

Measuring tissue oxygen tension: a review

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Sheffield PJ. Measuring tissue oxygen tension: a review. Undersea Hyper Med 1998; 25(3):179–188.—Because of technological advances in tissue oximetry, clinicians and scientists have a better understanding of the role of oxygen in wound healing. In wound care and hyperbaric medicine applications, an oximeter is principally used with vascular assessment to help determine amputation level and to estimate healing potential. With the current emphasis on cost savings in the managed care setting, transcutaneous oximetry (PtcO₂) has gained importance as a tool for predicting potential candidates for hyperbaric oxygen (HBO₂) therapy. It is used to identify the presence of hypoxia in wounded tissue, to predict the responders to hyperoxia and in some instances to determine when HBO₂ treatment is complete. This literature review describes the principal current methods for measuring tissue O₂ and the values obtained in normal and wounded tissue under both normobaric and hyperbaric conditions. The review includes the Jefferson C. Davis Wound Care and Hyperbaric Medicine Center protocol for PtcO₂ assessment of potential HBO₂ candidates and suggestions for obtaining reproducible PtcO₂ data.

tissue oxygen measurement, tissue oxygen tension, oxygen electrodes, normal oxygen tension, wound oxygen tension, transcutaneous oximetry, oxygen in wound healing, vascular assessment

Much has been written about the role of oxygen in wound healing and the technological developments to measure it. Hunt (1) and Niinikoski (2) demonstrated the hypoxic nature of wounds and postulated that healing and incidence of infection were related to O₂ supply. Silver (3) precisely analyzed the environment in and around experimental wounds and revealed an O2 gradient between the dead space of the wound and the wound edge. Investigations of severely hypoxic wounds in animal models suggested that elevation of O₂ tension up to normal would enhance healing, since the rate of healing is O_2 dependent (4). Direct relationships have been demonstrated between wound O2 tension and production of collagen (5), epithelialization (6), resistance to infection (7), and promotion of healing (1,7,8). Because of technological advances in O2 monitoring, clinicians and scientists now have a better understanding of wound healing mechanisms and the role of O2 in wound healing.

Sheffield (9) demonstrated that hyperbaric oxygen (HBO₂) elevated O₂ tension in severely hypoxic wounds. HBO₂ elevates arterial PO₂ (10), increases the steepness of the PO₂ gradient within tissue (3), and increases the O₂ diffusion distance from functioning capillaries (11). Thus, HBO₂ corrects severe hypoxia in some inadequately perfused wounds. In wound care and hyperbaric medicine applications, O₂ monitors are principally used with vascular assessment to help determine amputation level and estimate healing potential.

This review includes a discussion of methods used at the

Jefferson C. Davis Wound Care and Hyperbaric Medicine Center, San Antonio, Texas.

WHY MEASURE TISSUE OXYGEN TENSION?

Tissue oxygen tension is a direct, quantitative assessment of O_2 availability to tissue. Tissue O_2 data are used in medical decision making by several medical specialties, including wound care and hyperbaric medicine. With the current emphasis on cost savings in the managed care setting, tissue oximetry has gained importance as a tool for predicting potential candidates for HBO_2 therapy. These data aid in vascular assessment to help predict non-responders to treatment and to choose successful amputation sites. They are also used to select candidates for HBO_2 by identifying the presence of tissue hypoxia and the responders to hyperoxia. In some instances tissue O_2 data are used to determine when treatment is complete.

Present methods of measuring wound oxygen tension

Principal methods presently used in assessing tissue oxygenation are invasive polarographic electrodes, mass spectroscopy, tissue tonometry, blood gas analysis, and transcutaneous oximetry (12,13).

Invasive polarographic electrodes: The first application of O₂ polarography in animal tissues is credited to Davies and Brink (14) who used a 25-μm platinum bare-wire electrode to show that the O₂ value in tissue differed from that in a fluid-filled recess of the tissue. As electrode technology improved, the cathode was constructed of a

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semi-noble metal (silver, platinum, or gold) with a reference anode of silver/silver chloride. To prevent protein coating of the electrode, the cathode was usually coated with a collodion membrane that was permeable to water and oxygen. Clark and associates (15) improved the design by placing both the anode and cathode in an electrolyte solution behind a polyethylene membrane. Because of its large size, its application was limited to fluid and organ surfaces until it was miniaturized by Silver (16). The probe is calibrated before insertion into the tissue. An 18- to 22-gauge wire electrode has a sensing area of about 450 µm, 4-6 times the diameter of the tip (17). Invasive electrodes were used to confirm hypoxia in chronic, indolent human wounds and to show that oxygen is delivered to wounds during HBO₂ treatment (9).

Mass spectroscopy: The mass spectrometer can simultaneously analyze several tissue gases, such as O₂, carbon dioxide, nitrogen, and anesthetic gases. It has been used for both normobaric (18) and hyperbaric (19) applications. Two types of permeable membranes are commonly used for the tissue probe: a fast-response Silastic membrane for flowing blood, and a slow-response Teflon membrane for tissue. The inner face of the membrane is exposed to a low pressure (average, 10⁻⁶ mmHg), whereas the outside face is in contact with the tissue gas to be measured. Gas molecules from the tissue pass through the permeable membrane and are drawn via tubing into the mass spectrometer, where they are separated and measured quantitatively (20). Wells et al. (19) used mass spectroscopy to record Po2 and other tissue gas values for healthy subjects under normobaric and hyperbaric conditions (Table 1). The authors reported that following a HBO₂ exposure of 1 h, the subject's skin O₂ tension remained elevated >10% for up to 3 h.

Tissue tonometry: A method of tissue tonometry was developed by Hunt (21) in which about 14 cm of Silastic tubing was passed into the skin via an 18-gauge cutting-tip spinal needle. Physiologic saline was slowly injected into the tubing, allowing time for tissue gases to equilibrate with the saline in the tube. A 25-μm polarographic O₂ electrode was then inserted into the end of the tonometer for continuous measurement of O₂ tension. As an alternative, the saline can be withdrawn from the tonometer and measured by blood gas analysis. This method gives a single, integrated, mean value of extracellular fluid O₂ tension across the length of the tonometer. The reading is proportional to regional blood supply, arterial O₂ tension, and microvascular perfusion. The electrode can be calibrated in situ, and tissue O₂ tension can be read continuously (22).

