

# Running, walking, and hyperventilation causing asthma in children

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**ABSTRACT** To examine further the relation between type of exercise, workload, ventilation, and exercise-induced asthma, we compared treadmill walking with treadmill running and treadmill running with isocapnic hyperventilation in separate studies in children and adolescents. Inspired air conditions were identical during each pair of tests. Walking and running with similar minute ventilation and oxygen consumption were followed by similar falls in peak expiratory flow rate as were running and hyperventilation with similar minute ventilation and end-tidal carbon dioxide tension. This study supports the concept that hyperventilation is a central mechanism in exercise-induced asthma.

It has been suggested that different forms of exercise cause different degrees of bronchospasm in subjects susceptible to exercise-induced asthma (EIA), with free-range running, treadmill running, cycling, walking, and swimming in approximately diminishing order of effect (Anderson *et al*, 1971; Fitch and Morton, 1971; Anderson, 1972; Silverman and Anderson, 1972a). The reasons for this are unclear. Previous studies in children have shown treadmill running to cause more bronchospasm than treadmill walking, even when workload was well matched during the comparisons (Silverman, 1973; Eggleston and Guerrant, 1976; James *et al*, 1976). A recent study in adults, however, showed no difference in the bronchial response to treadmill running compared to cycling, where oxygen consumption and duration of exercise were the same (Miller *et al*, 1975).

The fact that minute ventilation was well matched during, for instance, comparisons between swimming and running exercise (Anderson, 1972) lead to doubt about the importance of hyperventilation in the mechanism of EIA. Yet an early study showed bronchospasm could be precipitated by isocapnic hyperventilation in some asthmatic subjects (Chan Yeung *et al*, 1971), suggesting hyperventilation could be an important factor in EIA. Later studies showed minimal or variable bronchospasm from hyperventilation (Vassalo *et al*, 1972; McFadden *et al*, 1977); one of these studies used a rebreathing circuit to obtain isocapnia (McFadden *et al*, 1977). A recent study using a non-rebreathing circuit showed that

isocapnic hyperventilation and hyperventilation from exercise produced similar degrees of airways obstruction (Zeballos *et al*, 1978). Other recent studies have shown a reduction in EIA with nose breathing as compared with mouth breathing (Shturman-Ellstein *et al*, 1978), the abolition of EIA by the inspiration of saturated air at body temperature (Chen and Horton, 1977), and the exacerbation of EIA by cold-air breathing during exercise (Strauss *et al*, 1977a). These observations imply that air conditions are crucial in the development of EIA and airways obstruction after hyperventilation.

We aimed to examine the relation between type of exercise, workload, ventilation, and induced asthma with two studies designed to reduce, as far as possible, the effects of environmental factors known to affect EIA.

## Patients and methods

### WALKING AND RUNNING

Eight boys and two girls, aged 11 to 17 years, performed a treadmill walking test and a treadmill running test on the same day. Anthropometric data and room temperature and relative humidity (Paper Hygrometer, Gallenkamp) at the start of each test are shown in table 1. All subjects had asthma, as defined by Scadding (1966), and all had previously experienced EIA after treadmill running. No subject was being treated with systemic corticosteroids, and all other drugs were omitted for at least 12 hours before tests. Initial peak

Table 1 Anthropometric data; inspired air temperature and relative humidity (RH)

Subject No	Age (yr)	Sex	Weight (kg)	Height (cm)	Walking-running comparison				Running-hyperventilation comparison			
					Walking		Running		Running		Hyperventilation	
					t (°C)	RH (%)	t (°C)	RH (%)	t (°C)	RH (%)	t (°C)	RH (%)
1	15	M	58.0	175	24.3	72	24.5	71	14.8	64	14.9	63
2	17	M	55.0	169	24.5	72	24.6	71	14.8	62	14.9	63
3	12	F	53.8	159	20.5	76	20.8	74				
4	13	F	51.0	161	20.0	76	17.8	74				
5	11	M	35.0	151	23.0	75	22.5	74	21.5	71	21.1	71
6	12	M	39.5	150	20.0	64	20.8	62	21.4	71	20.5	71
7	11	M	33.0	142	20.0	64	20.8	62				
8	14	M	55.0	170	19.5	75	20.5	75	17.5	57	18.0	56
9	13	M	38.5	153	17.5	74	17.5	74				
10	11	M	30.0	140	17.7	62	16.0	62				
11	13	M	35.0	150					21.2	72	21.3	72
12	13	M	45.9	152					12.5	54	12.8	53
13	13	F	53.7	163					15.3	58	15.6	52
				Mean	20.7	71.0	20.6	69.9	17.4	63.6	17.4	62.6
				SD	2.5	5.5	2.9	5.6	3.6	7.1	3.3	8.3

