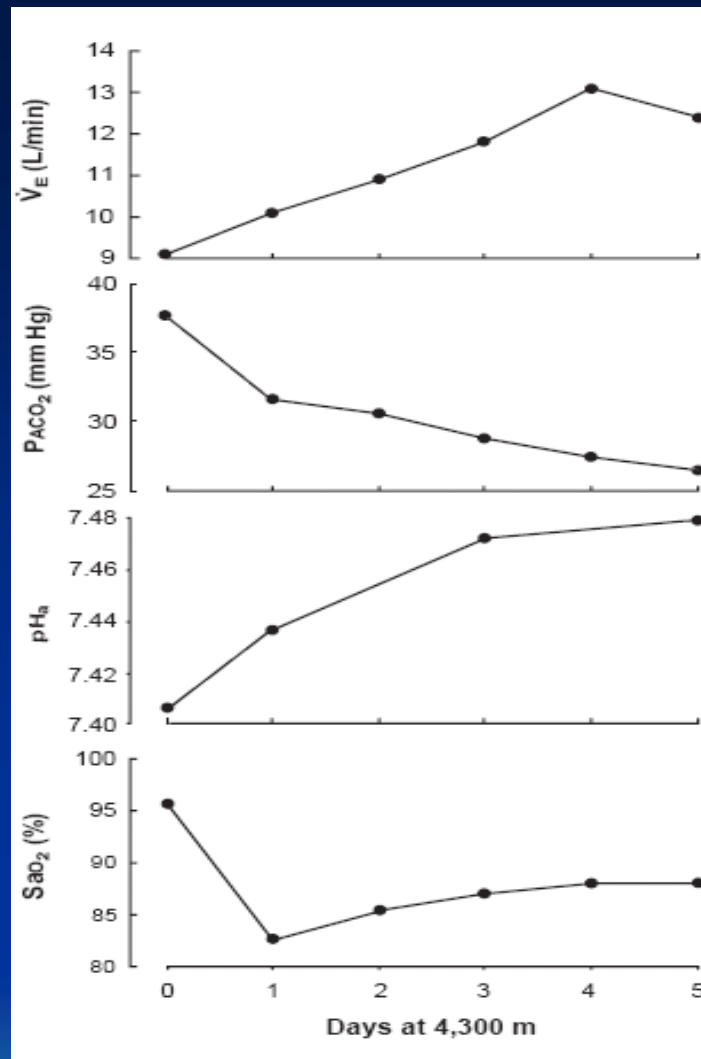
Sutton JR et al: J Appl Physiol 64:1309, 1988.



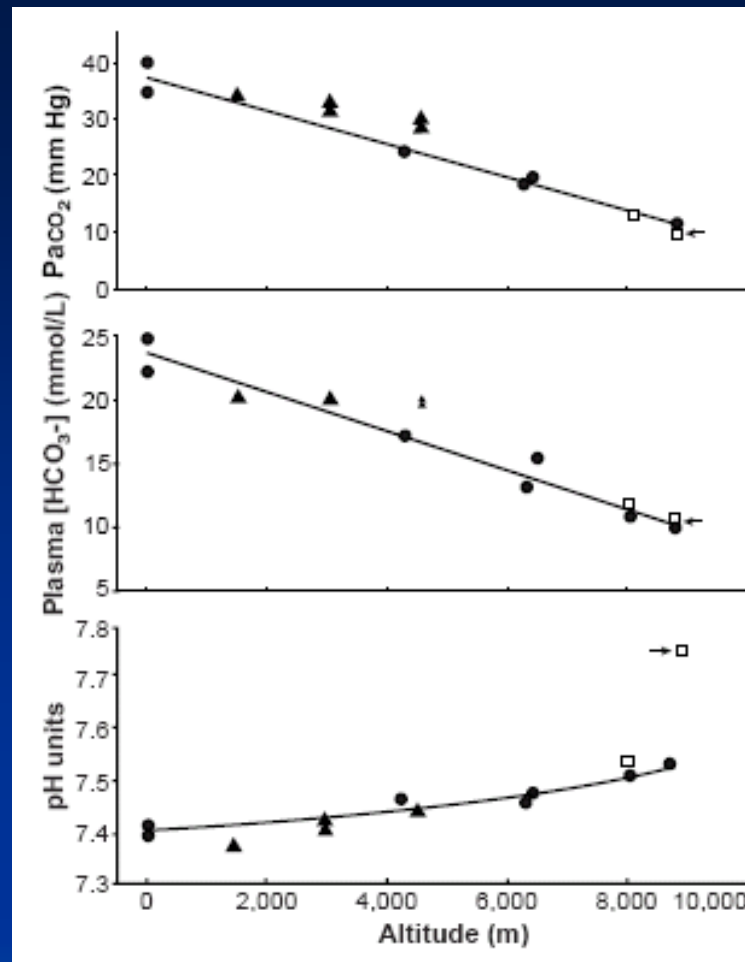
Lenfant C, Sullivan K. Adaptation to high altitude. *N Engl J Med*. 1971;284:1303.



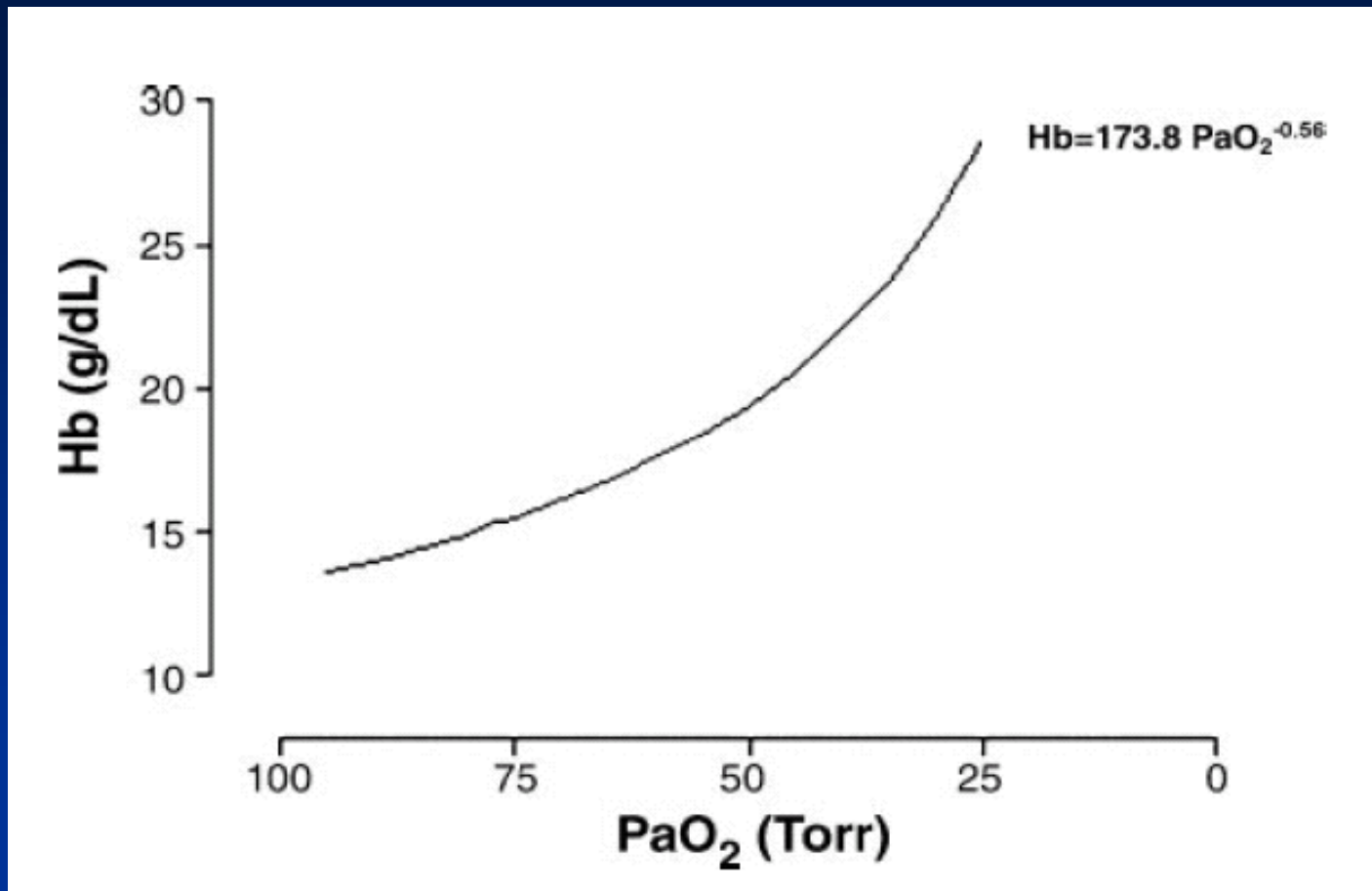
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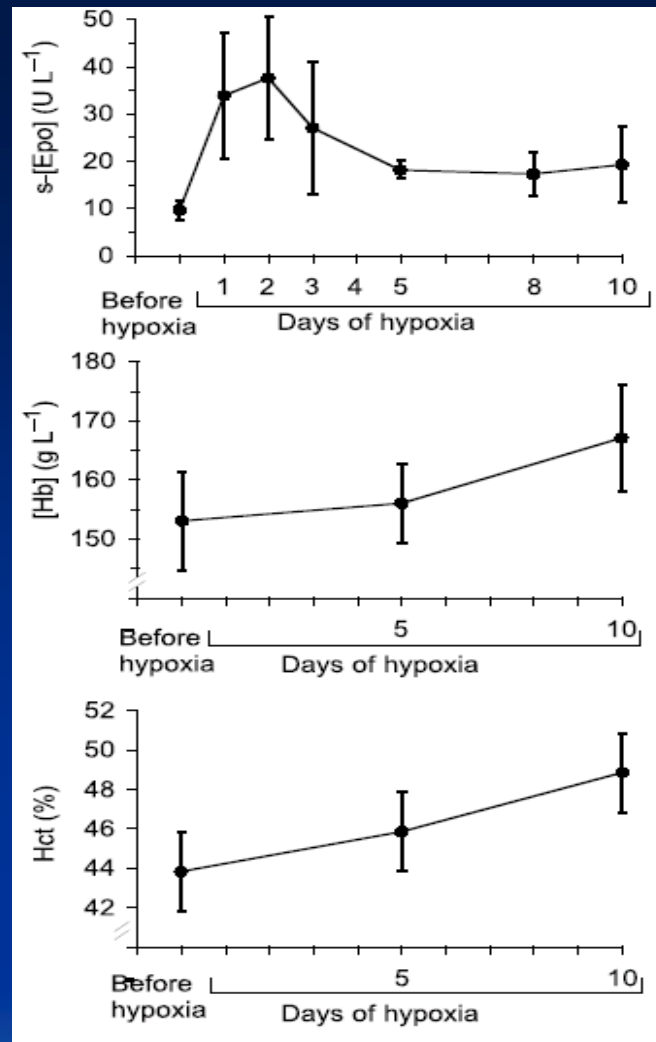
Huang SY, Alexander JK, Grover RF, et al. Hypocapnia and sustained hypoxia blunt ventilation on arrival at high altitude. *J Appl Physiol.* 1984;56:603.