Blood gas analysis: Procedures are now available for collecting arterial blood gases during HBO₂ treatments in

both walk-in and single-occupant chambers. The blood gas analyzer can be placed inside a walk-in chamber and the sample analyzed while under pressure (23). A portable, battery-powered blood gas analyzer (StatPal II Blood Gas Analyzer, PPG Industries) was evaluated and found to be effective at 2.5 atm abs (24), but when greater pressure is applied, the pressure-sensitive keys on the electronic keyboard are depressed, causing the analyzer to continually cycle and rendering it unusable (Martindale VE. 1994, personal communication). In a single-occupant chamber, arterial blood samples are aspirated from the subject and passed through the chamber wall for analysis outside the chamber (25). Analysis must be immediate to avoid false low readings due to off-gassing of the sample. Arterial blood gases measured at 1 atm abs have been used to predict arterial O₂ tension under HBO₂ conditions (26). Howe and associates (27) reported a correlation between arterial O2 tension and transcutaneous oxygen tension at the level of the chest, but on the leg and foot, the correlation is not clear.

Transcutaneous oximetry: Transcutaneous oximetry (PtcO₂, a.k.a. TcPO₂, TCOM) was developed as a non-invasive method to monitor tissue O₂ tension. Baumberger and Goodfriend (28) immersed a subject's finger in a phosphate buffer solution heated to 45 °C for about 60 min to demonstrate that the O₂ tension of the buffer solution approximated the subject's arterial O₂ tension. Huch et al. (29) developed PtcO₂ by modifying a Clark electrode with a heating element and thermistor to maintain a preset temperature of 42°-45°C. A fixation ring filled with contact solution is used to attach the electrode to the skin surface.

To aid O₂ migration to the skin surface where it can be analyzed, the noninvasive sensor must cause physiologic changes in the underlying tissue. Heating the sensor to 42°-45°C transfers heat to the skin surface, dilating capillaries, opening skin pores, decreasing O₂ solubility, and shifting the oxyhemoglobin curve to the right for ready release of O_2 (30). In the absence of heat, diffusion of O_2 from tissue to skin surface contributes less than 3.5 mmHg to the Po₂ at that location (31). A control electrode is placed on the skin at the second intercostal space of the chest. As of this writing, transcutaneous sensors will not adhere to a moist surface, thus PtcO2 values are collected near the wound at standardized locations. The interpreter uses periwound PtcO₂ in combination with other assessment methods (e.g., temperature, pulse, presence of hair, general appearance of the tissue) to predict the behavior of the wound.

Transcutaneous oximetry was originally used in neonatology (32), and is now commonly used in pediatric intensive

Table 1: Tissue Oxygen Tension Values for Progessively Increased Inspired Po2

Ambient Pressure (atm abs) / Breathing Media	1.0 Air	1.0 O ₂	2.0 O ₂	2.4 O ₂	3.0 O ₂		
de hij a rossantin sistema i	Representative tissue oxygen tension values, mmHg						
Ambient Po ₂ , mmHg	159	760	1520	1,824	2280		
Ideal alveolar Po ₂	104	673	1433	1,737	2193		
Ideal arterial Po ₂	100	660	1400	1,700	. 2150		
Arterial Po ₂ ^a		550±100	1,150±250	_	1,750±250		
Venous Po	36±4	60±9	101±36	W. Hilliam saled	_		
Muscle Po ₂ ^b	29±3	59±13	221±72				
Subcutaneous Po2b	37±6	53±10	221±72	- 5 -	-		
Subcutaneous Po ₂ c	65±7	_	-	-	-		
Subcutaneous Po ₂ ^d	30-50	90-150	200-300	250-500	-		
Typical chronic wound Po ₂ ^d	5-20	200-400		1,000-1,700	-		
Transcutaneous Po2e	69±6	440±95		1,350±220	71 1 1 2		
Transcutaneous Po2—chest	67±12	450±54	-	1,312±112	A 10 10 1		
Transcutaneous Po2—calf, male f	49±14	281±78	-	1,027±164			
Transcutaneous Po2—calf, female	59±12	367±59	-	1,174±127	-		
Transcutaneous Po2—midfoot	63±13	280±82	-	919±214	_		
Transcutaneous Po2—limbg	49	325	696		1		

^aBlood gas analyzer data from (10); ^bmass spectrometer data from (19); ^cimplanted tissue tonometer data from (40); ^dimplanted polarographic O₂ data from (12); ^etranscutaneous O₂ data from (41); ^dtranscutaneous O₂ data from (42); ^dtranscutaneous O₂ data from (43).

care units (33), plastic surgery (34), vascular surgery (35), anesthesiology (36). orthopedics (37), and hyperbaric medicine (38,39). In 1994, Matos and Nunez (38) reviewed a series of tissue oxygenation studies and concluded that PtcO₂ was clinically useful in determining healing potential, selecting amputation level, evaluating revascularization procedures, and assessing severity and progression of peripheral vascular disease.

MEASUREMENTS OF NORMAL O2 TENSION

No single value can be specified "normal" O₂ tension for all tissue. Rather, there exists a series of gradients, the steepness of which varies with arterial O₂ tension, type of tissue, inter-capillary distance, and cellular metabolic rate. Normal values will vary by method of measurement. Table 1 contains tissue O₂ values measured by blood gas analyzer (10), mass spectrometer (19), tissue tonometer (40), implanted polarographic electrode (12), and PtcO₂ (41–43) at pressures of 1.0, 2.0, 2.4, and 3.0 atm abs. The data of Dooley et al. (42) contain "normal" PtcO₂ values obtained in 72 subjects (53 males, 19 females) for chest, calf, and midfoot, and reveal a significant gender difference at the calf.

