Flap Monitoring Using Transcutaneous Oxygen or Carbon Dioxide Measurements

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Abstract

Free tissue transfer is a cornerstone of complex reconstruction. In many cases, it represents the last option available for a patient and their reconstruction. At high-volume centers, the risk of free flap failure is low but its occurrence can be devastating. Currently, the mainstay for flap monitoring is the clinical examination. Though reliable when performed by experienced clinicians, the flap exam is largely subjective, is performed discontinuously, and often results in significant time delay between detection of flap compromise and intervention. Among emerging flap monitoring technologies, the most promising appear to be those that rely on noninvasive transcutaneous oxygen and carbon dioxide measurements, which provide information regarding flap perfusion. In this article, we review and summarize the literature on various techniques but primarily emphasizing those technologies that rely on transcutaneous gas measurements. We also define characteristics for the ideal flap monitoring tool and discuss critical barriers, predominantly cost, preventing more widespread utilization of adjunct monitoring technologies, and their implications.

Keywords

- flap monitoring
- transcutaneous oxygen
- carbon dioxide
- systematic review

Introduction

Free tissue transfer is a cornerstone of complex reconstruction. In many cases, it represents the last option available for a patient and their reconstruction. At the highest volume centers, the risk of free flap failure is low but if it occurs, it may be a devastating result. Currently, the mainstay of flap monitoring includes clinical flap examination with possible adjunct monitoring techniques that vary by facility. Though reliable when performed by experienced clinicians, the flap examination is largely subjective, not done continuously, and often results in significant time delay between detection of flap compromise and intervention. Ideal postoperative surveillance of free flaps would consist of a monitoring system that is accurate, reliable, easily reproducible with a shallow learning curve, non- or minimally invasive, a realtime assessment, affordable, and provides an ability to differentiate arterial from venous compromise, in addition

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to vessel spasm from occlusion.¹ Flap failure is relatively uncommon with success or flap survival rates of 97 to 99%²; however, flap failure can lead to significant morbidity and, in some cases, mortality risk. Flap compromise generally occurs as a result of arterial or venous occlusion by thrombosis, external compression, vessel kinking, or hematoma.² Current flap monitoring mechanisms have led to a highly variable salvage rate of 40 to 80% with the greater salvage rates in instances of venous thrombosis than in arterial thrombosis.² Flap tissue oxygen saturation measurement has been found to significantly correlate with blood oxygenation readings without being confounded by blood pressure, supplemental oxygen (O_2) , flap type, perforator number, or vessel caliber.³ The literature shows flap success rates and salvage rates are generally equivalent, regardless of the monitoring system used, although transcutaneous oxygen and carbon dioxide monitoring have demonstrated the most promise in its resurgence with studies showcasing their versatility and ability to

© 2020. Society of Indian Hand & Microsurgeons. All rights reserved. Thieme Medical and Scientific Publishers Pvt. Ltd., A-12, 2nd Floor, Sector 2, Noida-201301 UP, India DOI https://doi.org/ 10.1055/s-0040-1718862 ISSN 0974-3227. identify early flap compromise. In this article, we performed a systematic review adherent to preferred reporting items for systematic reviews and meta-analyses (PRISMA) guidelines and summarize the literature on flap monitoring techniques, with particular attention to those using transcutaneous gas measurements.

Systematic Review of the Literature

Methods

A PubMed, Cochrane Database, and MEDLINE search adherent to PRISMA was performed for articles containing the terms: "flap," "monitoring," and/or "transcutaneous," "oxygen," "carbon dioxide." Search terms were stringed using PubMed's Automatic Term Mapping algorithm. Articles were limited to English with humans as the subjects for this study.

Retrospective studies and prospective analyses were included; case reports, case series fewer than five patients, commentaries, and editorials were excluded. The lead author (S.H.H.) reviewed the articles and two authors (T.M.S., A.S.H.) double-screened articles to decide which to include or exclude; discrepancies or indecisions were resolved via discussion with the other authors. Individual study bias was mitigated by reviewing and confirming the appropriate sources indicated. Studies were limited to those published since 2000.