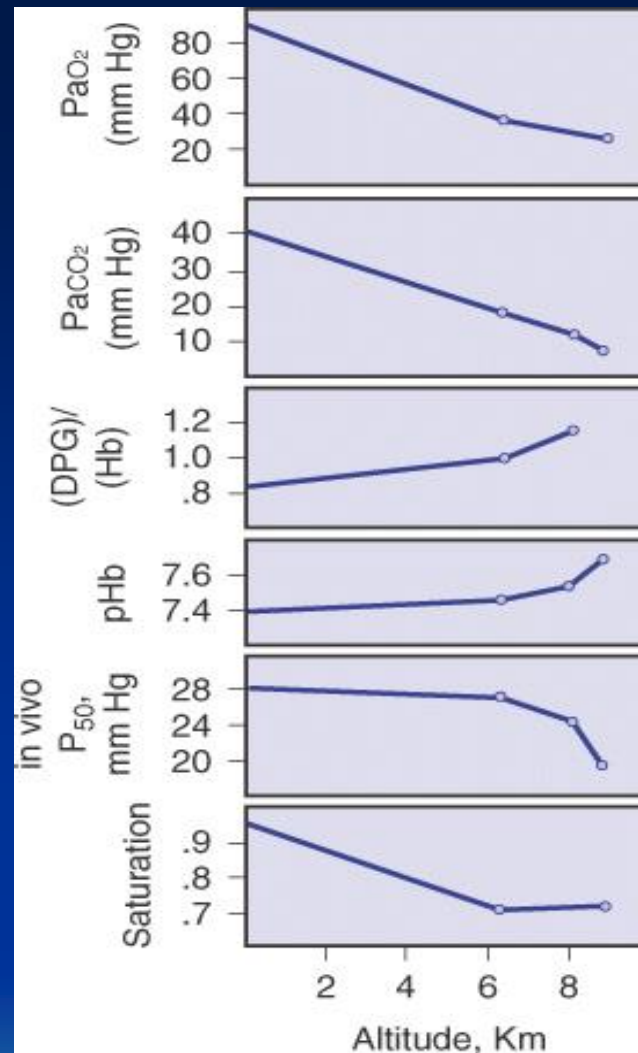
Sutton JR, Reeves JT, Wagner PD, et al. Operation Everest II: Oxygen transport during exercise at extreme simulated altitude. *J Appl Physiol.* 1988;64:1309–1321.



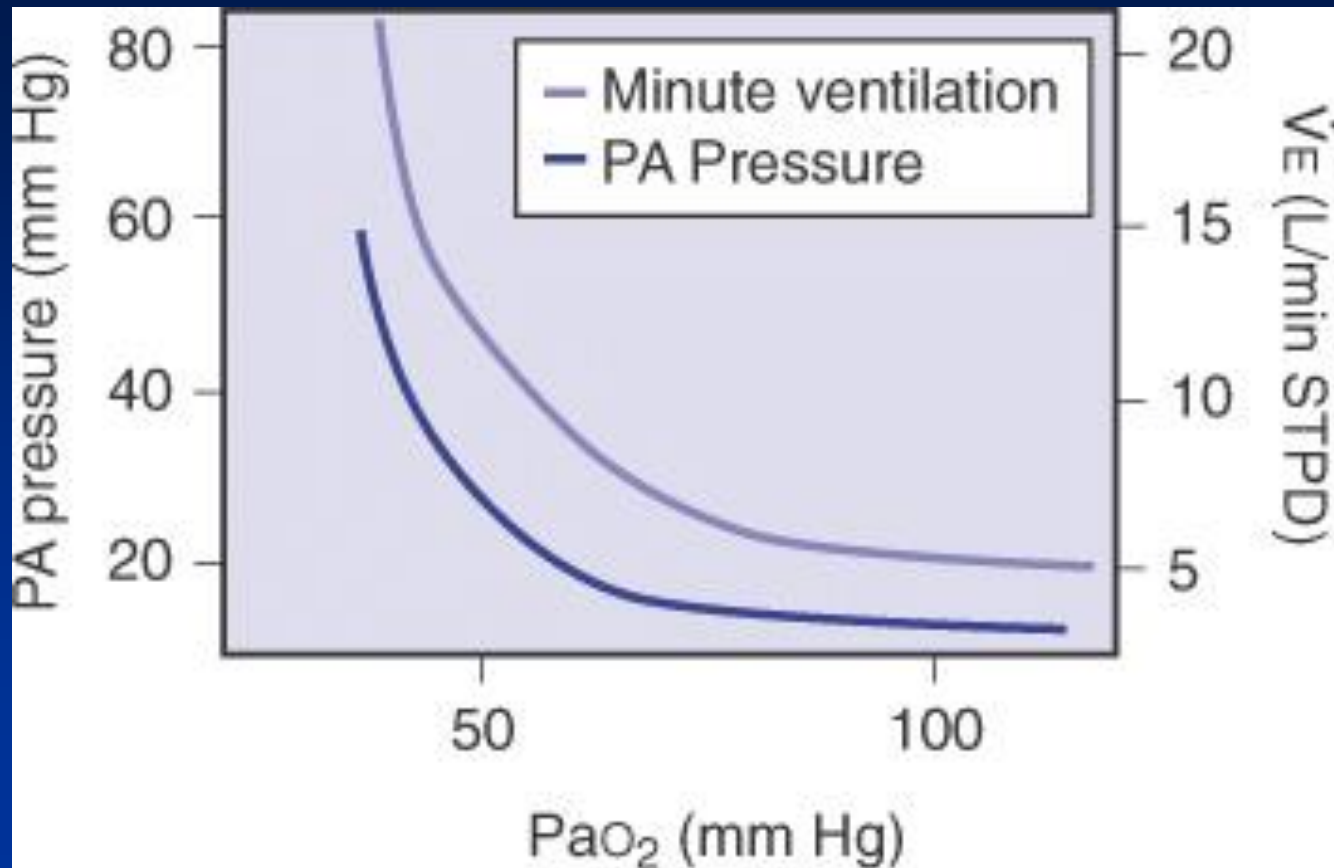
Villafuerte FC, Cardenas R, Monge CC. Optimal hemoglobin concentration at high altitude: A theoretical approach for Andean men at rest. *J Appl Physiol* 96: 1581–1588, 2004.