MEASUREMENTS OF WOUND OXYGENATION DURING A COURSE OF NORMOBARIC OR HYPERBARIC OXYGEN

Studies of human ischemic wounds revealed that many problem wounds were severely hypoxic and that HBO₂ corrected the severe hypoxia by daily bringing wound

oxygenation (PwO₂) up to normal levels (9). Specific findings were:

- Problem wounds have low PO₂: Invasive O₂ electrodes in chronic, indolent, soft-tissue human wounds revealed that non-healing wounds were often severely hypoxic, with PwO₂ values ranging from 5 to 20 mmHg as compared to control tissue of 30–50 mmHg (12). Values in the center of the wound were lower than values at the wound edge. PtcO₂ showed evidence of severe hypoxia adjacent to problem wounds, with PtcO₂ values adjacent to the wound falling below 20 mmHg, as compared to 65 ± 15 mmHg in healthy control tissue (44).
- Wounds respond to O₂: Invasive polarographic O₂ electrodes in two diabetic wounds revealed PwO₂ values less than 20 mmHg during air breathing, with subsequent increases of 10–400 mmHg during periods of 100% O₂ breathing at 1 atm abs (9).
- Wound PO₂ has minute-to-minute variability. Invasive
 O₂ electrodes revealed fluctuations in PwO₂ reflecting the
 local tissue demands for O₂ (9).
- Hyperbaric oxygen elevates wound PO₂: Invasive oxygen electrodes showed that HBO₂ demonstrably increased PwO₂ to above 1,000 mmHg in previously hypoxic wounds (9,45).
- Multiple HBO₂ exposures increase wound response: Measured invasively, baseline PwO₂ values that were below 15 mmHg while breathing air at 1 atm abs increased to above 30 mmHg as angiogenesis and vascularity improved with multiple (usually 20–30) HBO₂ exposures (9).

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• Irradiated tissue responds like a wound: Invasive electrode assessment during O2 breathing at 2.4 atm abs revealed PwO2 frequently exceeding 1,000 mmHg, but intact skin values rarely exceeded 500 mmHg, because of its ability to vasoconstrict. Invasive Po2 values in irradiated skin were 600-1,100 mmHg (like a wound) during the first 7 wk of HBO₂ treatment (45), but declined to 250-400 mmHg (like healthy skin) in subsequent treatments, indicating new blood vessel maturity in radiation-damaged tissue. PtcO2 attains values that are greater than invasive electrode values. Marx et al. (46) found that transcutaneous oxygen values in irradiated tissue changed from an initial value of 20% to an end value of 80% of non-irradiated control tissue over a course of about 20 HBO₂ treatments. In experimental burn wounds of rats, Ketchum et al. (47) had previously demonstrated extensive capillary proliferation by Day 18 of daily HBO₂ treatments. Angiogenesis in irradiated tissue was shown to exhibit a dose response to O₂ that improved as the partial pressure of oxygen was increased in 0.5 atm increments up to 3 atm abs (48,49).

TISSUE OXYGEN ASSESSMENT WITH PtcO₂

Transcutaneous oximetry has become increasingly popular as a tool for vascular assessment of amputation sites and for wound assessment for HBO₂ treatment. Tissue O₂ assessment is used to help define those patients within a given diagnosis that will benefit from HBO₂ treatment and, in some cases, to determine when treatment is complete.

Transcutaneous oximetry as a predictor of successful amputation: A number of studies of patients who did not receive HBO₂ have confirmed that normal wound healing of an amputation site requires a PtcO₂ value of about 40 mmHg or an extremity-to-chest ratio (limb PtcO₂ / chest PtcO₂) of about 0.6.

White and Klein (50) reviewed eight patient studies of PtcO₂ values for 260 amputees who did not receive HBO₂ and concluded that a PtcO₂ value of 40 mmHg or greater was required for successful outcome of amputation. Values less than 40 mmHg suggested an unsuccessful outcome unless revascularization could be combined with the amputation.

In a case-controlled study, Reiber et al. (51) computed odds ratios for a number of risk factors associated with

lower extremity amputation in diabetic patients, where <1 suggested a protective effect and >1 suggested increased risk. Patients with below-knee and dorsal foot PtcO₂ values above 40 mmHg served as a reference group against which the study groups were compared. Patients with PtcO₂ values of 20–40 mmHg had an odds ratio of 7.5 (Cl 4.0–14.1), whereas PtcO₂ values <20 mmHg yielded an odds ratio of 161 (CI 55–469). Thus, the lower the PtcO₂, the greater the risk of amputation.

In a blinded prospective study of 119 amputations, Harward and colleagues (52) compared PtcO₂ values before and after 10 min of 100% O₂ inhalation, and concluded that an increase above 10 mmHg was predictive of successful outcome.

Hauser (53) prospectively assessed 159 wounds (93 local debridements and 66 amputations) in 113 high-risk patients with diabetes mellitus and peripheral vascular disease who did not receive HBO₂. They measured PtcO₂ values at four sites on the limbs and compared extremity-to-chest ratios (RPI = wound PtcO₂ / chest PtcO₂) that reflected healing success. Of 93 local procedures, 48 healed (RPI = 0.72 ± 0.10) and 45 failed (RPI = 0.25 ± 0.12). Of 66 amputations, 45 healed (RPI = 0.64 ± 0.09) and 21 failed (RPI = 0.28 ± 0.11). Data in Table 2 reveal excellent outcome when RPI was >0.6, and poor outcome when RPI was <0.4.

Thus, patients whose PtcO₂ values at the amputation site are below 40 mmHg or whose PRI values are below 0.6 are potential candidates for HBO₂ and aggressive wound management (12,38,54). In many cases, mapping of the skin surface with PtcO₂ values at multiple amputation sites will offer the surgeon options for better limb salvage.

Transcutaneous oximetry as a predictor of wound healing: PtcO₂ values exceeding 30–40 mmHg are generally reported to be required for wound healing, but institutions vary in the absolute values.

Wyss et al. (55) reviewed a series of 188 diabetic patients with peripheral vascular disease. They reported that those with leg/foot PtcO₂ values below 20 mmHg were significantly more likely to have ulcers, rest pain, and amputation of the limb, as compared to those with PtcO₂ values above 20 mmHg.

