

# Hyperbaric Oxygen Therapy (HBOT): A Novel Treatment for Chronic Pain?

Raymond M. Quock, Ph.D. Honors College Distinguished Professor Psychology Department Washington State University Pullman, WA



#### Лечение комплексного регионарного болевого синдрома

Н.В. ТУТЕР, А.Б. ДАНИЛОВ, Л.В. ПОЛЯКОВА

Treatment of complex regional pain syndrome

N.V.TUTER, A.B.DANILOV, L.V.POLYAKOVA

Кафедра нервных болезней факультета послевузовского профессионального образования, межклиническое отделение гипербарической оксигенации Московской медицинской академии им. И.М. Сеченова

35 patients with complex regional pain syndrome (CRPS) were treated by hyperbaric oxygenation (HBO) and caffetin preparation. A significant decrease of pain was observed in all the patients after the treatment course. Intensity of pain was diminished according to visual analogous scale. Meanwhile considerable regression of autonomic disorders and weakening of anxious and depressive manifestations was noted too. The tendency to normalization of evoked skin potentials was also found. Some elevation of the threshold of nociceptive reflex was conditioned by displacement toward general increase of antinociception after the treatment. Effect of HBO therapy persisted during 6 months in 87% of the patients. Efficiency of caffetin was restricted by the time of its administration. The conclusion was made about possibility of successful treatment of CRPS patients by both methods.



The Journal of International Medical Research 2004; 32: 258 – 262

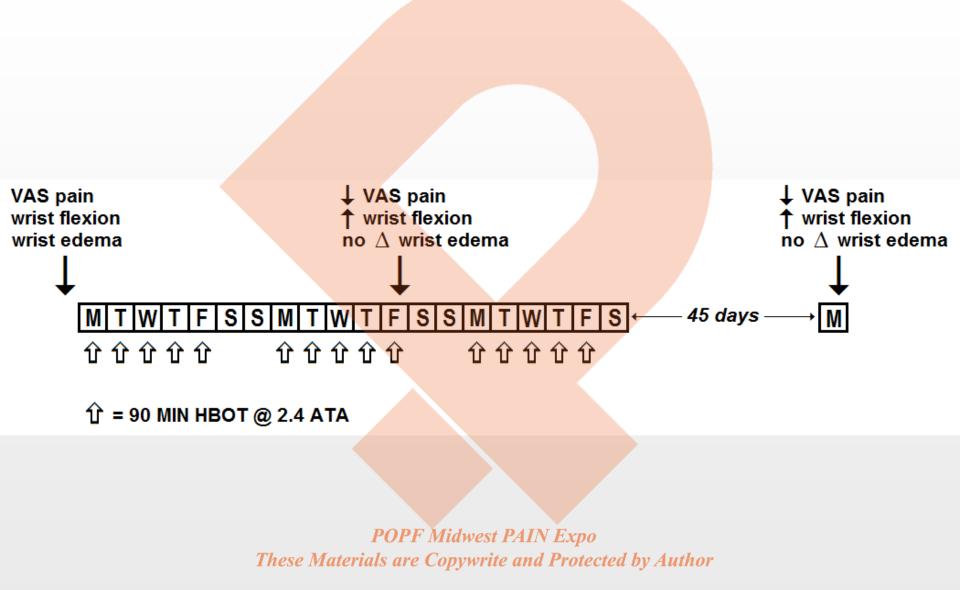
### Effectiveness of Hyperbaric Oxygen Therapy in the Treatment of Complex Regional Pain Syndrome

MZ KIRALP<sup>1</sup>, Ş YILDIZ<sup>2</sup>, D VURAL<sup>1</sup>, I KESKIN<sup>1</sup>, H AY<sup>2</sup> AND H DURSUN<sup>1</sup>

<sup>1</sup>Department of Physical Therapy and Rehabilitation and <sup>2</sup>Department of Underwater and Hyperbaric Medicine, Gülhane Military Medical Academy, Haydarpaşa Training Hospital, Istanbul, Turkey

