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Paulo Jorge Oliveira dos Santos
Effect of a swimming training session on the exhaled breath
temperature

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Effect of a swimming training session on the exhaled breath
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Área: Imunologia

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Dra Mariana Couto

E sob a Coorientação de:

Professor Doutor André Moreira

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Effect of a swimming training session on the exhaled breath temperature

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1 **Effect of a swimming training session on the exhaled breath**
2 **temperature**

3
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29 **Effect of a swimming training session on the exhaled breath**
30 **temperature**

31 **Pediatr Allergy Immunol**

32 **ABSTRACT**

33 **Background:** By exercising, an inflammatory response of the airways occurs, with
34 bronchial smooth muscle constriction and vasodilatation. The later could lead to an
35 increase in exhaled breath temperature (EBT), possibly more pronounced in subjects
36 with asthma. We aim to investigate the effect of a training session on EBT of elite
37 swimmers and to assess if the impact is different in asthmatic vs non-asthmatic
38 swimmers.

39 **Methods:** Swimmers that are annually screened for asthma in our Department were
40 invited to this prospective study. The regular screening includes 2 visits in which
41 subjects perform skin prick tests, spirometry before and after salbutamol inhalation and
42 bronchial challenge with methacholine. Diagnosis of asthma was according to IOC-MC
43 criteria. For those who agreed to participate, EBT was measured with X-halo
44 thermometer before and after a training session. SPSS ($p < 0.05$) was used to compare
45 baseline and post-training EBT; and Δ EBT between asthmatic and non-asthmatic
46 swimmers.

47 **Results:** 22 swimmers accepted to participate, of which 10 had asthma. EBT
48 significantly increased after the training session ($p = 0.020$). No significant differences
49 were observed in mean Δ EBT among asthmatic and non-asthmatic swimmers
50 ($p = 0.222$).

51 **Conclusions:** EBT increased after a training session. There was no difference in EBT
52 between asthmatics and non-asthmatics.

53 **Key words:** asthmatics, exhaled breath temperature, swimmers, training

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63 INTRODUCTION

64 Asthma is defined as a clinical syndrome of intermittent respiratory symptoms triggered
65 by viral infections, environmental allergens, or other stimuli, and is characterized by
66 nonspecific airway hyperresponsiveness and inflammation (1). Vasodilatation is a
67 critical feature of inflammation, and angiogenesis and vascular remodeling are features
68 of chronic inflammatory diseases, such as asthma (2). The increased vascularity of the
69 airways in asthma is partly due to the elevated number of vessels associated with
70 angiogenesis (3) and partly due to vasodilation caused by the release of vasodilator
71 mediators, such as histamine, bradykinin and nitric oxide (NO).

72 As part of this inflammatory process, the increased vascularization of the airway
73 mucosa that occurs in asthmatics, leads to increased heat exchange during expiration
74 (4) and previous studies have reported that patients with asthma present an elevated
75 exhaled breath temperature (EBT) compared to healthy controls (5). The elevation of
76 exhaled NO as an inflammatory marker seen in asthmatics, has previously been shown
77 to correlate with increased EBT (5, 6), and therefore EBT has been proposed as a non-
78 invasive new biomarker for asthma control (7, 8).

79 Although the pathogenesis of the association of sports practice and airway injury is not
80 fully elucidated, some hypotheses have been proposed, one of which focuses upon
81 cooling of the airways caused by hyperpnea during exercise. Vigorous exercise
82 requires an increased ventilatory rate to meet higher muscular oxygen needs, which
83 results in the inhalation of a large volume of relatively cold and dry air and the loss of
84 heat from the respiratory mucosa. This mechanic noxious stimulus could cause
85 epithelial damage, and therefore the influx of inflammatory cells and their mediators
86 release (9-11). Also, higher levels of airway vascular permeability have been shown to
87 be a good predictor of the severity of EIA in asthmatics, which has led to the
88 microvascular theory of EIA based on functional abnormalities of endothelial cells in
89 newly generated microvessels in asthmatic airways (12).