Wyss et al. (37) measured PtcO₂ at the foot and proximal

Table 2: Wound Healing vs. Regional Perfusion Index (Hauser)

Wound Status	RPI = 0-0.4	RPI = 0.41-0.6	RPI > 0.61	
Healed, $n = 86$	2	22	61	
Delayed healing, $n = 8$	2	5	1	
Failed, $n = 73$	60	11	2	

and distal to the knee in 162 patients who then had 206 amputations. An increasing probability of failure was correlated with decreasing PtcO₂. Values >40 mmHg were associated with good healing, 20–40 mmHg with intermediate healing, and <20 mmHg with poor healing.

Gorman (56) conducted a review of diabetic foot wounds in which he used relative PtcO₂ (percent of chest control) and ankle perfusion pressure to predict treatment outcome. He concluded that patients who have relative PtcO₂ values >85 % and ankle perfusion pressure >90 mmHg will heal without O₂. Conversely, those with relative PtcO₂ values <20% and ankle perfusion pressure <75 mmHg are unlikely to heal. Gorman's conclusions are summarized at Table 3.

In 1983, Sheffield and Workman (41,44) reported the first known PtcO₂ data recorded under hyperbaric conditions (100% O₂ at 2.4 atm abs, 238 kPa). Using a Radiometer TCM1 oximeter to monitor both healthy subjects (*n* = 18) and patients (*n* = 4), values exceeding 1,000 mmHg were reported (41). During the course of HBO₂, two patients healed (1 diabetic, 1 scleroderma) whose baseline air PtcO₂ values were elevated >30 mmHg, but two diabetic patients did not heal whose PtcO₂ values fell below 30 mmHg. In 1994, Matos and Nunez (38) concluded that transcutaneous oxygen measurement was the best presently available method to determine the need for HBO₂ before initiation of therapy.

Pecoraro et al. (57) found $PtcO_2$ values to be useful as predictors of healing in diabetic patients and in selecting patients for adjunctive HBO_2 to correct underlying tissue hypoxia, either alone or in combination with revascularization. Marx et al. (46) reported an RPI of 0.8 in irradiated tissue of the face and neck within 4 wk of starting HBO_2 therapy. Sheffield and Workman (58) reported improved baseline $PtcO_2$ at the wound site taken at 7-day intervals while the patients received HBO_2 therapy. Sheffield (59) retrospectively found that diabetic patients with forefoot wounds (n = 84) had an 8-fold increase in likelihood of successful outcome with HBO_2 when baseline transmetatarsal $PtcO_2$ values were >30 mmHg as compared to $PtcO_2$ values <30 mmHg (P < 0.05).

Measurements of PtcO₂ under hyperbaric conditions:

There is controversy as to which environment for measuring PtcO₂ has the best predictive value: atmospheric conditions or HBO₂ A number of investigators have suggested that the best predictor of wound healing success would be PtcO₂ conducted under hyperbaric conditions, but the suggested minimum values vary widely.

Myers and Emhoff (60) concluded that diabetic patients (n = 11), with below-knee PtcO₂ values <20 mmHg, would heal if 900 to 1100 mmHg could be achieved on the initial HBO₂ exposure (100% O₂ at 2.5 atm abs, 252 kPa). Healing did not occur if the values were below 900 mmHg.

Wattel et al. (61) found that in diabetic foot lesions (n =59), PtcO₂ values on normobaric O₂ did not discern between healing and failure. Conversely, PtcO2 values above 450 mmHg during HBO₂ (100% O₂ at 2.5 atm abs, 252 kPa) were predictive of healing in diabetic patients with plantar ulcers, with the healing group having significantly higher PtcO₂ than the failed group $(786 \pm 258 \text{ vs.})$ 323 ± 214 mmHg; P < 0.005). Campagnoli et al. (62) reported that diabetic patients (n = 24) healed if PtcO₂ values were above 400 mmHg during HBO2. The authors observed that the faster the rise, the greater likelihood of an efficient support microcirculation, and the better the patients' chances of a favorable outcome. Hart et al. (43) measured seven patients with ischemic limbs (100% O₂ at 2.5 atm abs, 202 kPa) and reported that limb salvage occurred in four patients with PtcO2 values above 100 mmHg, but three amputees had PtcO2 values below 100 mmHg. Strauss et al. (63) measured PtcO2 in 87 patients with problem foot wounds and reported that 98% healed when PtcO₂ measured over 200 mmHg during HBO₂ (100% O₂ for 90 min at 2 atm abs, 202 kPa). In contrast, only 17% (2 of 12) of foot wounds healed when PtcO₂ adjacent to the wound was less than 100 mmHg.

SPECIAL CONSIDERATIONS FOR REPRODUCIBLE PtcO₂ DATA IN THE CLINICAL SETTING

At this writing, PtcO₂ is as much an art as a science. Data collection requires a skilled technician using a rigorous protocol. Interpretation of data requires a complete understanding of the limits of the technology. To obtain repro-

Table 3: Use of PtcO₂ to Predict Healing of Diabetic Wounds of the Foot (Gorman)

	Ankle Pressure, mmHg	Relative PtcO2, % Chest Control	
Will heal without O2	>90	>85	
Probably will heal with O2	75–90	20-85	
Unlikely to heal	<75	<20 ^a	

[&]quot;Exception: when PtcO₂ rises >50% within 15 min of initiating O₂ breathing.

ducible results, it is important to standardize the technique. At this writing, however, there is no consensus of the best method to collect the data.

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Method of assessment: A number of methods have been used to assess healing potential: 1) a single PtcO₂ value adjacent to the wound; 2) a map of multiple sites around the wound; 3) a map of several sites on the affected limb; and 4) a comparison of periwound or amputation site values expressed as a percentage of control values at the chest. The number of sites to be measured is usually determined by the number of sensors available. The preferred method seems to be a map of at least three sites. But regardless of the assessment method, it is important to be consistent.

Selection of the monitor: Monitors from several manufacturers have been used for $PtcO_2$ studies (43,50,64). As of this writing, the TINA TCM3 and TCM30 (Radiometer, Copenhagen) are the only monitors with sensors that have been tested and shown to be compatible with operation in a chamber filled with pure O_2 (64). Although the sensor is inside the pure O_2 chamber, fire safety considerations require the monitor to remain outside, connected to the sensor via a through-hull penetrator.