Berglund B, Genser M, Ornhagen H, et al. Erythropoietin concentrations during 10 days of normobaric hypoxia under controlled environmental circumstances. *Acta Physiol Scand* 2002, 174, 225-229.



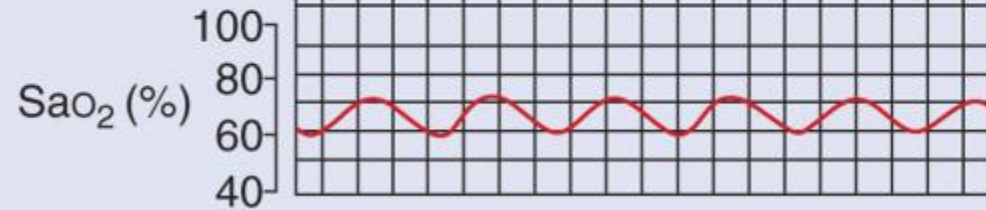
Winslow RM, Samaja M, West JB: Red cell function at extreme altitude on Mount Everest. J Appl Physiol 56:109–116, 1984



Reeves JT, Wagner, WW Jr, McMurtry F, et al: Physiological effects of high altitude on the pulmonary circulation. *In* Robertshaw D [ed]: International Review of Physiology, Environmental Physiology III. Baltimore: University Park Press, 1979, pp 289-310.

Placebo

Respiratory Pattern

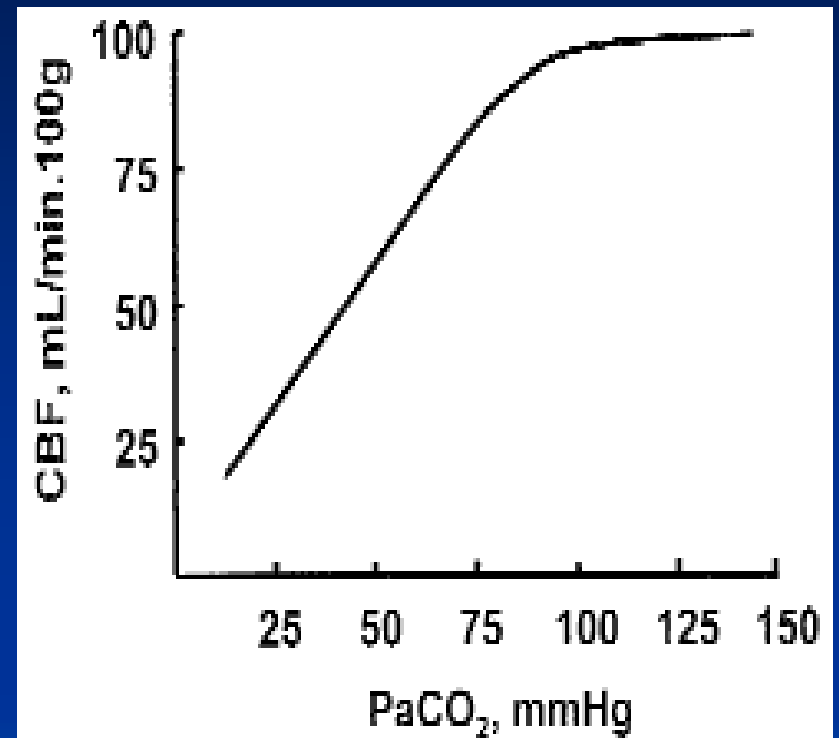
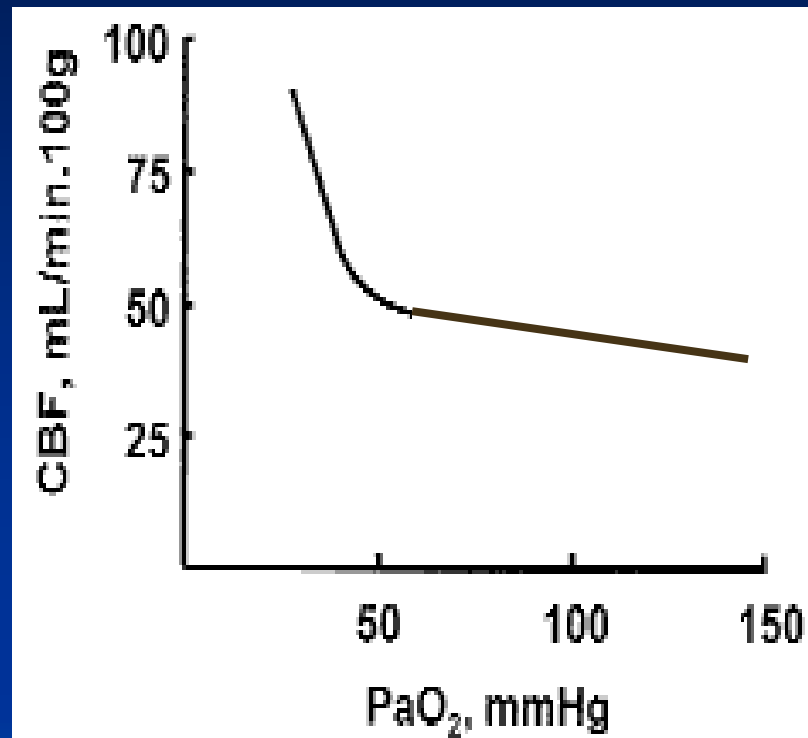


Acetazolamide

Respiratory Pattern



Hackett PH et al: Am Rev Respir Dis 135:896, 1987.



Acute Mountain Sickness



Acute Mountain Sickness

Symptoms

- Headache - Always Present
- Plus one or more of the following:
 - Nausea
 - Vomiting
 - Fatigue or weakness
 - Dizziness or lightheadedness
 - Difficulty Sleeping