Title and abstract reviews were performed once the initial list of studies was generated (► Fig. 1). The reviewers met prior to commencing study selection to ensure consistency in the application of the inclusion criteria. Studies were selected if they met the following inclusion criteria: (1) utilized transcutaneous gas measurements for flap monitoring, (2) included the details of flap survival and outcomes, and (3) detailed the measuring modality used for flap monitoring.

A full-text review was performed on all remaining studies following title and abstract review. Additional studies that did not meet inclusion criteria were excluded. The last search was performed on August 17, 2020.

Results

The initial PubMed/MEDLINE/Cochrane search revealed 17 articles eligible for full-text review; of these, 6 were included in both the qualitative and quantitative analysis.⁴⁻⁹ All studies included were prospective in nature (**Fig. 1**).

The studies collectively included 162 flaps, each with varying measurement modalities and devices. The overall flap failure rate was 4.3% (n = 7); the overall consensus from these studies demonstrated transcutaneous oxygen partial pressure (pO_2) monitoring in conjunction with physical exam findings of flap compromise were critical to avoiding flap failure and improving flap salvage rates. Notably, drops in pO_2 levels acutely (i.e., within 4–24 hours postoperatively) were indicative of a failing flap. Additionally, severely congested flaps requiring intervention demonstrated markedly increased levels of carbon dioxide partial pressure (pCO_2) levels. These studies are summarized in **~ Table 1**.

Study Quality and Risk of Bias

Given the paucity of well-designed studies on flap monitoring in the past 20 years, the studies included in our quantitative meta-analysis scored 7 or greater on the Newcastle–Ottawa scale (NOS),¹⁰ signifying studies were of sound quality. The major biases present in our included studies were selection bias, given the institutions conducting this high-level work are filled with well-experienced microsurgeons with likely hundreds to thousands of free flaps under their belt, with a team of clinicians well versed in flap monitoring. Additionally, the operative indications/criteria used in individual studies was seldom highlighted and thus suggests the patient population in the combined cohort varies considerably.

Current Flap Monitoring Methods

Creech and Miller first described the ideal characteristics for flap monitoring in 1975 and since then, authors have continued to detail optimal flap monitoring techniques.⁴ Ideal qualities for any such device may include but are not limited to accurate, reproducible, real time with continuous acquisition of data, quantitative, easy to use or train, operator independent, safe, and low cost. Additionally, in an era of rapidly evolving technological advances, it would ideally be able to communicate encrypted data wirelessly to an electronic medical record or a mobile device, as this has shown promise in increasing flap survival and decreasing delay in return to the operating room with flap compromise.^{5,6} However, no such device currently exists, thus understanding current techniques is critical for optimizing flap monitoring and survival.

While clinical examination (color, turgor, temperature, capillary refill, and bleeding to pin prick) remains the gold standard for flap assessment, there exists a plethora of flap monitoring devices.¹¹ We provide a brief overview of the available options, along with their advantages and disadvantages based on literature review (**-Table 2**).^{3,4} Laser Doppler flowmetry (LDF) and near-infrared spectroscopy are not well-suited for buried flaps. Conversely, implantable Doppler, flow coupler, and microdialysis are invasive monitoring methods suitable for buried flaps.⁷

Transcutaneous Oxygen Monitoring

Although various methods exist that detect a change in blood flow to the flap, none are direct indicators of cellular metabolism or oxygen availability to the tissue. An absolute measurement of arterial oxygen for noninvasive flap monitoring was not described until Achauer et al studied the use of the polarographic Clark-type electrode in measuring transcutaneous oxygen. This was accomplished by measuring the electrical current flowing into the electrode, which is dependent on the number of oxygen molecules available to be reduced as they diffuse through the tissue and out from the skin.⁸

Early studies evaluating the use of a transcutaneous oxygen monitor showed it to be a quick and reliable indicator of flap perfusion.⁹ In evaluating free flaps and digital replants, Smith et al found transcutaneous oxygen monitoring to be an



Fig. 1 PRISMA flow diagram for transcutaneous oxygen and/or carbon dioxide measurements for free flap monitoring⁶¹. PRISMA, preferred reporting items for systematic reviews and meta-analyses.