Calibration of sensor: It is advisable to install a new membrane weekly and to follow the manufacturer's instructions for calibration. It is also advisable to use a two-point calibration in the range of values that are anticipated and to correct for ambient temperature and barometric pressure. Some devices have the zero point determined electronically rather than requiring a zero-oxygen gas or zero-oxygen solution.

Selection of sensor temperature: Transcutaneous sensors measure the approximate arterial O₂ values at the site chosen. Heating the skin causes physiologic changes beneath the sensor: opens pores, dilates capillaries, increases skin perfusion, increases metabolism, and causes hemoglobin to release O2. A sensor temperature of 45°C produces maximum arterialization of Po, and can be tolerated by most patients. An occasional patient with poor perfusion will experience a blister beneath the electrode if it remains in place in excess of an hour. For this reason, a 44°C sensor temperature is used at the Jefferson C. Davis Wound Care and Hyperbaric Medicine Center. Thus, values obtained in this laboratory will be about 2% lower than other laboratories that collect PtcO₂ at 45 °C. For whatever sensor temperature setting is chosen, it is important to be consistent, and to report the temperature in scientific reports.

Standardized site of measurement: Considering normal circulation to the limb, there should be a standard approach to positioning the sensor. This will provide more consistent data and allow a means of comparing data among a group of patients.

Site preparation for electrode placement: The sensor site should be shaved, cleaned with alcohol, and stripped of superficial layers of dry, loose skin cells using adhesive tape. The sites should not overlie a bony prominence, superficial vessels, or pulse sites (65). Flat or slightly convex areas provide the most reliable sensor contact. The fixation ring should be checked for tightness of seal from the surrounding air. If multiple sites are to be measured, the fixation rings should be positioned in the same direction so that the electrode wires will all be positioned in the same direction. This reduces movement artifacts and risk of electrode displacement should the patient move.

Adequate time for electrode equilibration and PtcO2 evaluation: For 15 yr the Jefferson C. Davis Wound Care and Hyperbaric Medicine Center, has conducted PtcO₂ assessment of candidates for HBO2. The procedure has evolved into a 45-min assessment as shown in Table 4. Acquiring a baseline PtcO2 value of the supine patient takes 20 min. As the sensor heats the skin, equilibration of the electrode (Fig. 1) occurs within 10-15 min for subjects with normal circulation, but requires about 15-20 min for patients with compromised circulation (Vesterager P, 1997, personal communication). The limb is elevated 45° as a challenge to identify presence of large or small vessel disease that would cause Po₂ to fall and not recover until the patient is returned to the supine position. After the sensor is allowed to return to baseline, a 10-min O₂ challenge is administered to determine the response to O_2 of the wounded area. The sensor is allowed to return to baseline after each physiologic challenge to check for electrode drift. Some would argue that this assessment takes too much time and would be satisfied with a 30-min assessment that includes only baseline PO2 and the O2 challenge.

Interpretation of data: Methods of interpreting the data vary among investigators. Some investigators predict success by interpreting the raw data from the actual PtcO₂ values. Some use a "relative value", which is the value near

Table 4: Time Required for PtcO₂ Assessment by Mapping the Skin Surface

Assessment	Time Required, min		
Electrode equilibration, air	15		
Baseline PtcO2, air	5		
Elevate limb, air	5		
Baseline PtcO2, check for			
electrode drift	5		
100% Oxygen challenge	10		
Baseline PtcO2, check for			
electrode drift	5		
Total evaluation time	45		

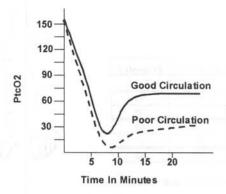


FIG. 1—Electrode behavior during equilibrium (as drawn by P. Vesterager, 1997).

the wound as compared to the chest control, or a mirror image site on the opposite limb, and expressed as a percentage. Others calculate a limb-to-chest ratio, called a "regional perfusion index" (RPI = limb PtcO2 / chest PtcO₂). As of this writing there is no consensus on the best method for interpreting the data. However, it is clear that interpretation demands careful assessment of the tissue on which the sensor is placed. PtcO₂ values can be elevated if the sensor is positioned over superficial vessels or if there is a leak under the fixation ring. PtcO₂ values can be lowered if the sensor is positioned over bony protuberances or if the patient uses tobacco products. PtcO2 values in smokers were 10% below nonsmokers (41) and remained significantly reduced for about an hour after smoking (40). But there is evidence of improved tissue oxygenation after a few days of smoking cessation (66).

Transcutaneous oximetry values may also be lowered by several pathologic conditions: edema, active infection/inflammation, thick or sclerotic skin, occluded vessels, severed vessels (flap), ischemic lesions, or irradiated tissue. Plantar PtcO₂ readings are different from non-plantar and should be avoided or used with an understanding of how they should be interpreted. The interpreter of the data must consider all these factors.

Interpretation of PtcO₂ in case AAI 530 TG. This 50-yr-old, non-insulin-dependant diabetic patient with acute arterial insufficiency had an open left 3rd ray amputation. There was an ischemic wound margin, edema, and inflammation in the wound area. The typical PtcO₂ map of standardized sites on the leg is at Fig. 2. PtcO₂ assessment showed presence of vascular disease (electrodes A, D, E, and F), severe hypoxia over the foot (electrodes C and E), but a good response to the O₂ challenge at 1 atm abs (electrodes A, B, D, and E). The physician interpreting the results recommended aggressive wound care and adjunctive HBO₂ to help control infection and prevent spread of necrosis (67).

A PROTOCOL FOR PtcO₂ ASSESSMENT OF HBO₂ CANDIDATES

Assessment protocol: The PtcO₂ assessment of potential HBO₂ candidates usually consists of four basic questions about whether the wound: 1) is severely hypoxic, 2) will respond to O₂, 3) will respond to HBO₂, and 4) is to the point that it will heal without further treatment (54). The questions and their respective tests are shown in Table 5.