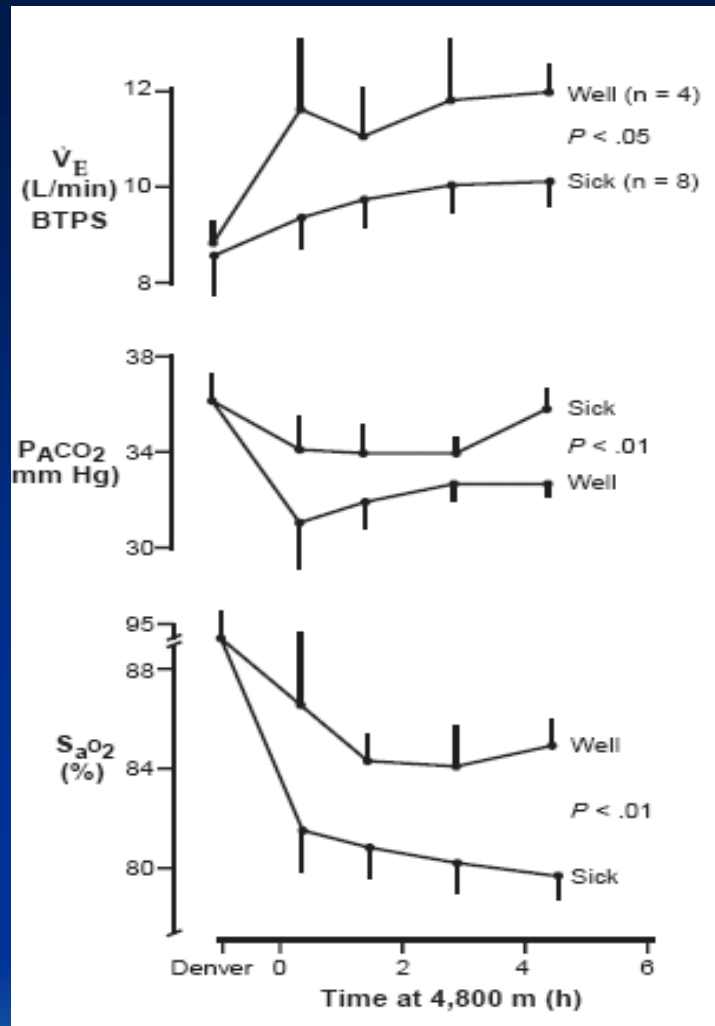


Acute Mountain Sickness

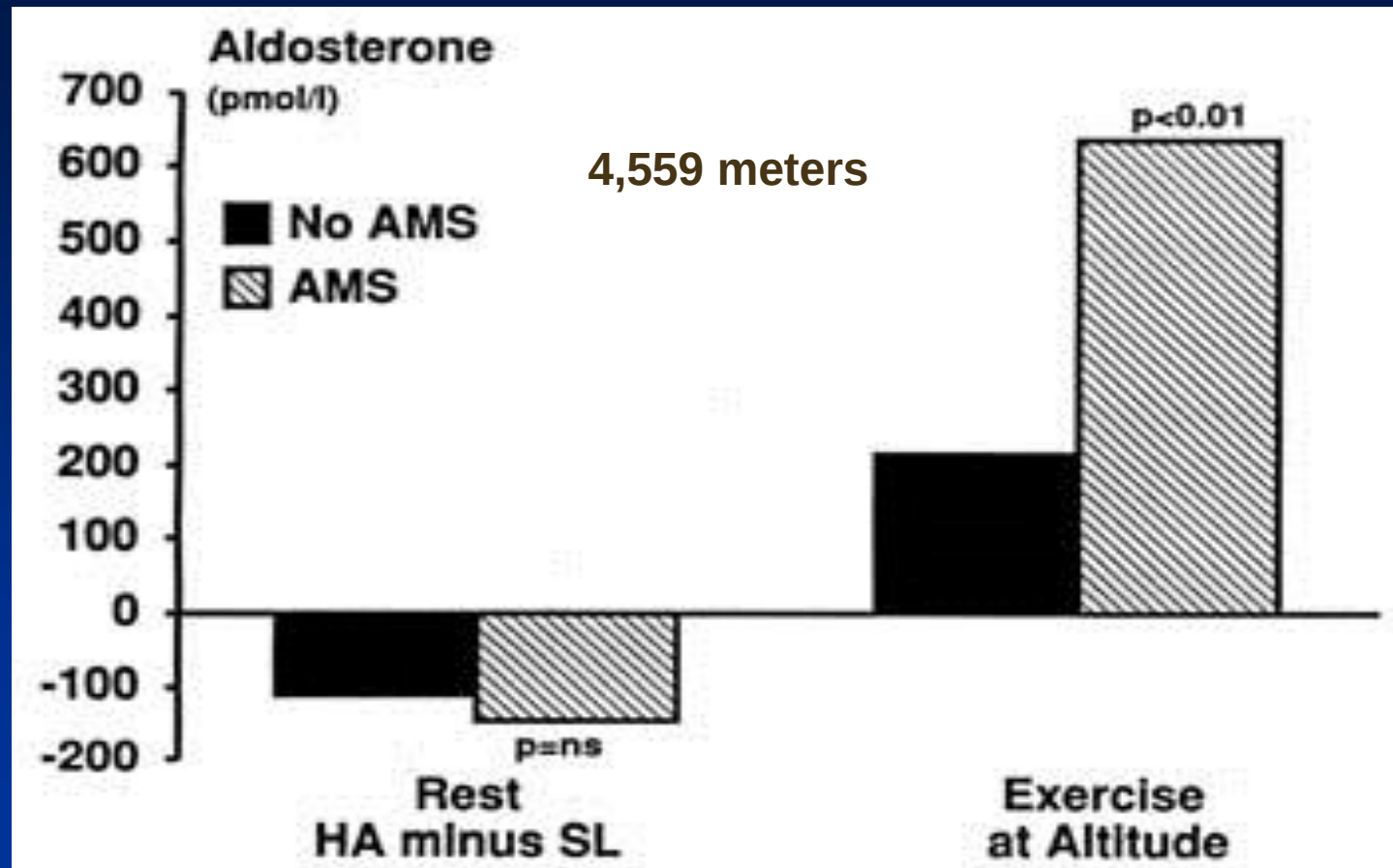
Epidemiology

- Typically occurs at altitudes > 8,000 ft
- Affects 50-60% of individuals at altitudes > 10,000 ft
- Symptoms are more severe with rapid ascent and exercise
- Symptoms are most severe on the 2nd and 3rd days after acute ascent
- Symptoms typically resolve after 3 to 5 days, as one becomes acclimatized.

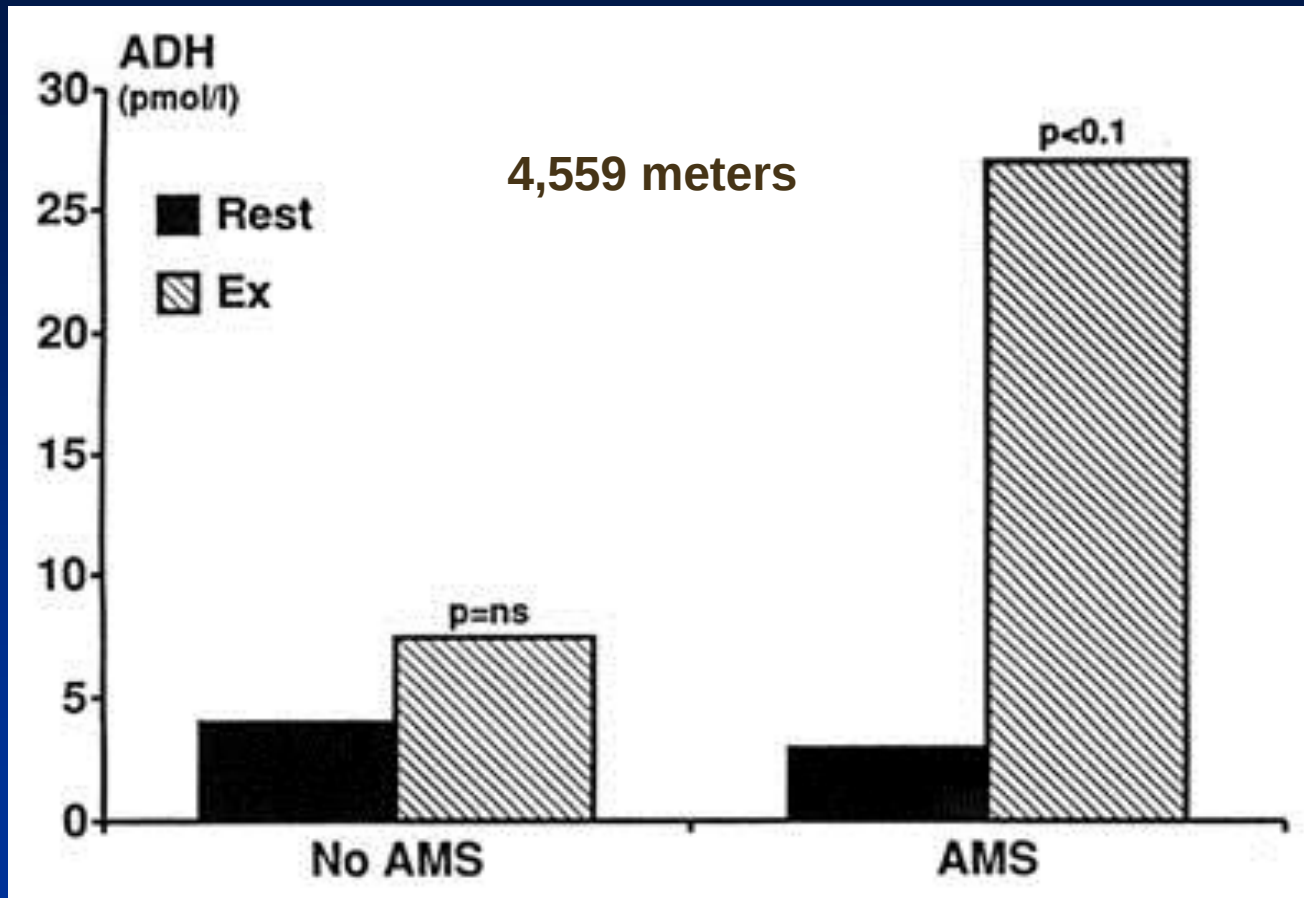




Moore LG, Harrison GL, McCullough RE, et al. Low acute hypoxic ventilatory response and hypoxic depression in acute altitude sickness. *J Appl Physiol.* 1986;60:1407–1412.