overall accurate indicator of flap viability and able to detect impending flap failure before clinical signs and temperature changes. Moreover, they showed an increase in the success rate of reintervention using these measurements: a metric largely dependent on the speed of detecting vascular compromise.¹⁰ Although Achauer et al had noted a significant variability between individuals partly due to differences in systemic arterial pO_2 ,⁸ Serafin et al found that variation in core temperature, systemic blood pressure, and pulse had negligible effects on the transcutaneous oxygen measurements.⁹ The residual interobserver variability could be resolved by comparing the flap site measurements with measurements taken from a nonoperated site and then expressing the results as a percent change from "normal."¹²

Despite these promising early findings, there were still numerous flaws with using the Clark electrode. Although some of the interindividual variability could be accounted for, eschar formation, edema, and the measurement site being over bone decreased the reliability of measurement.⁸ Several studies also found that while transcutaneous O_2 (*ptcO*₂) may be a rapid indicator of vascular compromise, it may be too sensitive, so that during low-flow conditions in the early postoperative period, even flaps that go on to survive show low *ptcO*₂ values indistinguishable from those of

Study reference	No. of flaps	Monitoring modality	Value measured	Summary	Flap loss n (%)
Schiltz et al ⁴	32	VisiSens	Transcutaneous pO_2 based on tissue flux of O_2 using ratiometric fluorescence imaging	Increased O ₂ flux was associated with increased venous congestion and flap failure	2 (6.7)
Trignano et al⁵	21ª	Licox Recon	Measures tissue pO_2 and temperature through a Clark-type polarographic micro-catheter posi- tioned in the tip of a flexible probe with an oxygen sensing area	Low pO_2 combined with exam findings were indicative of venous congestion and flap failure	1 (5.5)
Geis et al⁵	9	Luminescent lifetime imaging	Transcutaneous pO_2 measurements using dynamic phosphorescence imaging and sensors	Drops in <i>pO</i> ₂ below 10 mmHg acutely (within 4 hours) required surgical intervention for flap salvage	0 (0)
Schrey et a. ⁷	13	Licox Polarographic Probe	Tissue pO_2 was measured using a needle probe in Hartrampf perfusion zone 1 of the flap	Needle probes monitoring pO_2 are reliable to evaluate a single portion of the flap but are inadequate at measuring blood flow throughout the entirety of a flap	1 (7.7)
Hashimoto et al ⁸	27	Polarographic probes	Transcutaneous pO_2 and pCO_2 were measured using probe contains a Clark-type electrode and a pH-sensitive glass elec- trode, respectively	Physical exam is a necessary adjunct. pO_2 alone is inadequate to determine venous congestion; a congested flap with a pCO_2 more than 90 mm Hg requires further treatment	1 (3.7)
Kamolz et al ⁹	60	Licox micropolaro- graphic probe	pO_2 was measured with a microcatheter in some centralized portion of the flap	Failing flaps demonstrate a tissue pO_2 decrease of 10 mm Hg within a half-hour, or a tissue pO_2 drop below 10 mm Hg	2 (3.3)
Total	162				7 (4.3)

Table 1	Summary of studies	identified from a Pl	RISMA-quided search of	transcutaneous gas	monitoring in	free flaps
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Abbreviations: pCO_2 , carbon dioxide partial pressure; pO_2 , oxygen partial pressure. ^aFree as well as pedicled flaps.

failing flaps.¹³⁻¹⁶ Moreover, some have noted that measurement accuracy is highly dependent on probe placement and user expertise^{1,12,15,17-19} and the device may be difficult to use for continuous monitoring.^{1,20} Because normally very few oxygen molecules reach the skin surface by diffusion, this method may require inducing local hyperemia by heating the skin and increasing the amount of oxygen that can be measured. This is more cumbersome and may place the patient at risk for burns with long-term monitoring.