90 It has been shown a significant increase in EBT in asthmatic children after exercise
91 (13), but this increase is not different between asthmatics and controls (14). In athletes
92 this was never investigated before and is a relevant issue as they present an increased
93 risk for asthma, especially those who take part in endurance sports, such as swimming
94 (10). We hypothesize that exercise would increase EBT, and a more pronounced
95 increase in would be seen in asthmatic athletes as a greater degree of response to
96 exercise as compared to healthy ones. Therefore, our aim was to investigate the effect

97 of a training session on EBT of elite swimmers, and to assess if the impact is different
98 in asthmatic vs non-asthmatic swimmers.

99

100 **METHODS**

101 **Study design and subjects**

102 Elite swimmers of the FC Porto main swimming team who are annually screened for
103 asthma and atopy at Allergy, Asthma and Sports Unit were invited to participate in this
104 prospective study. The regular screening includes two visits, about 1 week apart, in
105 which subjects are evaluated with skin prick tests, lung volumes before and after
106 salbutamol inhalation, and airway responsiveness to methacholine. Swimmers are
107 asked to withheld anti-asthmatic or anti-allergic medication that they might be
108 possibly taking for both visits, according to European Respiratory Society guidelines
109 (15). Inhaled short acting β 2-agonists were withheld for 8 hours before testing; inhaled
110 long-acting β 2-agonists, theophylline, and leukotriene antagonists were withheld for the
111 last 72 hours; antihistamines were withheld for the last 7 days; and both inhaled and
112 orally administered corticosteroids were withheld for the last month. At that time, they
113 are diagnosed as having or not asthma, according to International Olympic Committee
114 criteria (16). Later, for those who agreed to participate in the study, EBT was collected
115 before and after a training session at their swimming pool.

116 In order to be eligible to participate in this study, a subject had to meet all the following
117 criteria: elite level swimmer; free from respiratory infection in the last 3 weeks; provided
118 signed and dated informed consent. A potential subject who met any of the following
119 criteria was excluded from participation in this study: pregnancy; recent episode of
120 hemoptysis; forced expiratory volume in the first second (FEV₁) lower than 60% of the
121 predicted value or 1.5 L; neurological or psychiatric illness; lack of collaboration or
122 coexistence of diseases that limit the patient's ability to carry out the tests; recent
123 stroke or heart attack.

124 All subjects gave written informed consent, and the study was approved by ethical
125 commission of Centro Hospitalar São João / Faculdade de Medicina da Universidade
126 do Porto.

127 **Procedures**

128 *Methacholine challenge*

129 Non-specific bronchial hyperresponsiveness was measured by methacholine challenge,
130 according to guidelines (15). Methacholine was delivered by inspiration triggered by an
131 automatic dosimeter that delivers a single dose soon after the onset of a deep breath.
132 The five-breath dosimeter method was used and the provocative dose causing a 20%
133 fall in FEV₁ (PD₂₀) was determined.

134 *Lung function*

135 Spirometry was carried out according to the American Thoracic Society criteria (17).
136 Lung function measurements were repeated 15 minutes after 400 µg of salbutamol in
137 aerochamber to assess reversibility.

138 *Skin prick tests*

139 Skin prick tests were carried out in accordance international guidelines (18) with a
140 standard battery of commercial extracts for common aeroallergens (Leti®, Madrid,
141 Spain). Histamine dihydrochloride and diluent were used as positive and negative
142 controls, respectively. Testing solutions were stored at +2 to +8°C when not in use. The
143 largest and perpendicular diameter of the wheal for each of the allergens is measured
144 and the following value calculated: largest + perpendicular diameter/2. A subject is
145 defined as atopic in the presence of at least one positive result (regarded if the value
146 calculated was ≥3 mm and controls showed adequate reactions) (18).

147 *Exhaled breath temperature*

148 EBT was measure using an X-halo device (Delmedica Investments, Singapore), 5
149 minutes before (baseline EBT) and 5 minutes after (post-exercise EBT) the swimming
150 training session, according to previously validated methods (20). Briefly, the swimmers
151 were requested to inhale freely through the nose and to exhale into the device at a rate
152 and depth typical of their normal tidal breathing rhythm. The maneuver was continued
153 until the built-in software of the instrument indicated that the measured value was
154 stable. The decision of collecting EBT 5 minutes after the exercise was based in
155 previous studies that have shown that this was the time point in which EBT reached the
156 highest values, and decreases thereafter (14).

157 The time needed to achieve the stable EBT was recorded. Before and between
158 measures, the device was kept at room temperature in order to maintain a stable
159 starting temperature.

