TRANSCUTANEOUS PARTIAL OXYGEN TENSION AND SKIN BLOOD FLOW MONITORING: CONTINUOUS, NONINVASIVE MEASURES OF CARDIORESPIRATORY CHANGE *

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Transcutaneous partial oxygen tension (tc pO_2) and skin blood flow (via heat clearance) were measured noninvasively in 22 male subjects who performed stress-inducing tasks (i.e. hand-grip exercise, cold pressor test, breath holding, hyperventilation and mirror-tracing). An analysis of variance and covariance was conducted for tc pO_2 , heat clearance, heart rate, respiration rate, finger pulse volume and systolic/diastolic blood pressure. Results indicate that tc pO_2 can depict phasic cardiorespiratory challenge. Heat clearance proved to be less sensitive to the demand conditions used here. An analysis of covariance revealed a negative correlation between tc pO_2 and blood pressure and respiration rate, as well as a positive correlation between tc pO_2 and heat clearance. This suggests that phasic changes in tc pO_2 are induced by both peripheral vasomotor activity and changes in arterial pO_2 .

1. Introduction

Cardiovascular psychophysiology is concerned with the physiological responses and response patterns elicited in human subjects during stress-inducing challenge (Ax, 1953; Fahrenberg, Walschburger, Foerster, Myrtek and Müller, 1983; Lacey, 1967). Since most experimental paradigms require healthy subjects to perform various tasks, physiological parameters should be measured noninvasively, because invasive measures could impair the subject's ability to respond and may entail certain risks. This limits researchers to a modest number of physiological measurements.

Though an important indicator of cardiorespiratory functioning, blood gas analysis has not been systematically conducted by psychophysiologists, be-

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cause it requires invasive methods. In addition, the procedure of blood taking is a challenge in and of itself. Such a measurement, therefore, may confound any effect due to experimental manipulation.

Polarographic measurement of transcutaneous pO_2

A method to estimate partial arterial oxygen pressure (a pO₂) through the skin, termed transcutaneous partial oxygen tension (tc pO₂), has been developed by Lübbers and his co-workers (Huch, Lübbers and Huch, 1972, Lübbers, 1981). Based on the polarographic measurement of oxygen in a liquid via the Clark method (Clark, 1956), a pO₂ can be estimated at the skin surface. To do this, the skin surface must first be heated to 44°C so that a state of blood flow independency is reached. The resulting hyperaemia provides the dermis with a rich amount of arterialized blood, thereby saturating the oxygen needs of the skin. The polarographic sensor has three platinum cathodes that are submerged in an electrolyte solution. Oxygen passes through a telfon-cellophane membrane covering the transducer and changes the potential of these cathodes.

To monitor skin blood flow, the amount of heat clearance can be directly measured by recording the electrical current needed to maintain a fixed temperature at the electrode. Since the temperature of the electrode is greater than that of blood, blood flow dissipates the heat of the electrode. It follows that the greater the blood flow the more current will be needed to maintain the electrode's temperature. Thus, a second measure is produced by the same electrode, namely, skin blood flow via heat clearance.

Cardiovascular physiologists and anesthesiologists have been concerned with tc pO_2 as a correlate of a pO_2 . Lübbers (1981) has demonstrated that under controlled conditions, tc pO_2 can reflect arterial pO_2 if blood flow independency is achieved. Huch, Huch and Lübbers (1982) monitored tc pO_2 and intra-arterial a pO_2 in subjects given 95% O_2 and 5% CO_2 . Their subjects further performed hyperventilation and breath holding. The correspondence between the resulting tc pO_2 and a pO_2 records is quite impressive, but correlation coefficients were not reported.

The present study has been conducted to explore the value of transcutaneous partial oxygen tension (tc pO_2) and skin blood flow monitoring in subjects exposed to typical laboratory challenge. These noninvasive microcirculatory measures should prove to be interesting variables for psychophysiologists because changes in tc pO_2 reflect changes both in arterial pO_2 and in skin blood flow due to vasoconstriction. As such, tc pO_2 and heat clearance give an index of cardiorespiratory functioning.

If to pO_2 and skin blood flow measures are to be useful, they must first be tested to see whether they are sensitive enough to reflect changes induced by demanding tasks. Therefore, we conducted measurements over a number of experimental conditions to first assess the sensitivity of to pO_2 and heat

clearance to various types of cardiorespiratory challenge. Significant differences from resting levels should be evident during adequate challenge. We also compared to pO_2 and heat clearance values with those of traditional variables used in cardiovascular psychophysiology.

2. Method

2.1. Subjects

Male student volunteers (N = 22, mean age, 26 yrs, SD = 2.3) free of any obvious illness participated in the experiment which lasted about two hours. They either received credit fulfilling requirements for their studies in psychology or a fee (\$8.00 US). Subjects were asked to abstain from caffeine or alcoholic beverages and tobacco for a three-hour period prior to examination. All subjects were right-handed.