In this double-blind, randomized, placebocontrolled study we aimed to assess the effectiveness of hyperbaric oxygen (HBO) therapy for treating patients with complex regional pain syndrome (CRPS). Of the 71 patients, 37 were allocated to the HBO group and 34 to the control (normal air) group. Both groups received 15 therapy sessions in a hyperbaric chamber. Pain, oedema and range of motion (ROM) of the wrist were evaluated before treatment, after the 15th treatment session and on day 45. In the HBO group there was a significant decrease in pain and oedema and a significant increase in the ROM of the wrist. When we compared the two groups, the HBO group had significantly better results with the exception of wrist extension. In conclusion, HBO is an effective and well-tolerated method for decreasing pain and oedema and increasing the ROM in patients with CRPS.







### Williams et al., Pain Physician 12:E335-339, 2009

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Case Report

#### Chronic Regional Pain Syndrome After Subtalar Arthrodesis Is Not Prevented by Early Hyperbaric Oxygen

#### Simon TB Williams, MBBCh<sup>1</sup>, Anthony Davies, FRCA<sup>2</sup>, and Phil Bryson, MBBS<sup>1</sup>

From: 'Diving Diseases Research Centre, Plymouth, UK; and 'Derriford Hospital, Plymouth, UK.

Dr. Williams and Dr. Bryson are with the Diving Diseases Research Center, Tamar Science Park, Plymouth, UK. Dr. Davies is a Pain Management Specialist with the Department of Anaesthesia, Derriford Hospital, Plymouth, UK.

> Address correspondence: Simon TB Williams, MBBCh Diving Diseases Research Centre Tamar Science Park Research Way Plymouth, Devon PL6 8BU United Kingdom E-mail: stbwilliams@hotmail.com

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> Free full manuscript: www.painphysicianjournal.com

Subtalar arthrodesis was performed on a 48-year-old, non-insulin-dependent diabetic with a history of chronic ankle instability and lateral ankle pain. In the early post-operative period he presented as an emergency with an infection at the operative site. This was treated with 2 returns to the operating theatre for washout and debridement. His wounds were left open and at 3 weeks after emergency admission he was referred for adjunctive hyperbaric oxygen (HBO) therapy to aid healing by secondary intention. He received a total of 19 hyperbaric sessions, at a pressure of 2.2 ATA, one treatment per day for 5 days a week.

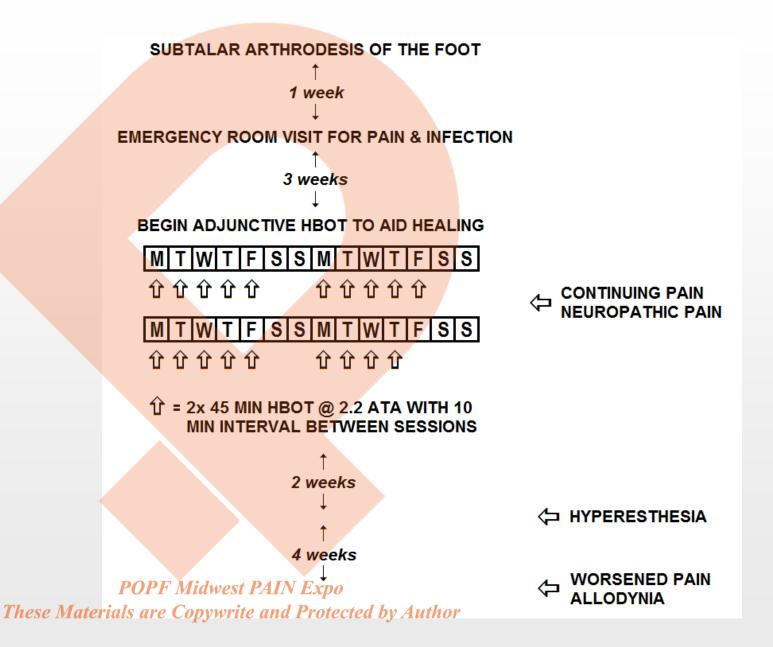
Shortly after commencing HBQ therapy his ankle became increasingly painful, despite the introduction of analgesia. By 7 weeks after emergency admission his wounds had virtually healed but hyperesthesia persisted over the dorsum of the foot. A computerized tomography scan at 5½ months post-operatively showed satisfactory joint fusion and revealed no evidence of infection. Symptoms and signs at this time were compatible with a diagnosis of chronic regional pain syndrome (CRPS).