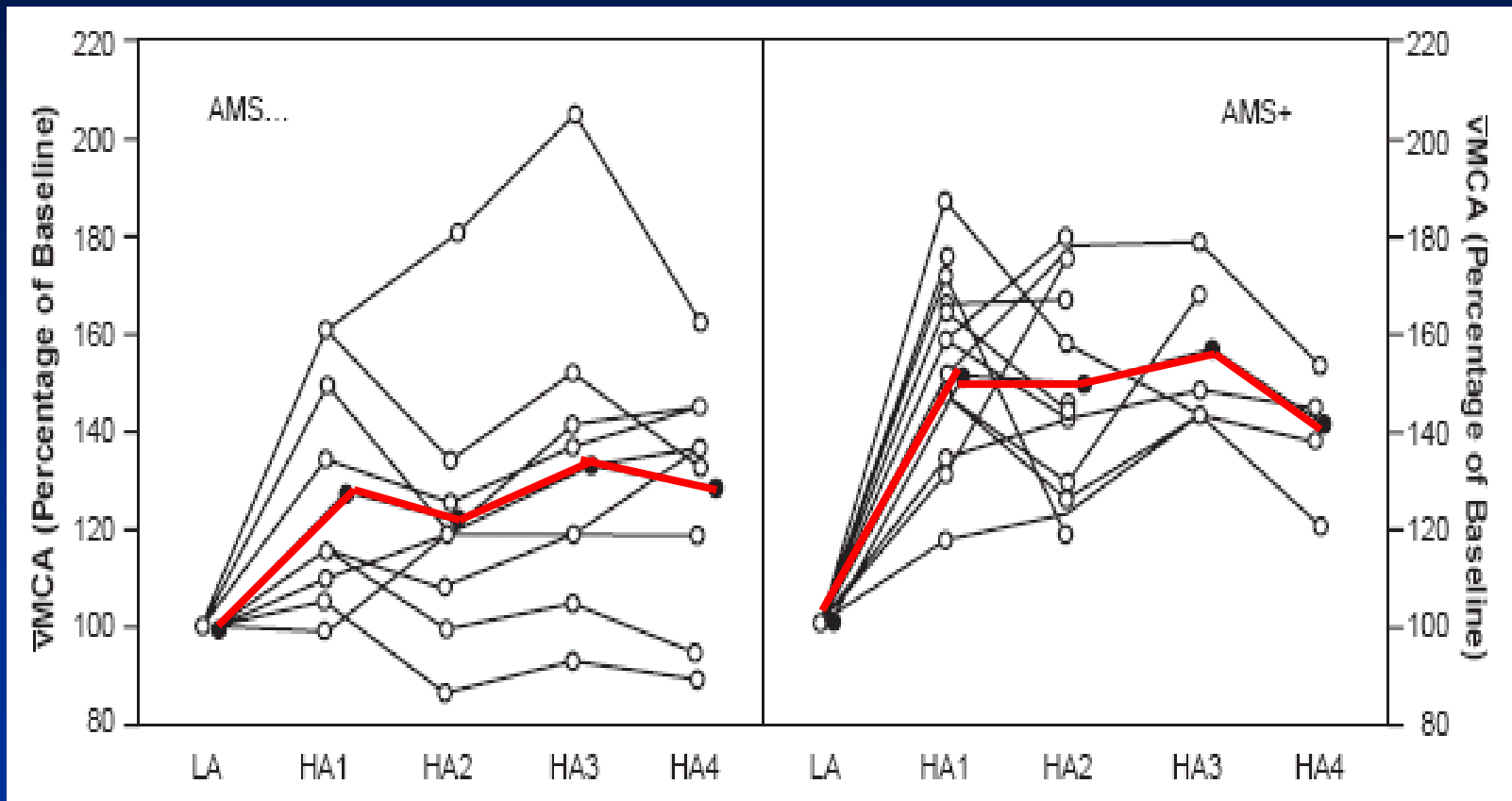


Bärtsch, P., M. Maggiorini, W. Schobersberger, S. Shaw, W. Rascher, J. Girard, P. Weidmann, and O. Oelz 1991a Enhanced exercise-induced rise of aldosterone and vasopressin preceding mountain sickness. *J. Appl. Physiol.* 71:136–143.



Bärtsch, P., M. Maggiorini, W. Schobersberger, S. Shaw, W. Rascher, J. Girard, P. Weidmann, and O. Oelz 1991a Enhanced exercise-induced rise of aldosterone and vasopressin preceding mountain sickness. J. Appl. Physiol. 71:136–143.

HA = 4.559 meters



Baumgartner RW, Bärtsch P, Maggiorini M, Waber U, Oelz O. Enhanced cerebral blood flow in acute mountain sickness. *Aviat Space Environ Med.* 1994;65:726–729.

Signa 1.5T FOR.IC0

Ex:7257
Se:2/2
Im:24/38
Ax S26.7

M 33
07/01/9
15:2

ET:8

R
1
0
0

FSE/V
TR:4000
TE:126/Ef
EC:2/2 16kHz

Head
FOV:20cm
5.0thk/2.5sp
38/04:16
256x256/1 NEX
St:I
556 L = 375



Acute Mountain Sickness

Physiological Profile

	NORMAL	AMS
V_E	++	+
P_aO_2	-	--
P_aCO_2	--	-
pH	++	+
Aldo -Exercise	+	++
ADH	+	++
ANP	+	++
Diffusion Capacity	++	+ / -
CBF	+	++
ARI	++	+

Acute Mountain Sickness Prevention

- Acclimatization at intermediate altitude is the most effective method:
 - For ascents between 8,000 and 14,000 ft, spend 2 to 4 days at 5,000 to 7000 ft
 - For ascents > 14,000 ft, spend an additional 2 to 4 days at 10,000 to 12,000 ft



Acute Mountain Sickness Prevention

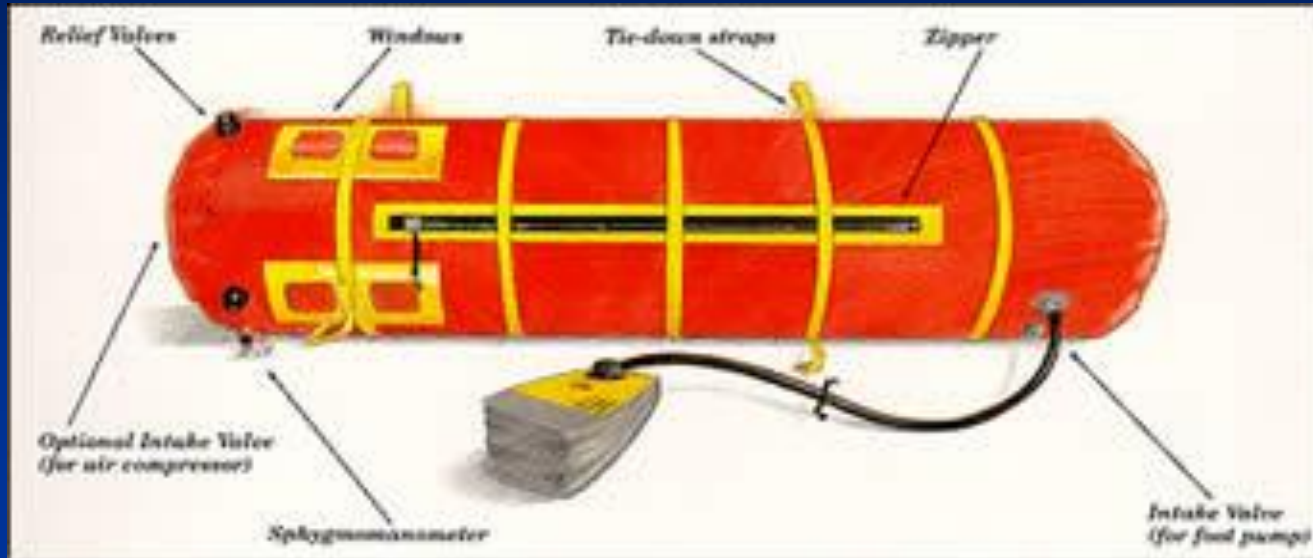
- Chemoprophylaxis
 - Acetazolamide
 - 500 mg of long acting preparation QD
 - Dexamethasone
 - 4 mg, Q6H
 - Acetazolamide plus dexamethasone
 - 500 mg of AZ QD + 4 mg DEX Q12H
 - 500 mg of AZ QD + 2 mg of DEX Q6H



Acute Mountain Sickness Treatment

- Descent to lower altitude !
- Continuous O₂ administration
- Dexamethasone
 - 4 mg Q6H
- Gamow bag







High Altitude Cerebral Edema

Symptoms

- Extreme form of AMS
- Severe headache
- Ataxia - early warning sign
- Confusion and impaired thinking
- Extreme lethargy
- Paresthesias
- Seizures
- Stupor and coma



High Altitude Cerebral Edema Treatment

- Descent to lower altitude ASAP !
- Continuous O₂ administration
- Dexamethasone
 - 8 mg STAT then 4 mg Q6H (PO or IV)
- If conscious - Gamow bag
- If unconscious or stuporous -
Intubation and hyperventilation



High Altitude Pulmonary Edema



High Altitude Pulmonary Edema

Symptoms

- Dyspnea
- Chest heaviness
- Non-productive cough
- Watery and frothy sputum
- Hemoptysis (in 20% of cases)
- Obtundation