Other than the Clark electrode, luminescence lifetime imaging (LLI), and luminescence ratiometric oxygen imaging (LROI) can also provide a direct measure of tissue oxygen by using a concept of oxygen-dependent fluorescent decay. LLI was found to be better able to produce accurate readings than the Clark electrode, as well as more responsive to micro-circulatory changes.⁸²¹ LROI could also differentiate between healthy and failing flaps, but its accuracy is dependent on a long acquisition period, examiner skill, and experience with the device. Furthermore, the probe cannot reach less accessible flaps such as intraoral and noncutaneous flaps.^{22,23} While the monitoring depth of LROI can be increased by performing

measurement alongside contrast-enhanced ultrasound (CEUS) to monitor noncutaneous flaps and flap portions, CEUS is a more invasive form of monitoring requiring the intravenous administration of contrast.²⁴

Since then, other measurement modalities have been developed to monitor tissue oxygen. Notably, near-infrared spectroscopy (NIRS) evaluates the relative concentrations of oxygenated and deoxygenated hemoglobin and oxidized cytochrome aa3 as an indicator of tissue oxygen supply and intracellular oxygen availability. Irwin et al first investigated the use of NIRS to monitor flap viability and found numerous advantages over the polarographic Clark-type electrode.²⁵ NIRS can monitor deeper depths, possibly even buried flaps, and is immutable to seroma or hematoma formation.^{6,26,27} Although measurement accuracy can be affected by probe movement, the movement must be dramatic. However, NIRS technology cannot provide absolute values and numerous other investigators have found relative trend or index values were better predictors of vascular compromise.^{25,28-31} Stranc et al confirmed these findings in a rat model by using an O₂ saturation index that increased the

Technology	Mechanism	Advantages	Disadvantages
Handheld Doppler US	Detects sound proportional to flap blood flow	 Inexpensive Easy to use Noninvasive Rapid assessment With experience, can differentiate venous from arterial signal 	 Operator dependent Not quantitative Intermittent monitoring Cannot surveil buried flaps
Color duplex US	Visualizes blood flow using US	 Relatively inexpensive Noninvasive Rapid assessment Can differentiate venous from arterial blood flow 	 Operator dependent Not quantitative Intermittent monitoring Typically requires technician Cannot surveil buried flaps
Implantable Doppler probe	Detects sound proportional to flap blood flow, near anastomosis on pedicle	 Relatively easy to use Continuous monitoring Rapid assessment Can monitor buried flaps 	 Has to be placed intraoperatively Not quantitative High cost
Flow coupler	Implantable Doppler incorporated into venous coupler	Same as "implantable Doppler probe"	 Same as "implantable Doppler probe"
Laser Doppler flowmetry	Blood flow velocity is estimated by illumi- nating a tissue sample with laser light and then analyzing the frequency distribution of the backscattered light	 Continuous monitoring Semiquantitative Very accurate Rapid assessment 	 Operator dependent Typically requires technician Cannot surveil buried flaps Very high cost
NIRS or VLS	Light of specific wavelength is emitted toward tissue and changes in intensity attributable to relative changes in hemo- globin concentrations, which correlate with tissue oxygenation	Continuous monitoringQuantitativeRapid assessment	 Operator dependent Cannot surveil buried flaps Extremely high cost
Microdialysis	Catheter is placed into tissue, sample extracted and presence of metabolites like lactate analyzed	 Quantitative Accurate Used in buried flaps	 Sample variability Intermittent monitoring Invasive Not real time Requires technician High cost
SFDI	Near infrared light projected and col- lected in real time, converted into spatial maps of O ₂ Hb and Hb concentrations	 Real time Quantitative Continuous Can distinguish arterial from venous complications 	High costCannot surveil buried flaps
LSI	Laser light reflected into video camera produces speckles that correlate with tissue blood flow	 Continuous Real time Noninvasive Can distinguish venous from arterial complications 	Cannot surveil buried flaps
Oxygen-sensing bandage	Molecule within paint detected by camera creates 2-map of oxygenation	 Real time Noninvasive Can be applied to high topography regions Low cost 	Intermittent monitoringNot for acute useSample variability
Thermal imaging	Temperature of skin measured by infrared sensor on handheld device	 Real time Noninvasive Low cost	 Operator dependent Intermittent monitoring Only useful for flaps with cutaneous island

 Table 2
 Overview of free flap monitoring technologies

Abbreviations: Hb, hemoglobin; LSI, laser speckle imaging, NIRS, near-infrared spectroscopy; SFDI, spatial frequency domain imaging; US, ultrasound; VLS, visible light spectroscopy.