Transcutaneous oximetry values used to select HBO₂ candidates. Not all problem wounds are hypoxic, but may fail to progress for other reasons. Some patients with PtcO₂ values near zero will heal, whereas others with values above 40 mmHg will not. As indicated in the above studies, there is a wide range of proposed threshold values for healing, but the numbers are not absolute. Each wound care and hyperbaric medicine center has established its own criteria for selection of HBO₂ patients.

At the Jefferson C. Davis Wound Care and Hyperbaric Medicine Center, transcutaneous mapping of the affected extremity is essential in the acceptance or rejection of patients for HBO₂ (54). Candidates who meet the following criteria are generally accepted for HBO₂ (54). Since at least six sites are being evaluated simultaneously, not all sites will satisfy every test. However, the sensors must show the presence of hypoxia (test 1) and a reasonable response to oxygen (test 2 or 3).

- Test 1: Baseline air value at 1 atm abs. If all measured sites are above 30 mmHg (>40 mmHg for diabetic patients), tissue oxygenation is considered adequate for healing and the patient is usually not selected for HBO₂. If sites near the wound fall below these values, wound healing is complicated by hypoxia and the patient is accepted for HBO₂, provided test 2 or 3 is successful. Values of 20–40 mmHg are considered to be of intermediate healing potential, and values below 20 mmHg are associated with poor healing. A PtcO₂ value that is less than 10 mmHg is usually not a good prognostic sign for successful HBO₂ outcome (54). However, individual cases may vary and some patients with PtcO₂ values near zero are treated if clinical judgment justifies it.
- Test 2: Oxygen challenge at 1 atm abs. An oxygen challenge is essential to the PtcO₂ assessment to determine if the wounded area will respond to O₂ administration. The value should at least double the baseline, and above 30 mmHg is desirable (59). However, an increase of 10 mmHg has been shown to be successful (52). If a baseline value of 10 mmHg only reaches 20 mmHg during O₂ challenge, the patient would be selected for HBO₂, but would probably have a prolonged course of treatment unless there is some underlying temporary cause for the low O₂ response. The O₂ challenge is

	L	Control	Electrode A	Electrode B	Electrode C	Electrode D	Electrode E	Electrode F
					PtcO2			
Supin	20 min. Air	57	44	50	2	45	9	58
45 Elev Supin.	5 min. Air	54	25	39	1	24	2	39
Supin	5 min. Air	50	40	41	2	43	7	49
Supin	10 min. O2	290	79	88	4	178	40	67
Supin	5 min. Air	96	48	47	3	47	10	62

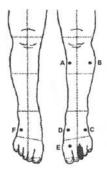


FIG. 2—Typical PtcO₂ mapping of the limb with 6 electrodes plus chest control.

Table 5: Protocol for PtcO2 Assessment

Question	Test		
Is wound healing complicated by severe hypoxia?	 Baseline air value at 1 atm abs; hypoxia exists if PtcO₂ <40 mmHg for diabetics, < 30 mmHg for non-diabetics. 		
Does the wound site respond to O2 breathing?	O₂ challenge at 1 atm abs; value should be at least double the baseline value.		
Does the wound site respond to HBO ₂ ?	 O₂ challenge 2–2.5 atm abs; value should be well above normobaric O₂ value. 		
Is the patient's wound at the point where it will heal without further treatment?	 Repeat PtcO₂ evaluations at 1 atm abs in 2- to 4-wk intervals; normalization of baseline values would indicate that the healing process is in place. 		

"Values are from experience at the Jefferson C. Davis Wound Care & Hyperbaric Medicine Center (Heimbach RD, 1998, personal communication).

sometimes conducted in the hyperbaric chamber (61).

- Test 3: Oxygen challenge at 2–2.5 atm abs. The HBO₂ challenge is used to document that the patient is responding to the prescribed HBO₂ dose. The test is usually limited to one to three sites. Recommend minimum values range from 100–450 mmHg (61,63). At Baystate Medical Center in Springfield, Massachusetts, pressure in the chamber is increased beyond 2 atm abs until the prescribed oxygen dose (PtcO₂ = 400–600 mmHg or above) is achieved (Emhoff TA, 1998, personal communication).
- Test 4: Repeat baseline air value at 1 atm abs. Baseline PtcO₂ values should increase over a course of HBO₂ (44). Normalization of the PtcO₂ values (usually above 30–40 mmHg) indicates that the healing process is in place. An increase in PtcO₂ baseline over a course of HBO₂ should achieve a value that is normal for that patient. If the chest control value in an elderly patient is only 25 mmHg, it is unlikely that a greater value will be achieved in the affected limb, even though wound healing is achieved.

In summary, tissue O₂ measurements have confirmed hypoxia in chronic, indolent human wounds and have demonstrated that HBO₂ elevates wound O₂ tension. Tissue O₂ assessment is a valuable tool in the medical decision-

making process for healing both acute and chronic soft-tissue problem wounds, and for helping define amputation levels. Sufficient clinical experience is published to justify PtcO₂, but measurement requires rigorous protocol and the interpreter must understand the limits of the technology. Best results are obtained from a dedicated, qualified technician who measures at standardized sites on the limb. The data suggest that the best predictor of success with HBO₂-treated cases would be PtcO₂ collected under hyperbaric conditions.

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REFERENCES

- Hunt TK, Twomey P, Zederfeldt B, et al. Respiratory gas tension and pH in healing wounds. Am J Surg 1967; 114:302–308.
- Niinikoski J. Effect of oxygen supply on wound healing and formation of experimental granulation tissue. Acta Physiol Scand, 1969; 334:1-72.
- Silver IA. The measurement of oxygen tension in healing tissue. In: Herzog H, ed. Progress in respiration research, III. Basel: S Karger, 1969:124–135.
- Hunt TK, van Winkle W Jr. Wound healing: normal repair. In: Dunphy JE, ed. Fundamentals of wound management in surgery. South Plainfield, NJ: Chirurgecom, 1976:1–68.
- Hunt TK, Pai MP. The effect of varying ambient oxygen tensions on wound metabolism and collagen synthesis. Surg Gynecol Obstet 1972; 135:561–567.