2.2. Apparatus

We used the Oxymonitor (Hellige GmbH, Freiburg) to measure to pO_2 and heat clearance. The oxygen sensor was calibrated to room air with reference to local barometric pressure. Following ten minutes of stable values, the sensor was positioned on the skin surface of the inside of the left forearm. A contact agent (Hellige, GmbH) and proper adhesive rings assured a direct interface between the skin and electrode. The electrode then hyperaemized the cutaneous tissue within a 15 min period.

The following parameters were also measured continuously during all rest and challenge phases: Heart rate, respiration rate (3 ECG chest leads), finger pulse volume (amplitude recorded on the left index finger using pneumatic plethysmography without venous occlusion, Boucker pressure transducer). Data were recorded on an 8-channel polygraph and analyzed off-line with a sampling rate of 20/min. Data were reduced to mean values for each subject on each measure in the given condition. These mean values were then used in the analyses of variance and covariance.

Additionally, blood pressure was measured using a manometer on the right arm directly following each challenge phase (except between hyperventilation and breath holding). A cuff was manually inflated to 170 mmHg and a microphone positioned over the antecubital artery picked up the Korokoff sounds.

2.3. Procedures

2.3.1. Electrode placement and calibration

Subjects were asked to enter the air-conditioned chamber (25 ± 1 °C, relative humidity 55%) and were seated in a comfortable chair. Instructions were given

concerning the nature of the tasks and the physiological measures to be made. After subjects gave their consent, electrodes were positioned and calibrated. Adequate time (15 min) was then given to allow for hyperaemization of the skin area beneath the electrode for the tc pO_2 recording. Subsequently, a five-minute baseline recording was conducted.

2.3.2. Cardiorespiratory challenge

First, various physical challenges were induced to test cardiorespiratory response to standardized stimulation. These were: (1) Ergometric exercise (self-paced hand-grip exercises lasting 2 min); (2) cold pressor (left forearm submerged in 4°C water which was stirred by a magnetically coupled propeller to prevent a warm layer from forming around the arm's surface, 1 min); (3) self-paced hyperventilation (1 min); (4) breath holding (at inspiratory level for as long as possible; maximal duration of 2 min). Following each challenge, subjects were allowed a 2-min rest period similar to baseline recording. The sequence of conditions was kept constant to allow analysis of between-subjects variance (see Data Analysis). Sufficient time for recovery minimized any carry-over effects.

2.3.3. Sensorimotor tasks

Subsequent to these challenges, subjects were instructed as to the nature of the sensori-motor test. It consisted of four mirror-tracing tasks of increasing difficulty. We used the Sensorimotor Training instrument (Zak Company, Munich) together with a tilt mirror and occluder. During tracing, whenever subjects traced outside of the pattern an error was signalled. The error was recorded and simultaneously a tone was produced to inform the subject of the error. To increase the competitive nature of the tracing tasks, performance norms, based on preliminary testing, were presented. Subsequent to performance, subjects were informed about their scores, which they then could compare to these norms. Tracing tasks were employed because they require concentration and do not require verbal responses, which would interrupt breathing and, thus, induce artifactual changes in to pO_2 .

2.4. Data analysis

2.4.1. Analysis of variance

An analysis of variance was used to test the main effects 'conditions' and 'subjects'. If to pO_2 and heat clearance are sensitive measures, then challenges eliciting changes in cardiorespiratory functioning should produce significant changes in these variables, thus yielding a significant main effect for 'conditions'. If, however, to pO_2 and heat clearance values exhibit considerable individual differences, then the main effect 'subjects' should be significant.

The amount of variance accounted for by these terms was used to indicate the degree to which these variables were comparable to traditional measures. In addition, the error term was assessed as a measure of reliability (a low error term corresponds to good reliability). The Scheffé test (Scheffé, 1953) is employed as a conservative test of significant pair-wise differences between initial rest and strain values.

2.4.2. Analysis of covariance

An analysis of covariance (see, Fahrenberg and Foerster, 1982) was conducted to assess the degree to which to pO_2 and heat clearance covary with standard measures of cardiorespiratory function during rest and challenge. The between-subjects coefficients were based on the reduced values on each variable averaged over all rest and challenge conditions for each subject separately. The resulting coefficients provided an index of covariance between variables after the differential effect of the various conditions had been partialled out by averaging each subject's value over the ten conditions. These coefficients were submitted to normal tests of significance with df = 21.

3. Results

Fig. 1 presents the mean values and standard errors across subjects for each physiological measure during the ten experimental conditions. As can be seen in fig. 1, to pO_2 increases during hand-grip exercise, hyperventilation and tracing tasks, whereas breath holding induces decreases. Heat clearance, on the other hand, fails to demonstrate significant changes from initial baseline values.

