Effect of high ambient temperature on the kinetics of the pupillary light reflex in healthy volunteers

N. K.-C. LEUNG, C. M. BRADSHAW & E. SZABADI*

Department of Psychiatry, University of Manchester, Stopford Building, Oxford Road, Manchester, M13 9PT

Miotic responses to brief light stimuli were studied in healthy volunteers under two ambient temperature conditions, 22° C and 40° C. The latency and amplitude of the light reflex did not differ between the two conditions, but the recovery time of the reflex was significantly shorter under the 40° C condition than under the 22° C condition. The results are consistent with the hypothesis that exposure to high ambient temperature results in an increased sympathetic drive to the iris dilator muscle but does not influence the parasympathetic light reflex.

Keywords ambient temperature pupillary light reflex sympathetic reflex

Introduction

Exposure to high ambient temperature results in changes in autonomic functions, for example enhancement of physiological tremor (Al-Eithan *et al.*, 1991) and hyperresponsiveness of eccrine sweat glands to intradermally injected cholinomimetics (Banjar *et al.*, 1987), which have been attributed to increased impulse flow in peripheral sympathetic nerves (e.g. Bini *et al.*, 1980; Szabadi *et al.*, 1988). We report here the effects of high ambient temperature on the kinetics of the pupillary light reflex. The latency and amplitude of the miotic response to light stimulation are determined by the parasympathetic reflex arc, whereas the recovery time of the response is believed to be determined in part by sympathetically mediated redilatation (Ishikawa *et al.*, 1977; Smith, 1988).

Methods

Subjects

Twelve drug-free healthy male volunteers (19–45 years) participated. All gave their written informed consent. The study was approved by the local Ethics Committee.

Apparatus

Pupil diameter was monitored using an infrared television pupillometer (Applied Science Laboratories). Twelve 200 ms light stimuli of graded intensities were delivered via an assembly of three light-emitting diodes (peak wavelength, 565 nm) mounted on a headband and positioned 1 cm from the cornea of the subject's right eye. The intensities of the stimuli (incident light intensity, measured 1 cm from the source) ranged from 5.3×10^{-5} to 3.59 mW cm⁻². Stimulus presentation was controlled by a BBC-B microcomputer, which also recorded the pupillometric data.

Procedure

Each subject participated in two sessions 7 days apart. In one session the test cubicle was maintained at $22^{\circ} C \pm 2^{\circ} C$, and in the other it was maintained at $40^{\circ} C \pm 2^{\circ} C$, the order of conditions being counterbalanced across subjects. At the beginning of each session the subjects were first acclimatized to red light (30 min) and to the ambient temperature in the test cubicle (20 min). Then resting pupil diameter was measured, followed by miotic responses to the twelve light stimuli. The stimuli were presented at 30 s intervals in the order of increasing intensity.

Analysis

Resting pupil diameter was compared between the two temperature conditions using Student's *t*-test (paired comparisons). The latency, amplitude and 75% recovery time of the miotic responses to the light stimuli (Bakes *et al.*, 1990) were analyzed by two-factor analyses of variance (temperature condition, light intensity), with repeated measures on both factors. Pearson's product-moment correlation coefficient was calculated for the

Correspondence: Dr C. M. Bradshaw, Department of Psychiatry, University of Manchester, Stopford Building, Oxford Road, Manchester, M13 9PT

*Present address: Department of Psychiatry, University of Nottingham, Queen's Medical Centre, Nottingham, NG7 2UH, U.K.

relationship between each parameter of the light reflex and the logarithm of the light intensity under each temperature condition.

Results

The mean (\pm s.e. mean) resting pupil diameter did not differ significantly between the two temperature conditions (22° C: 6.7 mm \pm 0.4 mm; 40° C: 6.8 mm \pm 0.2 mm; t(11) = 0.6; P > 0.1).