There is published evidence to suggest that HBO therapy may be a useful modality in the treatment of established CRP5. Here, we seek to publicize a case in which early treatment with HBO for another indication did not prevent the simultaneous development of CRP5 Type 1.

Key words: Subtalar arthrodesis, hyperbaric oxygenation, chronic regional pain syndrome

Pain Physician 2009; 12:E335-E339







### Outline

### HBOT and Complex Regional Pain Syndrome (CRPS)

• Three reports from the clinical literature

HBOT

- What is it?
- FDA-approved HBOT indications
- Main effects of HBOT
- Off-label uses of HBOT

### HBOT Basic Research Results

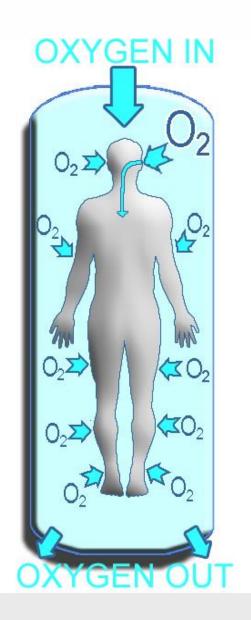
- Mechanism of pain relief may originate in the brain
- Mechanism of pain relief may involve the opioid system



### Hyperbaric Oxygen Therapy (HBOT)

HBOT is the clinical application of 100% oxygen at atmospheric pressures higher than sea level for limited periods of time (60– 90 min) to achieve therapeutic outcomes.

HBOT requires the use of a hard shell pressure vessel (hyperbaric chamber).





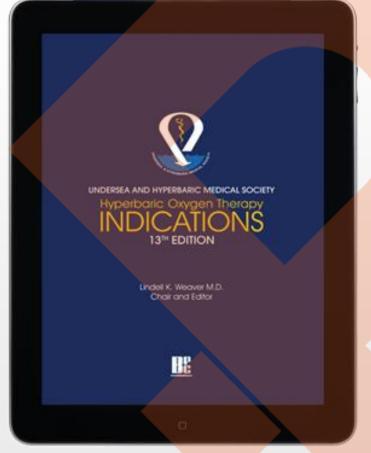
### **Hyperbaric Chambers**



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# **FDA-Approved HBOT Indications**



- Air or gas embolism
- Carbon monoxide poisoning
- Gas gangrene
- Acute traumatic ischemia
- Decompression sickness
- Arterial insufficiencies
- Severe anemia
- Intracranial abscess
- Necrotizing soft tissue infections
- Refractory osteomyelitis
- Delayed radiation injury
- Compromised skin grafts and flaps
- Acute thermal burn injury
- Idiopathic sudden sensorineural
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   hearing loss
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### **Mechanical Effect of Increased Pressure**

- Any free gas trapped in the body will decrease in volume as pressure increases (Boyle's Law).
- Reduction in bubble size may allow it to pass through the circulation.
- This effect is useful in the management of gas embolism and decompression sickness

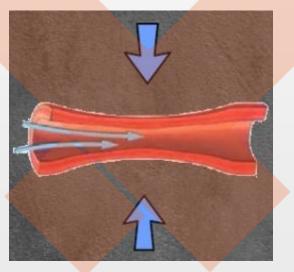
### Mass Action of Gases

- Flooding the body with oxygen forces the rapid elimination of other toxic gases such as carbon monoxide.
- The mechanic effect of increased pressure further accelerates the elimination process.