Table 1 presents the results of the analysis of variance for the main effects 'conditions' and 'subjects'. The columns (from left to right) present the sources of variance, the sums of squares, the degrees of freedom, the mean square, the F ratio, the significance of the F ratio, and the amount of variance accounted for by the term in question. In addition, the results of the Scheffé test for significant differences from initial resting values are given in the final column.

As can be seen in table 1, the F ratios for the main effects 'conditions' and 'subjects' are highly significant for most variables. The exceptions are the main effect 'conditions' for heat clearance (p=0.038) and for diastolic blood pressure (p=0.137; ns). Of more interest is the amount of variance accounted for by either of the main effects. For tc pO₂, the main effect 'subjects' accounts for 60% of the total variance, whereas the main effect 'conditions' accounts for only 12.8%. For heat clearance, 'subjects' accounts for 84.1% and 'conditions' merely for 1.8% of the total variance. Compared to heart rate (46.1% and 28.2% for 'subjects' and 'conditions', respectively), tc pO₂ and heat clearance are less sensitive to phasic changes induced by the challenges used here.

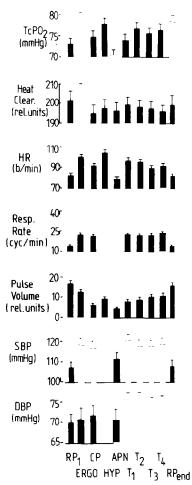


Fig. 1. Mean values and standard errors across subjects on physiological measures recorded during each experimental condition. Abbreviations: RP1 = rest period 1; ERGO = ergometric exercise; CP = cold pressor test; HYP = hyperventilation; APN = breath holding; T1-T4 = tracing tasks 1-4; RPend = final rest period; HR = heart rate; SBP = systolic blood pressure; DBP = diastolic blood pressure.

The results of the Scheffé tests further support these findings. Whereas to pO_2 values recorded during ergometric exercise and during the final rest phase significantly differ from initial rest values, heat clearance values recorded during the various challenges and final resting do not significantly differ from initial values.

Table 2 depicts between-subjects correlation coefficients for each physiological measure. These coefficients are based on each subject's averaged values

squares, the F-ratio, and the significance level of the F-ratio (p). The final two columns give the percent of variance accounted for by the term in question, and the results of the Scheffe tests. For the latter, only the experimental conditions are listed that yielded values that significantly differed Results of an analysis of variance for the main effects 'subjects' and 'conditions'; the first column (left) presents the source of variance for the variable in question; the second column the sums of squares, and the next column the degrees of freedom (df). In the next three columns are shown the mean from initial rest values

	Analysis of variance	e e					
Source	Sums of squares	đť	Mean squares	F	Ь	Percent of variance	Scheffé test
Variable: tc pO ₂							
Total	_	216	54.44			100.00	Ergometry
Subjects	7356.37	21	350.30	22.33	0.000	60.05	Final rest
Conditions	1484.25	6	164.92	10.51	0.000	12.17	
Error	2918.38	186	15.69			27.78	
Variable: Heat c	learance						
Total 82 392.	82 392.75	216	381.45			100.00	
Subjects	70085.75	21	3337.42	89.68	0.000	84.06	
Conditions	1904.75	6	211.64	3.78	0.038	1.81	
Error	10 402.25	186	55.93			14.13	
Variable: Heart 1	rate						
Total 4	47.755.77	216	221.09			100.00	Ergometry
Subjects	23 367.82	21	1112.75	18.67	0.000	46.08	Hyperventilation
Conditions	13 304.28	6	1478.25	24.81	0.000	28.21	•
Error	11 083.68	186	59.59			25.71	

Ergometry	Hyperventilation	Breath holding	Tracing tasks 1, 4		Cold pressor	Hyperventilation	Tracing tasks 1-4			Ergometry	Hyperventilation	Tracing tasks 1, 2, 4			Hyperventilation			
100.00	7.94	73.61	18.45		100.00	16.78	29.14	54.08		100.00	68.11	8.27	23.62		100,00	75.89	2.77	21.34
,	0.000	0.000				0.000	0.000				0.000	0.000				0.000	0.137	
, !	5.14	87.55				4.06	12.69				25.77	7.62				31.55	3.50	
87.85	71.16	1532.53	17.50		47.35	107.72	336.78	26.54		249.66	1641.31	485.24	63.70		133.36	966.34	105.62	30.63
216	21	6	186		216	21	6	186		188	20	œ	160		188	20	00	160
ation rate 18975.77	1927.25	13 792.80	3255.73	pulse volume	10228.50	2262.02	3031.0	4935.48	c blood pressure	46 899.31	32 826.18	3881.88	10191.24	dic blood pressure	25072.14	19326.81	845.00	Error 4900.34
Variable: Respiration rate Total 18975	Subjects	Conditions	Error	Variable: Finger pulse volume	Total	Subjects	Conditions	Error	Variable: Systolie	Total 46899.31	Subjects	Conditions	Error	Variable: Diasto	Total	Subjects	Conditions	Error

a) To compensate for missing data (comprising about 1% of all data), three degrees of freedom were subtracted from the total number. Blood pressure values are based on g measurements per subject since no measurement was conducted between hyperventilation and breath holding.