expiratory flow rates (PEFR) are shown, as percentage of expected normal values (Godfrey *et al*, 1970), in table 2. The test order was randomly selected, and the periods of exercise separated by two-and-a-half hours, a period long enough to avoid the refractory period known to follow EIA (Edmunds *et al*, 1978) but short enough to avoid potentially considerable day-to-day changes in air temperature and relative humidity. Exercise tests consisted of six minutes of non-stop walking or running on a treadmill at 5.3 kph. The treadmill slope was greater for the walking test, the aim being to achieve a similar oxygen consumption for each test, using the data of Silverman and Anderson (1972b) as a guide to the appropriate treadmill settings.

For two minutes before exercise, and during the

whole exercise, the subjects wore a noseclip and breathed through a mouthpiece attached to a low-resistance, low dead-space (50 ml) mobile valve. Inspired ventilation was measured with a dry gas meter (Parkinson Cowan CD 4), and the expired gas was mixed in a 4 l chamber and continuously sampled for CO<sub>2</sub> (measured by Godart infrared analyser) and O<sub>2</sub> (Servomex paramagnetic analyser). The data, including ECG, were continuously recorded by an ink jet chart recorder (Mingograf 81) during each exercise test. The analysers were calibrated before and after each test, using gases of concentration chemically determined by the Haldane technique.

Airway function was tested by measuring PEFR with a Wright's meter, immediately before and 1, 3, 5, 10, 15, and 20 minutes after exercise.

Table 2 Walking/running comparison: results

Subject No	First test	Walking				Running							
		Mean oxygen consumption (ml/min/kg)		Mean expired ventilation (l/min BTPS)		Initial PEFR (% of expected)	Fall in PEFR (%)	Mean oxygen consumption (ml/min/kg)		Mean expired ventilation (l/min BTPS)		Initial PEFR (% of expected)	Fall in PEFR (%)
		Whole test	Last 4 min	Whole test	Last 4 min			Whole test	Last 4 min	Whole test	Last 4 min		
1	Walk	29.7	31.8	54.0	57.5	74.6	63	29.4	31.4	50.0	50.8	70.6	54
2	Walk	26.4	28.3	37.5	41.5	101.1	35	27.4	29.3	42.0	45.8	97.9	22
3	Run	23.9	25.4	32.3	35.3	58.3	54	26.0	28.8	40.8	46.1	74.0	67
4	Run	25.6	28.2	38.2	42.5	100.5	15	27.9	32.2	44.0	50.5	89.8	17
5	Walk	27.7	30.0	30.7	34.0	43.4	55	30.4	32.9	33.3	36.0	46.8	58
6	Walk	29.3	32.1	33.5	36.8	67.2	52	29.7	32.0	32.7	36.8	73.9	55
7	Run	31.5	35.1	31.2	35.5	60.6	63	28.2	30.8	25.7	27.8	80.3	70
8	Walk	30.4	32.9	37.2	42.4	60.9	27	29.0	30.8	35.0	38.7	60.9	27
9	Run	27.3	29.7	26.4	29.5	86.6	51	30.7	32.9	29.1	32.1	90.4	36
10	Run	26.4	28.9	23.5	25.8	75.0	33	28.6	30.4	27.8	29.5	62.5	21
	Mean	27.8	30.2	34.5	38.1	72.8	44.8	28.7	31.2	36.0	39.4	74.7	42.7
	SD	2.8	2.8	8.9	8.7	18.8	16.3	1.4	1.4	7.9	8.5	15.6	20.3

The results of tests on six other children were excluded for the following reasons: three children failed to develop EIA (defined as fall in peak flow of 15% or more) with either walking or running; one child developed severe asthma necessitating bronchodilator treatment after his first test; and in two tests essential data were unavailable because of difficulties with measuring or recording equipment.

#### HYPERVENTILATION AND RUNNING

Seven boys and one girl, aged 11 to 17 years, performed a treadmill running test and a voluntary isocapnic hyperventilation test on the same day. Anthropometric data and air conditions at the start of each test are shown in table 1. Subject selection and test procedure were similar to that described for the walking-running comparison, except that only end-tidal CO<sub>2</sub> and inspired ventilation were measured during each six-minute exercise or voluntary hyperventilation period. As before, test order was randomly selected, and an interval of two-and-a-half hours was left between tests.