### Vasoconstriction

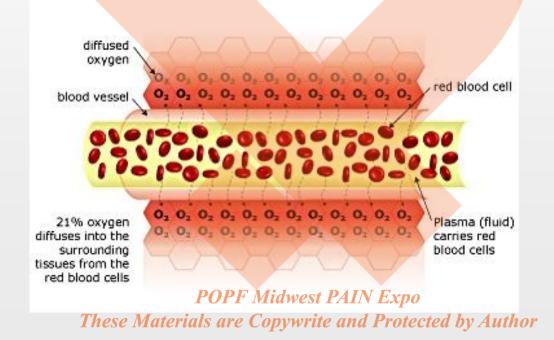
- Vasoconstriction can result in reduction of edema.
- Even with reduced blood flow, enough extra oxygen is carried by the blood so a net increase in tissue oxygen delivery occurs with HBOT.
- This effect is therapeutic in crush injury/compartment syndrome and thermal burns.





### Hyperoxygenation

- HBOT physically dissolves extra oxygen into the plasma (Henry's Law).
- This effect increases delivery of oxygen to ischemic or underperfused tissues, which can limit ischemic damage, inflammation and cell death.
- The additional oxygen helps the ischemic tissue meet the increased metabolic needs of the healing process.





### Antibacterial Effect

- Anaerobic bacteria do not flourish in the presence of oxygen.
- Many of the body's bacterial defense mechanisms are dependent on oxygen.
- HBOT slows the growth of anaerobic bacteria, allow the body's bacterial defense mechanisms to be reactivated, and combat clinical infections such as gas gangrene.



There are clinical reports of HBO<sub>2</sub>-induced therapeutic effects in the following conditions:

- Traumatic brain injury (TBI)
- Persistent post-concussion syndrome
- Cerebral palsy
- Multiple sclerosis
- Chronic fatigue syndrome
- Stroke
- Autism
- Cancer



## **HBOT-Induced** Pain Relief

There are clinical reports of HBO<sub>2</sub>-induced relief of pain in the following conditions:

- Complex regional pain syndrome (reflex sympathetic dystrophy syndrome)
- Fibromyalgia
- Trigeminal neuralgia
- Migraine and cluster headache
- Rheumatoid arthritis
- Chronic osteomyelitis



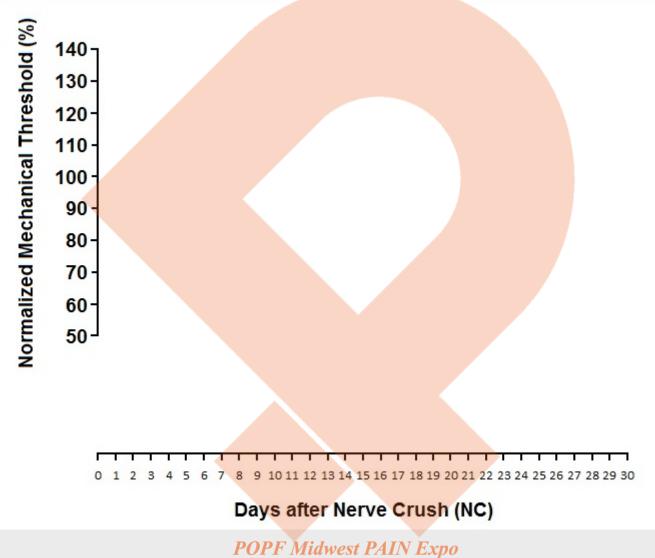
BRAIN RESEARCH 1537 (2013) 111-116

#### Brain Research Available online at www.sciencedirect.com ScienceDirect www.elsevier.com/locate/brainres **Research Report** Involvement of brain opioid receptors in the CrossMark anti-allodynic effect of hyperbaric oxygen in rats with sciatic nerve crush-induced neuropathic pain Carlee R. Gibbons<sup>a,1</sup>, Shulin Liu<sup>a,c,1</sup>, Yangmiao Zhang<sup>a</sup>, Casey L. Sayre<sup>d</sup>, Briana R. Levitch<sup>a</sup>, Sarah B. Moehlmann<sup>a</sup>, Donald Y. Shirachi<sup>e</sup>, Raymond M. Quock<sup>a,b,\*</sup> \*Department of Pharmaceutical Sciences, College of Pharmacy, Washington State University, Pullman, WA, USA <sup>b</sup>Translational Addiction Research Center, Washington State University, Pullman, WA, USA Department of Diving Medicine, Second Military Medical University, Shanghai, China <sup>d</sup>Faculty of Pharmacy, University of Manitoba, Winnipeg, MB, Canada \*Department of Physiology and Pharmacology, Thomas J. Long School of Pharmacy and Health Sciences, University of the Pacific, Stockton, CA, USA ABSTRACT ARTICLE INFO Article history: Earlier research has demonstrated that hyperbaric oxygen (HEO<sub>2</sub>) can produce an antinoci-Accepted 25 August 2013 Available online 30 August 2013