- Winter GD, Perrins DJD. Effects of hyperbaric oxygen treatment on epidermal regeneration. In: Wada J, Iwa T, eds. Proceedings of the fourth international congress on hyperbaric medicine, Baltimore, MD: Williams & Wilkins, 1970:363

 –368.
- Hunt TK. Disorders of repair and their management. In: Hunt TK, Dunphy JE, eds. Fundamentals of wound management. New York: Appleton-Century-Crofts, 1979:68–168.
- Niinikoski J, Henghan C, Hunt TK. Oxygen tensions in human wound. J Surg Res 1972; 12:77.
- Sheffield PJ. Tissue oxygen measurements with respect to softtissue wound healing with normobaric and hyperbaric oxygen. Hyper Oxygen Rev 1985; 6:18–46.
- Lanphier EH, Brown IW Jr. The physiological basis of hyperbaric therapy. In: Fundamentals of hyperbaric medicine. Washington DC: National Academy of Sciences-National Research Council, 1966:33-55.
- Krogh A. The number and distribution of capillaries in muscle with calculations of the oxygen pressure head necessary for supplying the tissue. J Physiol 1919; 52:409

 –415.
- Sheffield PJ. Tissue oxygen measurements. In: Davis JC, Hunt TK, eds. Problem wounds: the role of oxygen. New York: Elsevier, 1988:17–51.
- Camporesi E, chairman. Tissue oxygen measurements. In: Hyperbaric oxygen therapy, a committee report. Kensington, MD: Undersea and Hyperbaric Medical Society, 1996:63–64.
- Davies PW, Brink F. Microelectrodes for measuring local oxygen tension in animal tissues. Rev Sci Instr 1942; 13:524–533.
- Clark CL Jr, Wolf R, Granger D, et al. Continuous recording of blood oxygen tensions by polarography. J Appl Physiol 1953; 6:189-193.
- Silver IA. Some observations on the cerebral cortex with an ultramicro, membrane covered, oxygen electrode. Med Electr Biolog Eng 1965; 3:377–387.
- Fatt I. Polarographic oxygen sensor: its theory of operation and its application in biology, medicine, and technology. Cleveland, OH: CRC Press, 1976.
- Brantigan JW, Gott VL, Martz MN. A teflon membrane for measurement of blood and intramyocardial gas tensions by mass spectroscopy. J Appl Physiol 1972; 32:276–282.
- Wells CH, Goodpasture JE, Horrigan DJ, et al. Tissue gas measurements during hyperbaric oxygen exposure. In: Smith G, ed. Proceedings of the sixth international congress on hyperbaric medicine. Aberdeen, Scotland: Aberdeen University Press, 1977:118–124.
- Woldring S, Owens G, Woolford D. Blood gases: continuous in vivo recording of partial pressures by mass spectrography. Science 1966; 153:885–887.
- Hunt TK. A new method of determining tissue oxygen tension. Lancet 1964; 2:1370–1371.
- Gottrup F, Firmin R, Chang N, et al. Continuous direct tissue oxygen tension measurement by a new method using an implantable silastic tonometer and oxygen polarography. Am J Surg 1983; 146:399–403.
- Sheffield PJ, Davis JC, Bell GC, Gallagher TJ. Hyperbaric clinical support: multiplace. In: Davis JC, Hunt TK, eds. Hyperbaric oxygen therapy. Bethesda, MD: Undersea Medical Society, 1977:25–39.
- Martindale VE, Stewart KM, Norris TD. Comparison of operating characteristics of the StatPal II blood gas analyzer at 45 fsw. Abstracts of UHMS Gulf Coast and Midstate Chapters Annual Scientific Meeting, Memphis TN, March 24–27, 1994
- 25. Weaver LK, Howe S. Normobaric measurement of O2 tension of

- blood in subjects exposed to hyperbaric oxygen. Chest 1992; 102:1175-1181.
- Moon RE, Camporesi EM, Shelton DL. Prediction of arterial Poduring hyperbaric treatments. In: Bove AA, Bachrach AJ, Greenbaum LJ Jr, eds. Proceedings of the ninth international symposium on underwater and hyperbaric physiology, Bethesda, MD: Undersea and Hyperbaric Medical Society, 1987:1125–1131.
- Howe S, Hein S, Weaver LK, et al. Response of arterial oxygen tension (PaO2) and transcutaneous oxygen (TcPO2) to changes in alveolar PO2. Undersea Hyper Med 1995; 22 (suppl):76.
- Baumberger JP, Goodfriend RB. Determination of arterial oxygen tension in man by equilibration through intact skin. Fed Proc 1951; 10:10–11.
- Huch R, Lubbers DW, Huch A. Quantitative continuous measurement of partial oxygen pressure on the skin of adults and newborn babies. Pflugers Arch 1972; 337:185–198.
- Lubbers DW. Theoretical basis of the transcutaneous blood gas measurements, Crit Care Med 1981; 9:721–733.
- Evans NTS, Naylor PFD. The systemic oxygen supply to the surface of the human skin. Respir Physiol 1967; 3:21–37.
- Huch R, Huch A. Fetal and maternal PtcO2 monitoring. Crit Care Med 1981; 9:694–697.
- Ravindranath T. Non-invasive monitoring in the pediatric ICU, part I: Transcutaneous oxygen monitoring (PtcO2). Indian J Pediatr 1990; 57:169–173.
- Serafin D, Lesense CV, Mullen RY, et al. Transcutaneous PO2 monitoring for assessing viability and predicting survival of skin flaps: experimental and clinical correlations. J Microsurg 1981; 2:165-178.
- Rooke TW. The use of transcutaneous oximetry in the noninvasive vascular laboratory. Int Angiol 1992; 11:36–40.
- Dennhardt R, Ricke MF, Huch A, et al. Transcutaneous PO2 monitoring in anaesthesia. Eur J Intens Care Med 1976; 2:29–33.
- Wyss CR, Harrington RM, Burgess EM, et al. Transcutaneous oxygen tension as a predictor of success after an amputation. J Bone Jt Surg 1988; 70-A:203-207.
- Matos LA, Nunez AA. Enhancement of healing in selected problem wounds. In: Kindwall EP, ed. Hyperbaric medicine practice. Flagstaff AZ: Best Publishing; 1994:589–612.
- Simanonok J. Transcutaneous oximetry. Triage 1996; 8:1,7.
- Jensen JA, Goodson WH, Hopf HW, Hunt TK. Cigarette smoking decreases tissue oxygen. Arch Surg 1991; 126:1131–1134.
- Workman WT, Sheffield PJ. Continuous transcutaneous oxygen monitoring in smokers under normobaric and hyperbaric oxygen conditions. In: Huch R, Huch A, eds. Continuous transcutaneous blood gas monitoring, New York: Marcel Dekker, 1983:649–656.
- Dooley J, King G, Slade B. Establishment of reference pressure of transcutaneous oxygen for the comparative evaluation of problem wounds. Undersea Hyper Med. 1997; 24:235–244.
- Hart GB, Meyer GW, Strauss MB, et al. Transcutaneous partial pressure of oxygen measured in a monoplace chamber at 1, 1.5, and 2 atm abs oxygen. J Hyper Med 1990; 5:223–229.
- Sheffield PJ, Workman, WT. Transcutaneous tissue oxygen monitoring in patients undergoing hyperbaric oxygen therapy. In: Huch R, Huch A, eds. Continuous transcutaneous blood gas monitoring. New York: Marcel Dekker, 1983:655-660.
- Sheffield PJ, Dunn JM. Continuous monitoring of tissue oxygen tension during hyperbaric oxygen therapy. In: Smith G, ed. Proceedings of the sixth international congress on hyperbaric medicine. Aberdeen, Scotland: Aberdeen University Press, 1979:125–129.
- 46. Marx, RE, Johnson RD, Kline SN. Prevention of osteoradionecro-