For two minutes before each test, and during the whole of each test, the subjects breathed through a valve as described above. During the voluntary hyperventilation test each subject was encouraged to hyperventilate to achieve a "target" minute ventilation and respiratory frequency, based on the values obtained during a previous treadmill running test. End-tidal CO<sub>2</sub> was held constant by adding 100% CO<sub>2</sub> through a Rotameter flowmeter, at 0.4–1.5 l/min, to the wide-bore corrugated respiratory line, 1.3 m from the valve. This did not produce measurable changes in inspired air temperature or relative humidity. PEF<sub>R</sub> was measured as for the walking-running comparison.

The "target" consisted of an electrically driven, continuously revolving pointer fixed over the dial of the gas meter, so that both the additional pointer and the hand of the gas meter could be easily seen by subjects. Subjects were asked to breathe in time to a metronome, with sufficient depth to match the rotation of the hand of the gas meter with that of the electrically driven pointer. Thus any target ventilation could be achieved merely by the setting of the additional pointer.

In the analysis of results, paired observations were compared using the *t*-test (for absolute values) or Wilcoxon's rank sum test (for percentages).

## Results

#### WALKING AND RUNNING

Table 2 summarises the results of the walking-running comparison. In addition to the data shown, paired comparisons were made on air temperature and relative humidity, barometric pressure, heart rate during exercise, tidal volume, and absolute fall in PEF<sub>R</sub> (as opposed to percentage fall). No significant differences were found in any of the parameters examined. In addition there were no differences in initial PEF<sub>R</sub> or subsequent fall in PEF<sub>R</sub> between the first and second tests, or when running or walking was the first test performed.

In individual subjects during both walking and running, oxygen consumption and minute ventilation increased in a similar pattern over the first three minutes and tended to be stable over the last four minutes, though with greater variation in the sixth minute than in the third, fourth, and fifth minutes. While the additional oxygen consumption related to the exercise cannot be determined, it is thought that similar oxygen consumption between the two forms of exercise implies similar workload. Greater individual variations were noted with minute ventilation between the two forms of exercise, though mean values were similar, and there were no significant differences between walking and running. The initial PEF rates of 43–101% (mean 74%) showed the subjects generally had mild to moderate asthma before testing, and falls in PEF<sub>R</sub> of 15–70% (mean 43%) showed a pronounced, though not extreme, response to the level of exercise chosen.

#### RUNNING AND HYPERVENTILATION

Table 3 summarises the results of the running-hyperventilation comparisons. No significant differences were found between values of mean end-tidal CO<sub>2</sub>, mean inspired ventilation, initial PEF<sub>R</sub>, or fall in PEF<sub>R</sub>; respiratory frequency was greater during voluntary hyperventilation (*p* just <0.05). Though there were considerable individual variations in fall in PEF<sub>R</sub> between the two tests, mean values were similar, and there was no significant difference between running or voluntary hyperventilation (whether expressed as percentage fall or absolute fall in PEF<sub>R</sub>).

## Discussion

#### WALKING AND RUNNING

We have shown that under identical environ-

Table 3 Treadmill running/isocapnic hyperventilation comparison: results

Subject No	First test	Running					Hyperventilation				
		Mean PetCO <sub>2</sub> (kPa)	f (l/min)	Mean V <sub>1</sub> (l/min BTPS)	Initial PEFR (% of expected)	Fall in PEFR (%)	Mean PetCO <sub>2</sub> (kPa)	f (l/min)	Mean V <sub>1</sub> (l/min BTPS)	Initial PEFR (% of expected)	Fall in PEFR (%)
8	Run	4.8	22	37.3	58.8	27.0	5.1	33	36.2	54.6	46.0
5	Hypervent	4.6	43	26.3	50.0	33.3	4.9	55	28.4	48.5	43.7
6	Hypervent	5.1	51	22.3	40.5	16.7	4.3	62	19.8	40.5	30.0
11	Run	5.0	38	32.0	117.0	51.0	5.0	44	35.5	115.0	47.0
12	Run	5.3	37	49.2	110.0	40.0	5.0	46	50.5	105.5	43.4
2	Hypervent	5.2	40	52.8	98.0	45.6	4.4	49	55.5	98.0	46.7
1	Hypervent	4.5	38	60.0	63.6	62.5	4.5	32	49.2	75.5	65.7
13	Run	4.6	44	58.0	78.0	53.0	4.7	78	50.0	83.5	18.3
	Mean	4.9	39.1	42.2	77.0	41.1	4.7	49.9	40.6	77.6	42.6
	SD	0.3	8.3	14.7	28.5	15.0	0.3	15.2	12.6	27.7	13.8

On paired *t* tests, no significant differences except for frequency (0.05 > *P* > 0.01).