Keywords: Hyperbaric oxygen Anti-allodynia Naltrexone Opioid receptors Neuropathic pain Sciatic nerve crush Rat ceptive effect in models of acute pain. Recent studies have revealed that HBO2 can produce pain relief in animal models of chronic pain as well. The purpose of the present investigation was to ascertain whether HBO2 treatment might suppress allodynia in rats with neuropathic pain and whether this effect might be blocked by the opioid antagonist naltrexone (NTX). Male Sprague Dawley rats were subjected to a sciatic nerve crush under anesthesia and mechanical thresholds were assessed using an electronic von Frey anesthesiometer. The time course of the HBO2-induced anti-allodynic effect in different treatment groups was plotted, and the area-under-the-curve (AUC) was determined for each group. Seven days after the nerve crush procedure, rats were treated with HBO2 at 3.5 atm absolute (ATA) for 60 min and exhibited an anti-allodynic effect, compared to nerve crush-only control rats. Twenty-four hours before HBO2 treatment, another group of rats was implanted with Alzet® osmotic minipumps that continuously released NTX into the lateral cerebral ventricle for 7 days. These NTX-infused, HBO2-treated rats exhibited an allodynic response comparable to that exhibited by rats receiving nerve crush only. Analysis of the AUC data showed that HBO2 significantly reduced the nerve crush-induced allodynia; this anti-allodynic effect of HBO, was reversed by NTX. These results implicate opioid receptors in the pain relief induced by HBO2.

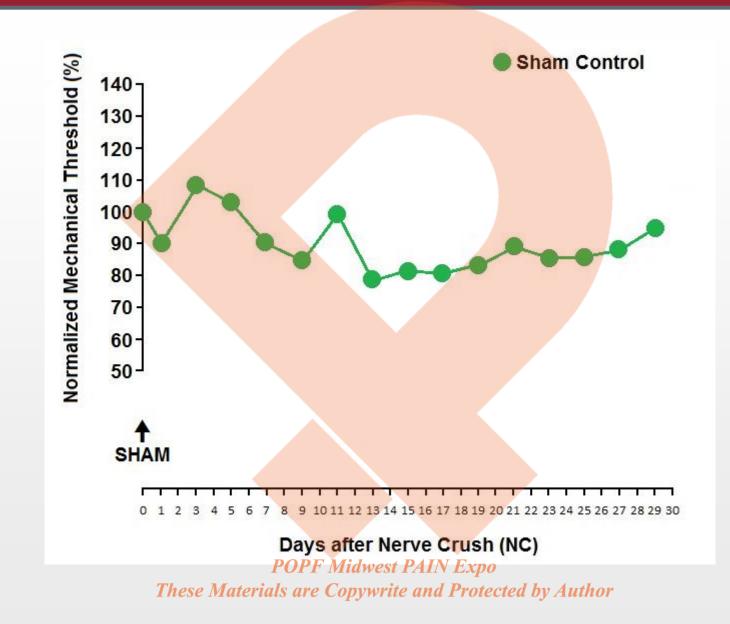
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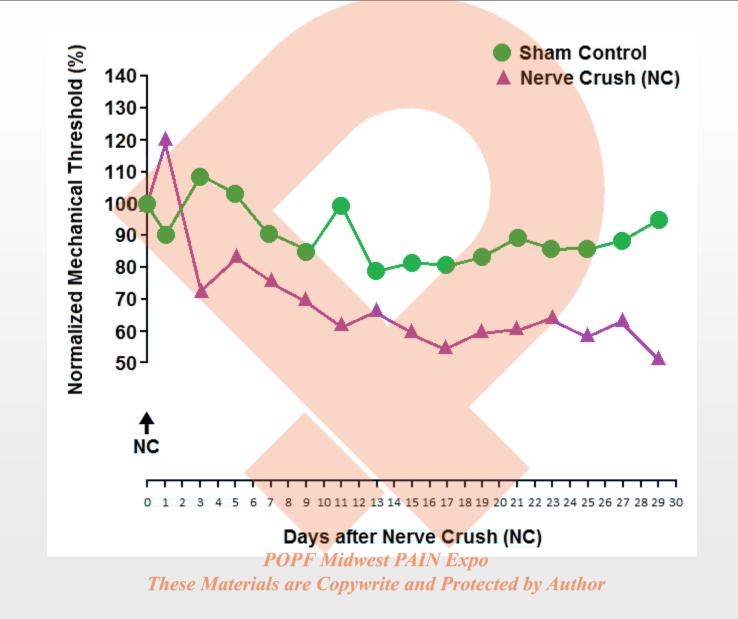