Precedes death by 6 to 12 hours



High Altitude Pulmonary Edema

Clinical Signs

- Hyperpnea
- Rales
- Tachycardia
- Cyanosis
- Hypotension





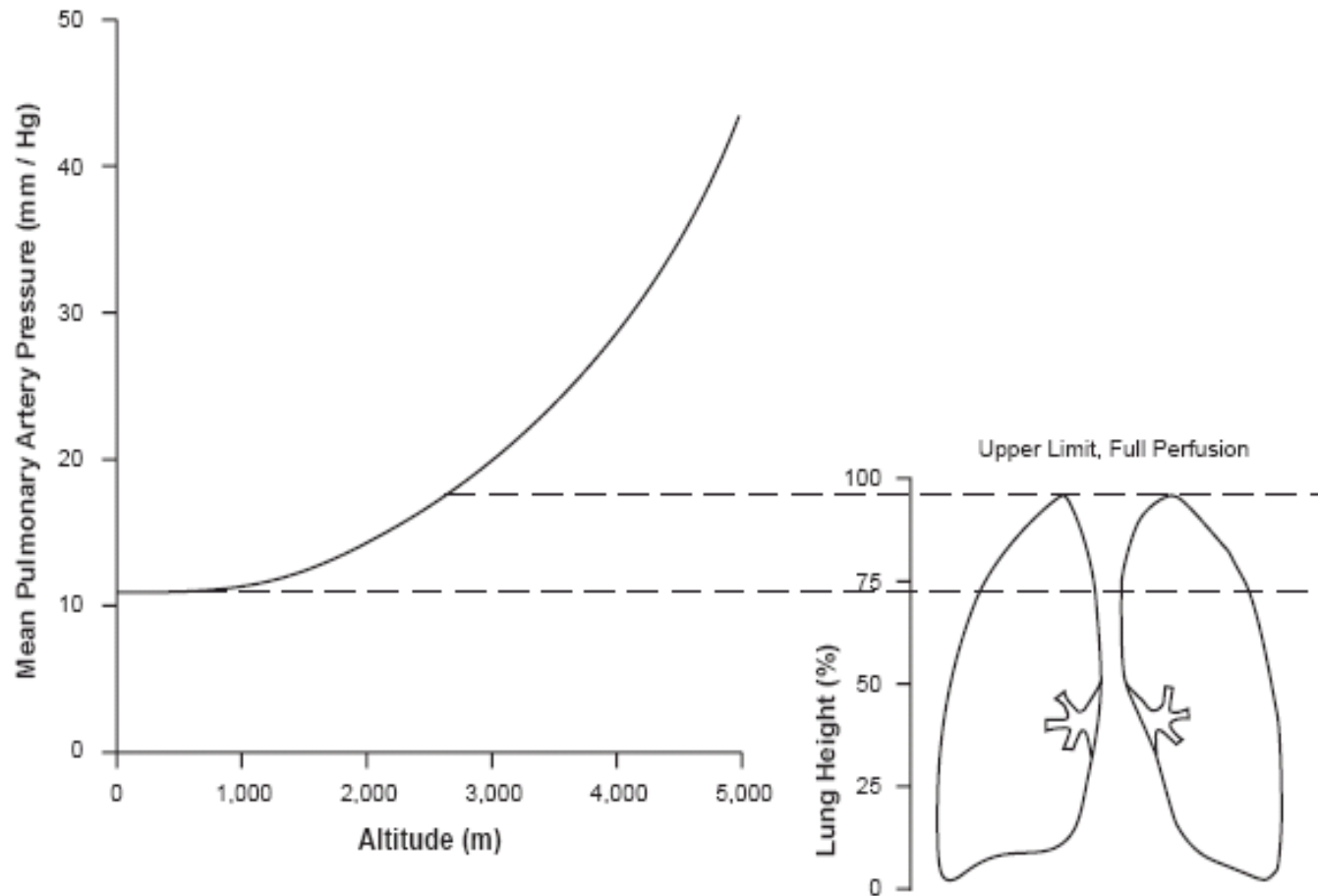
High Altitude Pulmonary Edema Epidemiology

- Typically occurs at altitudes > 10,000 ft
- Affects 5% of individuals who make an acute ascent to high altitude
- 12% have had previous episodes
- Typically occurs 3 to 5 days after acute ascent
- Exercise and cold are contributing factors
- Mortality rate is 27%