Table 2 Between-subjects covariance coefficients are presented for the physiological measures recorded. These are based on values averaged for each subject over the ten experimental conditions; df = 21. Significance levels: r = 0.41, p = 0.05; r = 0.54, p = 0.01

	1	2	3	4	5	6	7
(1) Tc pO ₂							
(2) Heat clearance	0.37						
(3) Heart rate	-0.35	-0.09					
(4) Respiration	-0.39	-0.13	0.25				
(5) Pulse volume	0.00	-0.02	-0.05	-0.31			
(6) Systolic BP	-0.40	-0.30	0.46	0.13	0.09		
(7) Diastolic BP	-0.42	-0.29	0.56	0.04	0.17	0.74	

from the 10 experimental conditions. These coefficients reflect the degree to which between-subject differences covary after eliminating the various effects of the demanding tasks by averaging.

As can be seen in table 2, covariance between physiological measures is generally modest to low. The main exceptions are found between to pO_2 and heat clearance (0.37), to pO_2 and respiration rate (-0.39), to pO_2 and systolic blood pressure (-0.40), to pO_2 and diastolic blood pressure (-0.42) and systolic/diastolic blood pressure and heart rate (0.46 and 0.56, respectively). Note that heat clearance only covaries well with to pO_2 and very little with heart rate (-0.09).

4. Discussion

The results of the analysis of variance indicates that to pO_2 is sensitive enough to reflect changes induced by cardiorespiratory challenge. Heat clearance fails, however, to satisfactorily depict changes in skin blood flow. In addition, the analysis of variance shows that the 'subjects' term accounts for most of the variance for to pO_2 and heat clearance.

It may be argued that the 'subjects' term consists of both true individual differences and between-subjects measurement error due to differences in transducer placement, etc. However, if the variance accounted for by 'subjects' is due primarily to error, a significant correlation between tc pO_2 and diastolic blood pressure (-0.42) would not be expected. As this coefficient is based on the subjects' average value over all conditions, the 'conditions' variance is partialled out, thus leaving only the 'subjects' and error terms. Error variance does not covary between independent measures, which indicates that the 'subjects' term does consist of meaningful variance.

To further substantiate between-subjects differences, a replication with the same subject sample using identical conditions would, in some respects, be

preferable, but not completely unproblematic. Experiencing the same challenges for a second time, subjects would surely respond differently. Therefore, responses would vary depending on how well subjects habituate to the conditions.

In summary, the present findings suggest that to pO_2 can depict changes elicited by cardiorespiratory challenge. Heat clearance is not, however, sensitive enough to depict changes in skin blood flow induced by the challenges administered here. The significant amount of variance accounted for by the 'subjects' term and the significant correlation with diastolic blood pressure indicate that tonic to pO_2 values can reflect individual differences in cardiorespiratory functioning.

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References

Ax, A.F. (1953). The physiological differentiation between fear and anger in humans. Psychosomatic Medicine, 15, 433-442.

Clark, L.C. Jr. (1956). Monitor and control of blood tissue oxygen tensions. Transactions American Society of Artificial Organs, 2, 41–48.

Fahrenberg, J. and Foerster, F. (1982). Covariation and consistency of activation parameters. Biological Psychology, 15, 151-169.

Fahrenberg, J., Walschburger, P., Foerster, F., Myrtek, M. and Müller, W. (1983). An evaluation of trait, state and reaction aspects of activation processes. Psychophysiology, 20, 188–195.

Huch, R., Huch, A. and Lübbers, D.W. (1982). Transcutaneous pO₂. Thieme: Stuttgart.

Huch, R., Lübbers, D.W. and Huch, A. (1972). Quantitative continuous measurement of partial oxygen pressure on the skin of adults and newborn babies. Pflügers Archive, 337, 185–198.

Lacey, J.I. (1967). Somatic response patterns and stress: Some revisions of activation theory. In: Appley, M.H. and Trumbull, R. (Eds.). Psychological Stress: Issues in Research. Appleton-Century-Crofts: New York.

Lübbers, D.W. (1981). Theoretical basis of transcutaneous blood gas measurement. Critical Care Medicine, 9, 721-733.

Scheffé, H. (1953). A method for judging all contrasts in the analysis of variance. Biometrika, 40, 87-104.