- sis: a randomized prospective clinical trial of hyperbarie oxygen versus penicillin. J Am Dent Assoc 1985; 3:49-54.
- Ketchum SA III, Thomas AN, Hall AD. Angiographic studies of the effects of hyperbaric oxygen on burn wound revascularization.
 In: Wada J, Iwa T, eds. Proceeding of the fourth international congress in hyperbaric medicine. Baltimore, MD: Williams & Wilkins, 1969:388–394.
- Marx RE, Ehler WJ, Tayapongsak P, et al. Relationship of oxygen dose to angiogenesis induction in irradiated tissue. Am J Surg 1990; 160:519-524.
- Ehler WJ, Marx RE, Paleo MJ. Oxygen as a drug: a dose response curve for radiation necrosis. Undersea Hyper Med 1993; 20 (suppl):44–45.
- White RA, Klein SR. Amputation level selection by transcutaneous oxygen pressure determination. In: Moore WS, Malone JM, eds. Lower extremity amputation. Philadelphia, PA: WB Saunders, 1989:44-49
- Reiber GE, Pecoraro RE, Koepsell TD. Risk factors for amputation in patients with diabetes mellitus. Ann Intern Med 1992; 117:97–105.
- Harward TRS, Volny J, Golbranson F, et al. Oxygen inhalationinduced transcutaneous PO2 changes as a predictor of amputation level. J Vasc Surg 1985; 2:220–227.
- Hauser CJ. Tissue salvage by mapping of skin surface transcutaneous oxygen tension index. Arch Surg 1987; 22:1128–1130.
- Brakora MJ, Sheffield PJ. Hyperbaric oxygen therapy for diabetic wounds. In: Lavery LA, Dennis KJ, eds. Clinics in podiatric medicine and surgery: the diabetic foot. Philadelphia, PA: WB Saunders, 1995; 12:105–117.
- Wyss CA, Matsen FA III, Simmons CW, et al. Transcutaneous oxygen measurements on limbs of diabetic and nondiabetic patients with peripheral vascular disease. Surgery 1984; 95:339–345.
- Gorman DF. Oxygen therapy in the diabetic foot. In: Frykberg RG,
 ed. The high risk foot in diabetes mellitus. New York: Churchill

- Livingstone, 1991:441-447.
- Pecoraro RE, Ahroni JH, Boydo EJ, Stensel VL. Chronology and determinants of tissue repair in diabetic lower-extremity ulcers. Diabetes 1991; 40:1305–1312.
- Sheffield PJ, Workman WT. Noninvasive tissue oxygen measurements in patients administered normobaric and hyperbaric oxygen by mask. Hyper Oxygen Rev 1985; 6:47–62.
- Sheffield PJ. Measuring tissue oxygen tension: a review. Undersea Hyper Med 1998; 25(3):179–188.
- Myers RAM, Emhoff TA. Transcutaneous oxygen measurements in the non-healing diabetic wound. In: Program and abstracts. Eight international congress on hyperbaric medicine, Bethesda, MD: Undersea Medical Society, 1984:185–186.
- Wattel FE, Mathieu MD, Fossati P, Neviere RR, Coget JM. Hyperbaric oxygen in the treatment of diabetic foot lesions. Search for healing predictive factors. J Hyper Med 1991; 6:263–268.
- Campagnoli P, Oriani G, Sala G, et al. Prognostic value of TcPo₂ during hyperbaric oxygen therapy. J Hyper Med 1992; 7:223–227.
- Strauss MJ, Breedlove JW, Hart GB. Use of transcutaneous oxygen measurements to predict healing in foot wounds. Undersea Hyper Med 1997; 24(suppl):15.
- Marotte S, Larson-Lohr V, Weaver LK. Performance evaluation of transcutaneous oxygen (TcPO2) monitors. Undersea Hyper Med 1993; 20(suppl):28.
- Clarke D. Transcutaneous monitoring of pO2 in hyperbaric medicine. Patient focus circle. Copenhagen, Denmark: Radiometer Medical A/S, 1997.
- Strauss AG, Hart GB, Strauss MB. Effect of smoking cessation on transcutaneous oxygen measurements—a case report and review. Undersea Hyper Med 1997; 24 (suppl):36.
- Stegmann BJ. PtcO2 Interpretation in wound assessment. Proceedings of the clinical management of problem wounds symposium VI. San Antonio, TX: International ATMO, 1997.