160 The swimmers performed their regular training session and no changes were imposed
161 by the investigators. The intensity of the training session was recorded (categorized as
162 1: mild aerobic training; 2: moderate/intense aerobic training; 3: mild anaerobic training;
163 and 4: moderate/intense anaerobic training) in order to identify a possible confounding
164 effect. The training was performed at a chlorine disinfected open-air swimming-pool.

165 *Body temperature*

166 Oral temperature has been proposed to be related to the airways and/or the oral cavity,
167 as both are part of the respiratory tract and are affected differently than systemic
168 temperature during exercise (14), and therefore we have chosen axillary temperature
169 to evaluate body temperature. It was measured with an axillary thermometer
170 (MedCare®) before collecting baseline EBT.

171 **Statistical analysis**

172 Categorical variables are expressed as counts (%) and continuous variables as mean
173 (standard deviation - SD) or, if not normally distributed, as median (interquartile range -
174 IQR). Paired samples t-test was used to compare the differences between baseline
175 and post-exercise EBTs. Differences between asthmatic and non-asthmatic swimmers
176 were assessed with independent samples t-test for normally distributed data, Mann-
177 Whitney for non-normally distributed data or Chi-Square for categorical variables. A
178 new variable was created (Δ EBT = post exercise EBT – baseline EBT), and used to
179 assess if the variation of EBT was different among asthmatic and non-asthmatic
180 subjects. Correlations were assessed with Spearman's test.

181 All analyses were performed using SPSS and STATA and considering a $p < 0.05$ for
182 statistical significance.

183

184 **RESULTS**

185 Twenty-two elite swimmers accepted to participate, of which 10 had asthma. No
186 differences were observed between asthmatic and non-asthmatic swimmers for
187 demographic and personal characteristics, except for the expected lower PD₂₀ among
188 those with asthma (**table 1**).

189 **Table 1:**

190 ***Demographic and personal characteristics***

	Asthmatics (n=10)	Controls (n=12)	p
Men, n (%)	7 (70)	3 (23)	0.084
Age	17 ± 2.8	17 ± 2.9	0.946
Atopy, n (%)	4 (40)	4 (33)	1.000
PD20 methacholine	0.71 ± 0.6	4.39 ± 2.4	<0.001
FEV ₁ , Liters	4.26 ± 0.75	4.07 ± 0.92	0.622
FEV ₁ , % of predicted	111.10 ± 14.54	115.50 ± 8.89	0.393
FVC, Liters	5.13 ± 0.94	4.60 ± 1.20	0.271
FVC, % of predicted	114.50 ± 10.49	114.08 ± 11.48	0.931
FEV ₁ /FVC	83.52 ± 7.32	89.36 ± 7.18	0.074
FEF ₂₅₋₇₅ , Liters	4.22 ± 1.20	4.67 ± 1.17	0.387
FEF ₂₅₋₇₅ , % of predicted	97.3 ± 26.67	113.42 ± 22.54	0.140

191 Data presented as mean±SD, unless otherwise stated.

192 **FEF2575**: forced expiratory flow middle portion of FVC; **FEV1**: forced expiratory volume in one second;

193 **FVC**: forced vital capacity; **L**: liters; **PD20**: provocative dose inducing a 20% decrease in FEV1.

194 *Exhaled breath temperature*

195 Baseline and post-exercise EBT were not significantly different between asthmatic and
196 healthy swimmers (**table 2**). EBT significantly increased after exercise in all subjects
197 ($p=0.020$) (**figure 1**). This increase (Δ EBT) was not significantly different among
198 asthmatic swimmers and the healthy ones ($p=0.222$) (**figure 2**). Correlation between
199 PD_{20} and Δ EBT for asthmatic swimmers was not significant ($r=-0.103$, $p=0.777$).
200 Because 6 (60%) asthmatic swimmers were under therapeutic with inhaled
201 corticosteroids at the time of EBT collection, a second analysis was performed and no
202 differences were observed between those with and without therapy ($p=0.853$).

203 Correlations between axillary temperature and both baseline and post-exercise EBTs
204 were not significant ($p=0.972$ and $p=0.597$, respectively).

205 Regarding time to collect EBT, no differences were observed from baseline to post-
206 exercise measurements in the global sample ($p=0.822$). Also, for baseline EBT
207 collection, no differences occurred between asthmatic and non-asthmatic swimmers
208 ($p=0.366$). However, for EBT collection after exercise, asthmatic swimmers took
209 significantly less time to complete the measurement (4.2 ± 0.79 vs 5.0 ± 0.85 min,
210 $p=0.035$).