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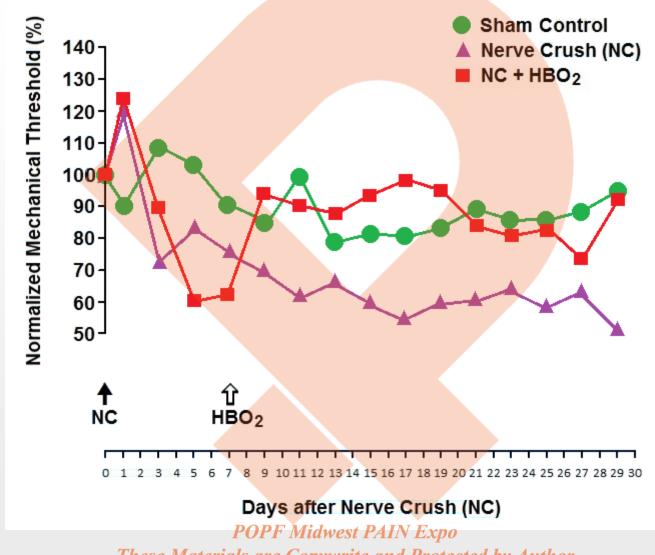






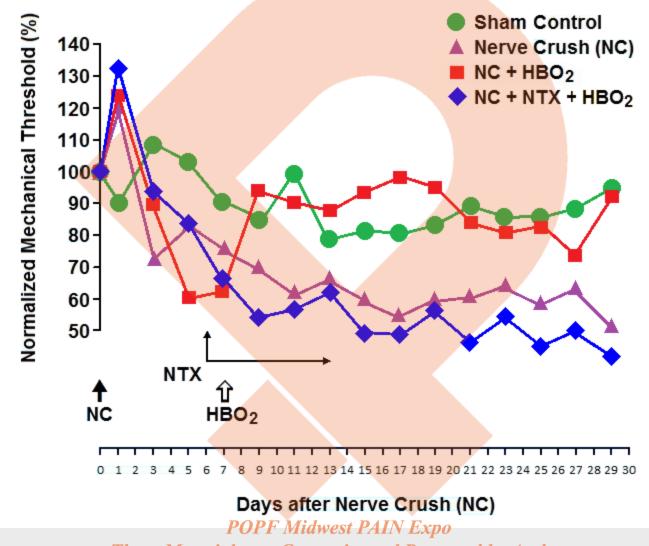






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## Summary

- 1. HBOT is a medical treatment that delivers 100% oxygen under higher-than-normal atmospheric pressure.
- 2. There are fourteen clinical indications for HBOT in the U.S. that are approved by the AMA and the FDA.
- 3. There are several "off-label" indications that have been shown to be responsive to HBOT, including several conditions of chronic pain.
- 4. There are clinical reports that HBOT can produce pain relief in complex regional pain syndrome.
- 5. Basic science research indicates that HBOT-induced pain relief may originate from an action in the brain and possibly involve the endogenous opioid system.



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