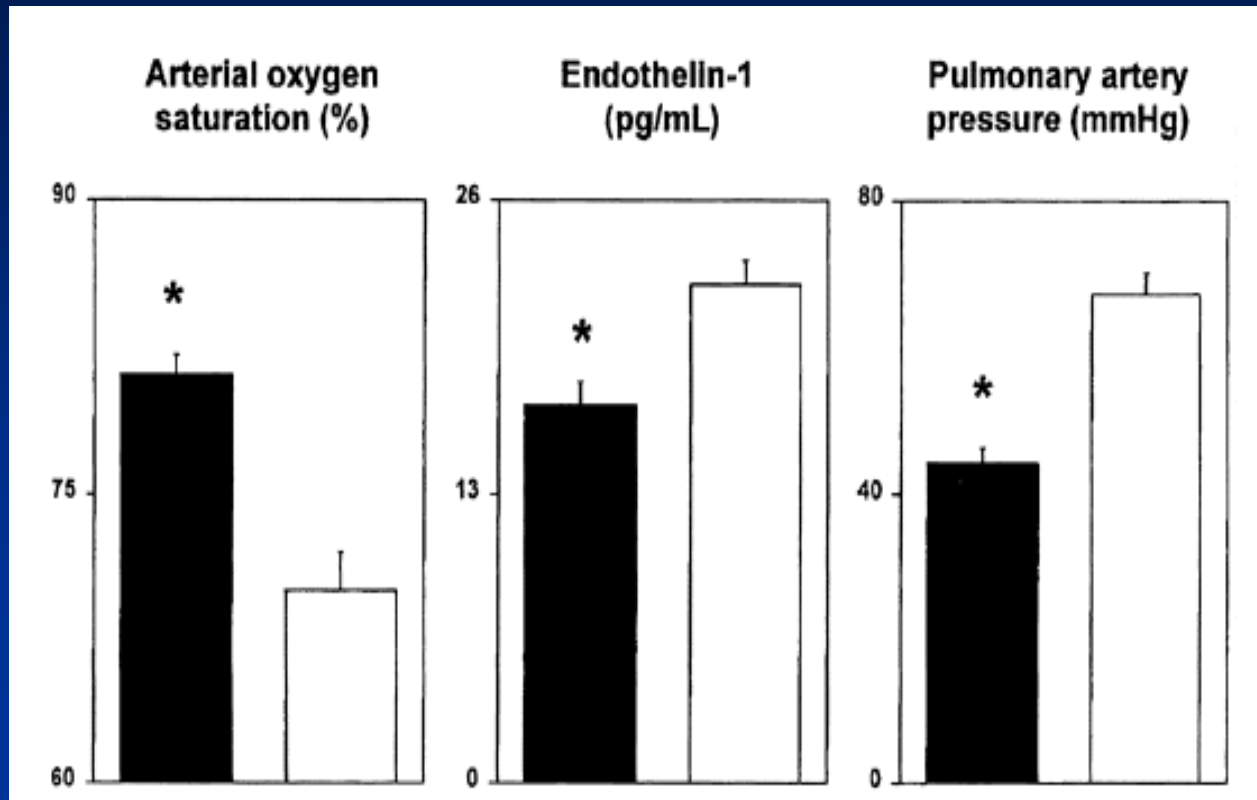




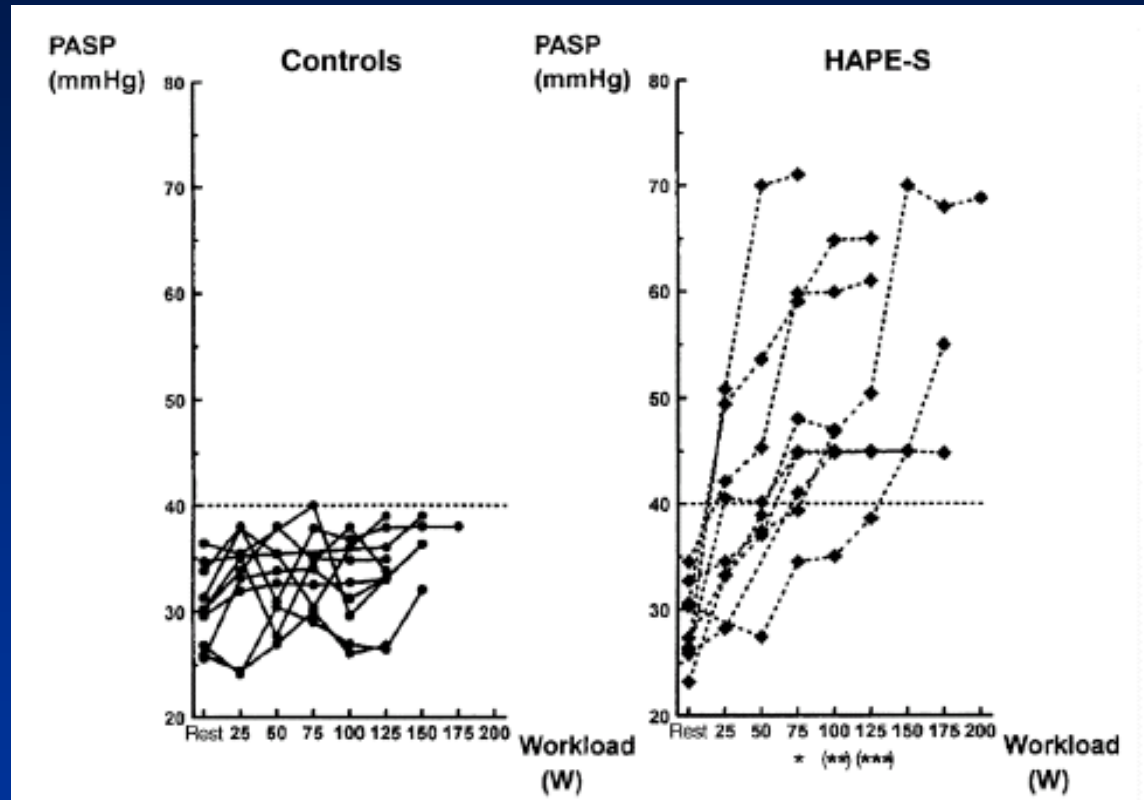
Jerusalem, Aug. 23, 2001. Amram J. "Ami" Cohen, a prominent heart surgeon who built a program that brought hundreds of children from developing countries to Israel for live saving cardiac operations died on August 16 while climbing Mount Kilimanjaro in Tanzania. He was 47.



Lenfant C, Sullivan K. Adaptation to high altitude. *N Engl J Med*. 1971;284:1303.



Sartori C., Vollenweider L., Loffler B.-M., Delabays A., Nicod P., Bartsch P., and Scherrer U. (1999b). Exaggerated endothelin release in high-altitude pulmonary edema. *Circulation* 99:2665–2668.



Grunig E., Mereles D., Hildebrandt W., Swenson E.R., Kubler W., Kuecherer H., and Bartsch P. (2000). Stress Doppler echocardiography for identification of susceptibility to high altitude pulmonary edema. *J. Am. Coll. Cardiol.* 35:980–987.

High Altitude Pulmonary Edema

Physiological Profile

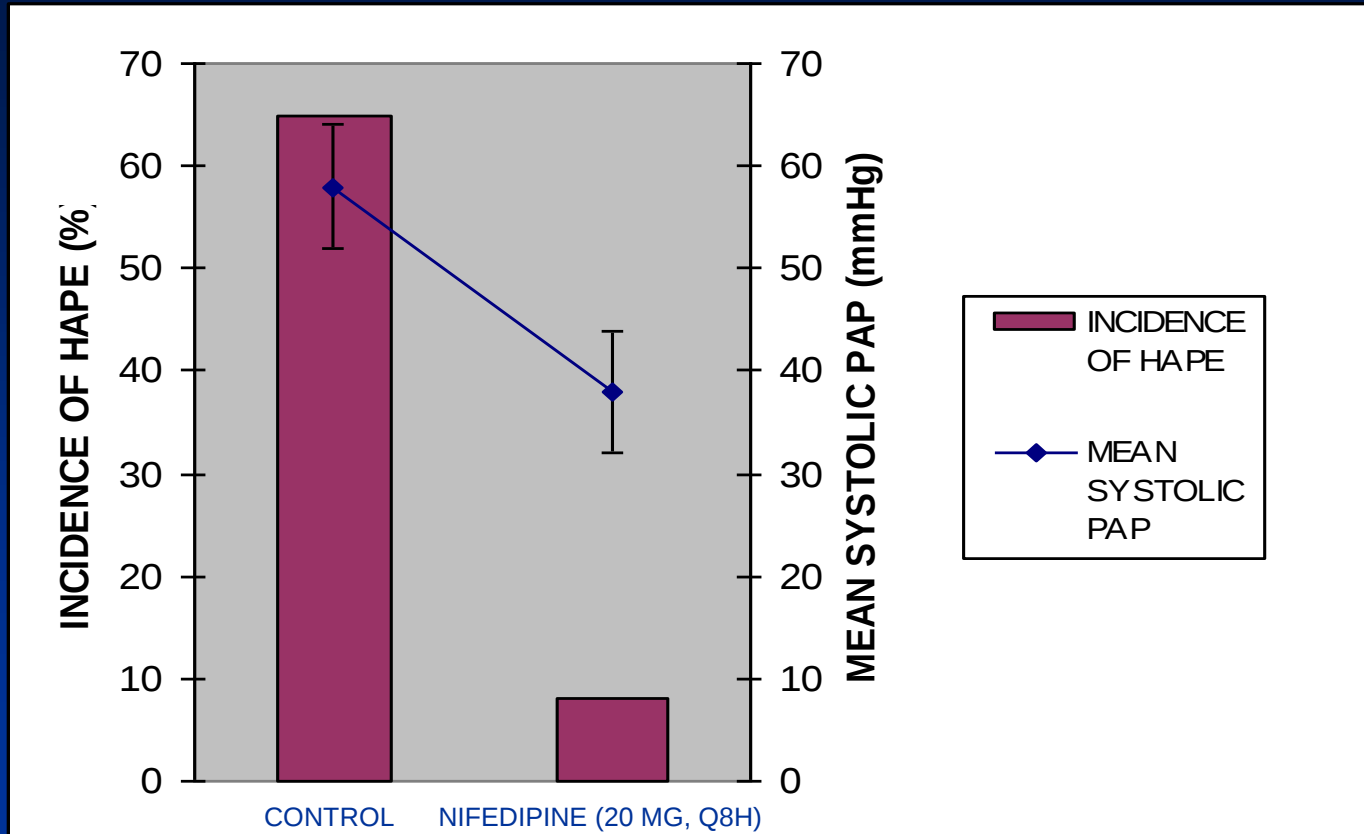
	NORMAL	HAPE
PA Blood Flow	+	+ +
PA Pressure	+	+ +
PC Pressure	+	+ +
Endothelial Breaks	-	+
Exhaled NO	+ +	+
Adrenergic Stimulation	+	+ +
V _E	+ +	+



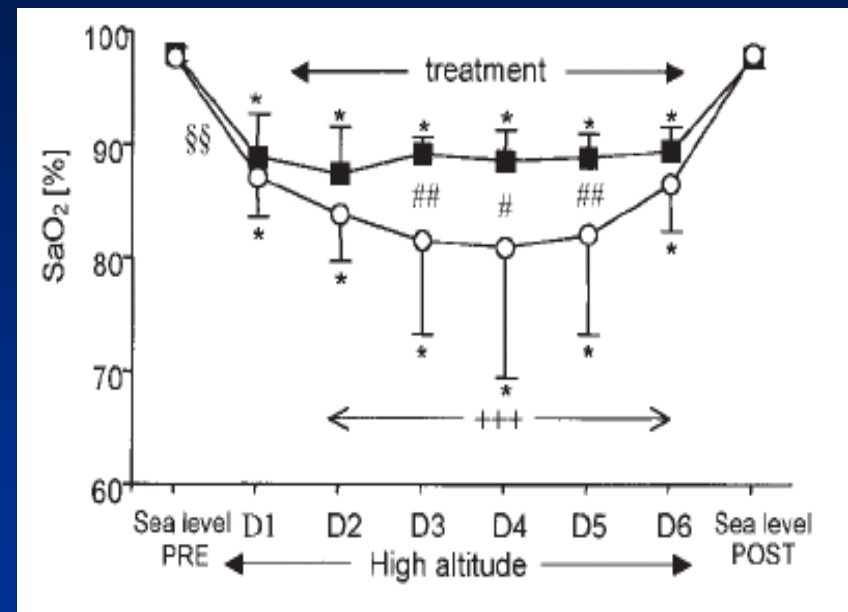
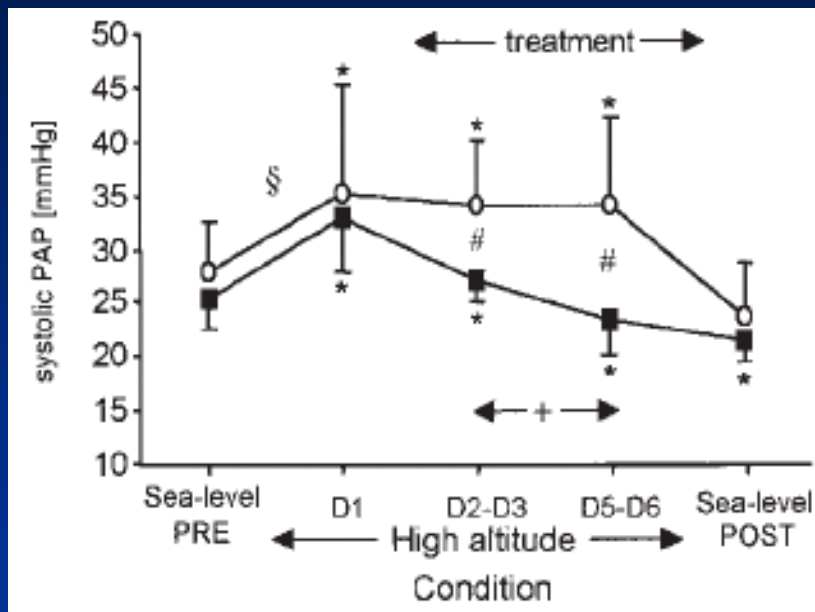
High Altitude Pulmonary Edema Prevention