211 *Training*

212 No significant differences were observed between asthmatic and controls regarding the
213 intensity of the training session performed (**table 2**). The number of hours trained in the
214 week of EBT collection was similar in the two groups (**table 2**).

215 **Table 2:**

216 ***Temperatures measurements and training evaluation***

	Asthmatic swimmers (n=10)	Healthy swimmers (n=12)	p
Baseline EBT, °C	34.08 ± 0.60	33.49 ± 0.95	0.102
Post-exercise EBT, °C	34.23 ± 0.55	33.94 ± 0.64	0.275
Body Temperature*	35.97 ± 0.42	36.02 ± 0.31	0.769
Hours of training in that week	9.4 ± 2.6	9.3 ± 3.4	0.910
Intensity of the session, <i>n</i>			0.121 [#]
Mild aerobic	8	3	
Moderate/severe aerobic	1	0	
Mild anaerobic	1	5	
Moderate/severe anaerobic	2	2	

218 Data presented as mean±SD, except otherwise stated. *Axillary temperature was used as a measure for
219 body temperature. [#]Chi-square test.

220 **DISCUSSION**

221 The hypothesis on the basis of this study was that elite swimmers would present
222 increased EBT after a training session, due to the previously suggested idea that
223 exercise would cause inflammation, and furthermore that asthmatic swimmers would
224 experience a higher increase when compared to healthy controls. Interestingly, this
225 could not be confirmed by this study. Although EBT was significantly increased in all
226 subjects after training, supporting the hypothesis of heat loss during exercise, no
227 differences were observed between swimmers with previously diagnosed asthma and
228 healthy controls. It is therefore tempting to speculate that the theory of the heat loss as
229 an ethiopathogenic mechanism of exercise-induced asthma lacks evidence.

230 Our study has some limitations. The sample size might under power the study to draw
231 solid conclusions. Moreover, a cause that may have affected the results could be the
232 20 minutes of recovery (swimming with low effort) that swimmers made in the end of
233 the training session and that decreases cardiac rhythm, respiratory frequency and the
234 hyperpnea that makes EBT increase. If the temperature has been taken in the end of a
235 more vigorous exercise instead at the end of the training this could lead to a marked
236 results in Δ EBT.

237 Our results are consistent with previous studies that also have show an increase in
238 EBT after exercise (13), but no differences between asthmatic subjects and healthy
239 controls (14). That was different of other studies that shown differences in EBT of
240 asthmatic patients compared to healthy controls (5). This raises the question whether
241 the EBT could be a good inflammatory biomarker of asthmatics.

242 Previously has been hypothesized that dehydration of the airways with increased
243 osmolality in the mucosal surface liquid leads to mast cell degranulation (21, 22),
244 releasing bronchoconstriction mediators, such as bradykinin and histamine, and
245 activating endothelial nitric oxide synthase (23), witch increase the vasodilation effect.
246 This could suggest that elevated EBT could be a potential marker of a temporary effect
247 on the airways, rather than of the long-term inflammation of the asthmatic disease. So
248 the increase of EBT could be a physiological response of the airways to the increase of
249 ventilatory rate, rather than a pathological pathway which can be involved in
250 development of exercise-induced asthma.

251 This study leads us to hypotheses that the increase of EBT and there for the elevation
252 of airway inflammation shown in previous studies (5, 6), during exercise (13), could be

253 a protect measure of the airways against heat loss and dehydration, that occurs with
254 the increase of ventilatory rate, rather than a pathological result.

255

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ANEXO

1 - Normas da revista: *Pediatric Allergy and Immunology*

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If the manuscript is longer, reasons for increase in length, figure or table number or reference number should be stated in the cover letter. In general, the printed version of the manuscript should not occupy more than 6 pages.

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Books and monographs: Stiehm ER, Fulginiti VA. Immunologic disorders in infants and children. Philadelphia: WB Saunders 1973.

Chapter in book: Holt PG, Turner KJ. Regulation of IgE synthesis in man and experimental animals. In: Lessof MH, Lee TH, Kemeny DM, eds. Allergy, an international textbook. New York: John Wiley 1987: 69-87.

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