sensitivity of detecting tissue hypoxia and correlated with necrosis onset.³² They found that using this index gave highly reproducible results allowing for early prediction of flap survival.³² In the clinical setting, many studies demonstrated that NIRS technology could not only predict flap compromise before clinical manifestation without false positives or negatives, but could also detect and distinguish between arterial and venous occlusion.^{6,27,29,33-35} In addition, NIRS monitoring was found to have greater sensitivity and specificity than clinical examination alone²⁷ and led to lower flap failure rates for one investigator.³⁴ Moreover, it may decrease the amount of time spent in postoperative recovery, the length of intensive care unit (ICU) stays, and the amount of nursing care required for monitoring.³³ Another advantage of NIRS technology include ease-of-use requiring minimal training.^{6,33,34,36}

Flaws with the NIRS monitoring system have also been reported. Contrary to earlier reports, Scheufler et al found hematoma formation affected the accuracy of NIRS monitoring.28,35 In addition, they found nonviable tissue may still be perfused even after prolonged ischemia.28 Serosanguinous buildup, systemic oxygen saturation, and systemic blood pressure variations may affect measurement accuracy and careful interpretation of measurements is needed.³⁷ Salgarello et al described a negative relationship between regional oxygen saturation and body mass index, as well as flap size, and suggested the current NIRS thresholds may need to be reevaluated in patients with high body fat content.³⁸ Perhaps the most prohibitive flaw to NIRS monitoring is expense as reported costs range from \$8,000 to 50,000 per unit.^{3,33,34,36} It has yet to bet determined whether the hospital cost savings can offset the expense associated with currently available NIRS flap monitoring technology.33

To offset these flaws, some have proposed concurrent use of LDF, even though it has been found to be inferior to NIRS.^{28,35} In response, a system combining the two technologies was developed, which Hölzle et al used with some success.^{39,40} They found that the combined NIRS/LDF detected all vascular complications with no false positives or negatives and was able to establish a threshold indicating vascular compromise for different flap types. In addition to detecting impending flap failure before clinical signs were present, the combined modalities prevented unnecessary return to the operating room when clinical signs would suggest otherwise. However, continuous monitoring was difficult, and the system was unable to survey buried flaps. Other proposed improvements to measuring tissue oxygenation, such as broadband measurements to detect cytochrome aa3 and Erlangen microlightguide spectrophotometry, failed to show significant superiority to existing technology.^{41,42} However, Olenczak et al noted that a more reliable measurement of cytochrome aa3 may be a better indicator of flap compromise than measures of hemoglobin oxygenation and concentration.41

One the newer technologies for flap monitoring involving transcutaneous oxygen monitoring is the oxygen-sensing paint-on bandage. The oxygen-sensing molecule embedded in the bandage allows for smart-phone cameras to acquire images that can be converted into a two-dimensional map of tissue oxygenation. Early studies indicate the oxygen-sensing bandage is comparable to NIRS in detecting tissue oxygenation during vascular compromise.⁴³ Despite ease of use, this technology remains costly and not widely available.

Other modalities of note include spatial frequency domain imaging (SFDI), visible light spectroscopy (VLS), laser speckle imaging (LSI), and thermal imaging. SFDI provides similar measurements as NIRS but surveys a larger area of interest and does not require cutaneous contact. While it could reliably distinguish between arterial and venous occlusion with low variability in measurements, it may be difficult to use as it requires correct calibration and relies on patients remaining still for a long period of time. In addition, results may be confounded by thinner flaps, room illumination, and skin pigmentation.44,45 While SFDI has demonstrated potential usefulness for intraoperative planning, it has not yet been evaluated for postoperative flap monitoring in human subjects. Visible light spectroscopy also allows similar measurements and may be more sensitive to smaller changes in tissue oxygenation.^{11,46-48} It is more applicable for smaller flaps, such as those used in the upper extremity, including digital replantation.¹¹ Kagaya et al found that NIRS was better for evaluating fat perfusion or the perfusion of buried flaps, while visible light spectroscopy is more suited to thinner flaps.⁴⁶ Despite promising results, high interobserver variation precluded establishment of a threshold for flap failure.47 Laser speckle imaging involves capturing reflected laser light of a tissue as speckles on a video-camera detector array which correlates with blood flow as the speckles move in relation to the red blood cells moving.⁴⁹ This technology is fast, easy to use, noninvasive, and may be a valuable adjunct for use intra- and postoperatively for evaluating tissue perfusion of free flaps.⁷ Another relatively new technology for monitoring free flaps is use of thermal imaging cameras. Cruz-Segura et al found that a difference of more than 2°C between flap and surrounding skin as detected by thermal imaging may allow for diagnosis of flap vascular compromise up to 12 hours earlier than clinical examination.⁵⁰ Of note, Ricci et al combined the transcutaneous pO₂ monitoring with contemporary communications modalities to notify the surgeon of flap compromise via text message alert to expedite surgical intervention as needed, which highlights how modern technology can continue to improve flap care.