- Acclimatize at intermediate altitudes
- Ascend using supplemental oxygen
- Stay warm
- Chemoprophylaxis





Bartsch P, Maggiorini M, Ritter M, et al. Prevention of high-altitude pulmonary edema by nifedipine. *New Engl J Med* 1991; 325:1284-1289



■ Sildenafil 40 mg Q8H ○ Placebo

Richalet JP, Gratadour P, Robach P, et al. Sildenafil inhibits altitude-induced hypoxemia and pulmonary hypertension. *Am J Respir Crit Care Med* 2005; 171: 275–281.

High Altitude Pulmonary Edema Prevention

- Nifedipine - 20 mg, Q8H, for 5 days
- Sildenafil - 40 mg, Q8H, for 5 days



High Altitude Pulmonary Edema Treatment

- Descend to lower altitude !
- Oxygen
- Gamow bag
- Nifedipine - 10 mg SL + 20 mg Q6H



High Altitude Pulmonary Edema Treatment

- Acetazolamide is not effective
- Furosemide is not effective
- Morphine is not effective



Sea-Level Problems at High Altitude



COPD

Predictors of Poor Performance at Altitude

- Dyspnea on exertion
- $P_aO_2 < 70$ mm Hg at or near sea-level
- $P_aO_2 < 50$ mmHg at altitude
- $MVV < 40$ L/min
- CO_2 retention



COPD

Recommendations

- If any one of the predictors of poor performance at altitude are present:
 - Do not ascend to high altitude (> 8,000 feet)
 - Supplemental O₂ for ascent to moderate altitudes (5,000 to 8,000 feet)
 - 2 L/min if not on baseline supplemental O₂
 - If on baseline supplemental O₂ - increase flow rate by 33%



COPD

Prediction of P_aO_2 at Altitude

$$P_aO_2 \text{ Alt} = 0.453 \times P_aO_2 \text{ Ground (mm Hg)} \\ + 0.386 \times (\text{FEV1 \% predicted}) + 2.44$$

P_aO_2 Alt is the predicted in-flight oxygen tension at aircraft cabin altitude in mm Hg

P_aO_2 Ground is the baseline arterial oxygen tension measured at ground altitude in mm Hg

FEV1 (% predicted) is the percentage of the predicted forced expiratory volume of air in 1 s measured during spirometric testing at ground altitude.

Dillard TA, Berg BW, Rajagopal KR, et al. Hypoxemia during air travel in patients with chronic obstructive pulmonary disease. Ann Intern Med 1989; 111:362-367



Restrictive and Interstitial Lung Disease

Predictors of Poor Performance at Altitude

- Dyspnea on exertion
- O_2 desaturation with mild exercise
- $P_aO_2 < 70$ mm Hg at or near sea-level
- $P_aO_2 < 50$ mmHg at altitude
- Moderate to severe impairment of D_LCO



Restrictive and Interstitial Lung Disease Recommendations

- If any one of the predictors of poor performance at altitude are present:
 - Do not ascend to high altitude (> 8,000 feet)
 - Supplemental O₂ for ascent to moderate altitudes (5,000 to 8,000 feet)
 - 2 L/min if not on baseline supplemental O₂
 - If on baseline supplemental O₂ - increase flow rate by 33%



Restrictive and Interstitial Lung Disease

Prediction of P_aO_2 at 8,000 feet

$$P_aO_2 (8,000 \text{ feet}) = 0.74 + (0.39 \times [P_aO_2 \text{ sea level}]) + (0.33 \times [TL_{co} \% \text{ predicted}])$$

P_aO_2 (8,000 feet) is the estimated arterial oxygen tension at 8,000 feet in kPa;

$[P_aO_2 \text{ sea level}]$ is the baseline arterial oxygen tension at sea level in kPa

$[TL_{co}\% \text{ predicted}]$ is the percentage of the predicted diffusion capacity for carbon monoxide at sea level.

Christensen CC, Ryg MS, Refvem OK, et al. Effect of hypobaric hypoxia on blood gasses in patients with restrictive lung disease. Eur Respir J 2002; 20:300-305



Altitude Simulation Test

- Hypobaric chamber
 - Can simulate a variety of altitudes and duration of hypoxia, depending upon capabilities of the chamber
 - Measure arterial blood gasses before and at the end of chamber study
 - Most commonly used in research
- Hypoxic gas mixture
 - 15% O₂ for 20 minutes simulates an altitude of 8,000 feet
 - Measure arterial blood gasses before and after exposure to hypoxic gas mixture



Pulmonary Hypertension

- Patients with pulmonary hypertension (either primary or secondary) are at risk for complications at high altitude
- Even a mild increase in hypoxia at altitude may precipitate a significant increase in pulmonary artery pressure
- Considerable variability in magnitude of hypoxia induced PA vasoconstriction



Pulmonary Hypertension

- Any degree of pulmonary hypertension is a contraindication to ascent to high altitude (i.e., altitude greater than 8,000 feet)
- Patients with Eisenmenger syndrome or severe pulmonary hypertension (Class III and IV) should not ascend above 5,000 feet - even with supplemental oxygen.
- Patients with mild to moderate (Class I and II) pulmonary hypertension may ascend to moderate altitude (5,000 to 8,000 feet) with supplemental oxygen at 2-4 L/min.

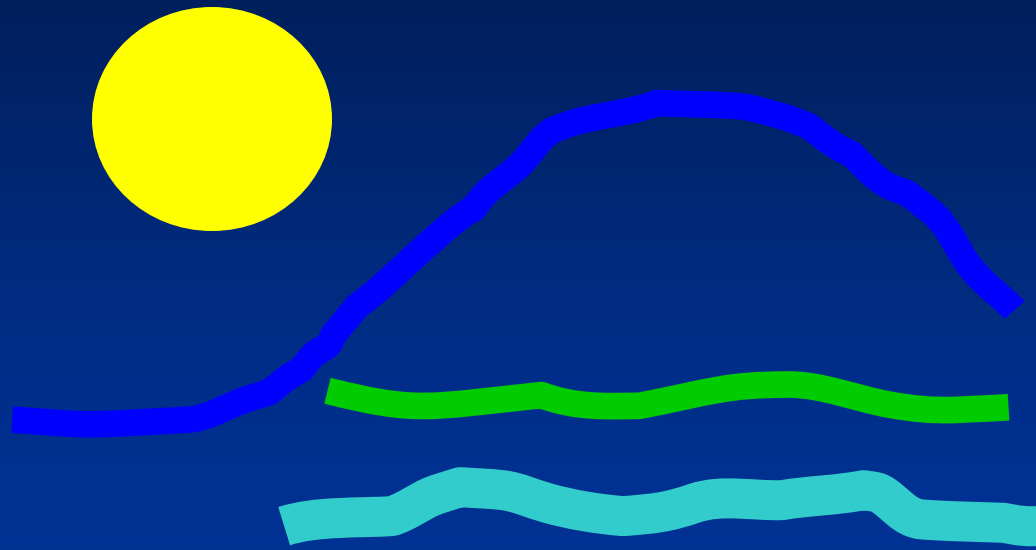


Pneumothorax

- Any evidence of pneumothorax
 - *Absolute contraindication* for ascent to altitudes > 5,000 ft
- After successful drainage
 - Wait 2-3 weeks before ascent to > 5,000 ft
 - F/U CXR recommended before ascent to > 5,000 ft

Pneumothorax

- Patients with severe bullous emphysema who have had a previous pneumothorax from a ruptured bulla:
 - Risk of a recurrent pneumothorax is significantly greater than individuals without bullous lung disease
 - Prudent to wait at least one year after the resolution of a previous pneumothorax before considering ascent to altitude > 5,000 ft



Thank You !