Key: PetCO<sub>2</sub>=end tidal PCO<sub>2</sub>; f=Frequency of breathing; V<sub>1</sub>=Inspired minute ventilation.

mental conditions when the severity of exercise is closely matched, EIA is equally severe after running and walking exercise. These results differ from previous studies on this subject. Some investigators attempted to standardise workload using heart rates without measurement of ventilation or oxygen consumption (Eggleston and Guerrant, 1976; James *et al*, 1976). Eggleston and Guerrant (1976) used a longer period of walking than of running, and workload and ventilation for the two forms of exercise were probably not comparable. In another walking versus running comparison with five subjects significantly greater airways obstruction occurred after running, despite achieving similar values of expired ventilation and oxygen consumption over the last minute of exercise (Silverman, 1973). Our study suggests the last minute of exercise may not be representative of the whole period of exercise, or even of the last four minutes of exercise during which oxygen consumption, expired ventilation, and other indices of workload are generally stable. In addition the subjects studied by Silverman (1973) performed running and walking tests on different days, so that variations in patient response, air temperature, and relative humidity could have led to different airway responses.

In a running versus cycling comparison in adults, Miller *et al* (1975) found no significant difference in airway response with matched total oxygen consumption, although the minute ventilation was not identical for the two forms of exercise. Possibly the level of energy expenditure with both forms of exercise was sufficient to provoke a maximal airway response in individual subjects at the time of testing.

Strauss *et al* (1977b) comparing arm and leg work, found greater EIA from arm work with a

similar external workload, and proposed a relation between workload, muscle mass, and EIA. The explanation is probably the significantly higher ventilation found in their subjects during the arm work.

On the evidence currently available, it appears that different types of exercise produce similar degrees of EIA providing ventilation is similar, and provided that environmental conditions are similar. The differences between matched periods of exercise carried out in different environments (laboratory, open-air or heated swimming-pool) may possibly be due entirely to the local conditions of air temperature and humidity (Chen and Horton, 1977).

#### RUNNING AND HYPERVENTILATION

The results of our treadmill running versus isocapnic hyperventilation study are similar to the recently published study of Zeballos *et al* (1978), which compared treadmill walking and isocapnic hyperventilation. In contrast to our study testing was carried out on different days of the same week, periods of three and 10 minutes both of walking and hyperventilation were compared, a wide variety of respiratory function tests were done, though only once after exercise, and similar respiratory frequencies were achieved during both exercise and hyperventilation. In both studies rigorous attention was given to maintaining isocapnia in non-breathing circuits. Despite differences in design, both studies appear to confirm that isocapnic hyperventilation causes airways obstruction as severe as that produced by exercise performed at the same minute ventilation. Hyperventilation is probably the central mechanism in EIA.

Earlier reports of the airway response to volun-



tary isocapnic hyperventilation varied, one study (Chan Yeung *et al*, 1971) showing consistent bronchospasm, while others showed variable (Vasallo *et al*, 1972) or minimal (Silverman, 1973) response. Some later studies, including that of McFadden *et al* (1977), showed no response to isocapnic hyperventilation. Recent studies have shown the importance of inspired air temperature and relative humidity in EIA, with reduction or abolition of EIA by warming and humidifying the inspired air (Chen and Horton, 1977; Strauss *et al*, 1978). The same phenomenon would appear to apply in hyperventilation-induced asthma, explaining the negative response to hyperventilation where partial rebreathing of expired air was used to maintain the  $PCO_2$  during hyperventilation (Silverman, 1973; McFadden *et al*, 1977).

Moreover, a correlation has been shown between respiratory heat and water loss from increased ventilation and the severity of EIA (Chen and Horton, 1977; Strauss *et al*, 1978). The exact sites of the receptors and the subsequent mediating mechanisms remain to be elucidated, although oropharyngeal epithelial receptors probably play an important part (Shturman-Ellstein *et al*, 1978). The confirmation of an important oropharyngeal reflex mechanism in asthma is obviously of great importance, opening up new avenues for the prophylaxis and treatment of asthma.

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