Transcutaneous Carbon Dioxide Monitoring

Minimal work has been done regarding clinical efficacy evaluation of transcutaneous carbon dioxide partial pressure $(pCO_2 \text{ or } ptcO_2)$ measurement in flap monitoring. Early uses of this technology include detection of skin necrosis in necrotizing fasciitis, bullous pemphigoid arteriosclerosis obliterans, distal ischemia of canine pedicle flaps, and pressure ulcer monitoring.⁵¹⁻⁵⁵ One modality recently developed involves a bifunctional probe taped to the skin and which measures both transcutaneous $ptcO_2$ and $ptcCO_2$.¹⁹ While this technology is at least semiquantitative and can detect total venous occlusion, it does not appear sensitive enough to detect partial

venous occlusion. Other drawbacks of current technology include the need for substantial training in calibration and placement of the probe for accurate measurement.¹⁹ Similar to *ptcO*₂ monitoring, *ptcCO*₂ monitoring may ideally be conducted at supraphysiologic temperatures (~44°C) to increase the gas permeability of skin.¹⁴ However, other studies argue that flaps can be monitored effectively at 37°C.²⁰ Without having to heat the skin, the probe does not require constant repositioning to spare the underlying skin, thereby improving the accuracy of *ptc*CO₂ measurements.²⁰ The ability to attach the probe for long durations raises the possibility of ptcO₂ and ptcCO₂ continuous flap monitoring.^{20,56-59} Combined ptcO₂ and ptcCO₂ monitoring has also shown promise in selecting the number and duration of leech therapy.⁶⁰ While *ptcO*₂ and *ptcCO*₂ have been measured together, each has its own advantages and disadvantages. Measurements of *ptcO*₂, respond more rapidly to occlusion than *ptcCO*₂, but *ptc*-CO₂ is influenced far less by external factors than *ptc*O₂.¹⁴

Future Directions

Flap monitoring is a necessary part of postoperative care following free tissue transfer. In addition to a conventional flap clinical examination, which relies on an experienced clinician and is done intermittently, there is a critical need to develop a method or device that can provide accurate, reliable, rapid, easy-to-use, and continuous assessment of flap perfusion. As discussed earlier, techniques that leverage measurements of $ptcO_2$ and $ptcCO_2$ measurements may be the most promising as tissue oximetry is associated with higher salvage rates compared with clinical examination alone. Transcutaneous gas measurements in flaps are not susceptible to core temperature, blood pressure, or pulse rates. If these technologies can be reduced in cost and made more widely available, it may result in systematic improved protocol for flap salvage.

With free flap reconstruction currently being a generally uncommon procedure and flap loss rates generally less than 5%, cost may be the most critical barrier for most institutions investing in newer, promising technology. If a device could be designed and manufactured for much less cost, more microsurgical teams would have access to it, leading to more use and ultimately data collection, more technological refinements to optimize its utility, and thus more universal adoption of a newer adjunct technique for flap monitoring. Some of the newer technologies, such as thermal imaging and oxygen-sensing bandages, require little more than a digital camera or smartphone add on which could prove invaluable for widely distributing these devices. More work needs to be done to reduce the cost of acquiring high fidelity ptcO₂ and ptcCO₂ data as an adjunct to the clinical examination to potentially improve free flap salvage rates.

Conflict of Interest

None declared.

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