Under both temperature conditions, the latency, amplitude and 75% recovery time of the miotic responses were approximately linearly related to the logarithm of the light intensity (Figure 1, Table 1). Analysis of variance revealed a significant main effect of light intensity on each parameter (latency: F(11, 121) = 49.7; amplitude: F(11, 121) = 117.9;75% recovery time: F(11, 121) =26.8; P < 0.001 in each case). In the case of latency and amplitude, there was no significant main effect of temperature condition (latency: F(11, 121) = 1.1; amplitude: F(11, 121) = 0.8; P > 0.1 in each case). 75% recovery time showed a significant main effect of temperature condition (F(1, 11) = 7.3; P < 0.02), and a significant interaction effect (F(11, 121) = 3.1; P < 0.01), reflecting a shortening of the recovery times of responses to higher light intensities under the 40° C condition.

Table 1Correlation coefficients (r) for relationshipsbetween parameters of the light reflex and log lightintensity under the two temperature conditions (c.f.Figure 1)

	Temperature	
	22° C	40° C
Latency	-0.947*	-0.993*
Amplitude	+0.990*	+0.991*
75% recovery time	+0.971*	+0.964*

```
*P < 0.001
```

Discussion

The latency and amplitude of the miotic response to light are determined by activity in the parasympathetic reflex arc, whereas the recovery time of the response is believed to be determined, in part, by active redilatation brought about by the sympathetic innervation of the radial muscle of the iris (Smith, 1988). The present finding that exposure to high ambient temperature shortened the recovery time of the light reflex without affecting the latency or amplitude of the response is thus consistent with the hypothesis that exposure to high temperature results in an increased sympathetic drive to the radial muscle, but does not influence the parasympathetic reflex. The present results are also consistent with our previous findings that systemic treatment with clonidine, a drug which is believed to suppress sympathetic outflow (Kobinger, 1978), prolongs the recovery time of the light reflex (Morley et al., 1991).

The present results may have implications for the changes in autonomic function seen in clinical anxiety states. Such changes have traditionally been ascribed to sympathetic overactivity (see Szabadi & Bradshaw, 1988). However, we recently observed that the amplitude of the light reflex (an index of parasympathetic activity) was smaller in a group of patients suffering from generalized anxiety disorder than in a matched group of healthy volunteers, whereas the 75% recovery time of the reflex did not differ between the two groups (Bakes et al., 1990), suggesting that there may be suppression of the parasympathetic drive, rather than facilitation of the sympathetic drive to the iris in these patients. This interpretation is strengthened by the demonstration that interventions believed to suppress (clonidine) or enhance (high ambient temperature) impulse traffic in sympathetic nerves selectively alter the recovery time of the light reflex, without affecting its amplitude.

This work was supported by the North Western Regional Health Authority. We are grateful to R. W. Langley for skilled technical assistance.

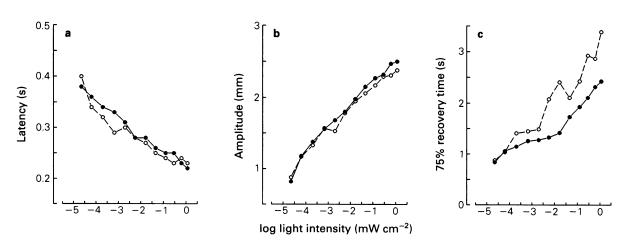


Figure 1 Relationship between parameters of the light reflex and the intensity of the light stimulus. Ordinates: A, latency (s); B, amplitude (change in pupil diameter, mm); C, 75% recovery time (s). Abscissa: light intensity (mW cm⁻², logarithmic scale). Points are mean data (n = 12); symbols indicate treatment conditions: $\circ 22^{\circ}$ C; $\bullet 40^{\circ}$ C.

References

- Al-Eithan, M., Banjar, W. M. A., Gazzaz, J., Bradshaw, C. M. & Szabadi, E. (1991). Effects of high ambient temperature and terbutaline on physiological tremor. Br. J. clin. Pharmac., 31, 603P.
- Bakes, A., Bradshaw, C. M. & Szabadi, E. (1990). Attenuation of the pupillary light reflex in anxious patients. *Br. J. clin. Pharmac.*, **30**, 377–382.
- Banjar, W., Longmore, J., Bradshaw, C. M. & Szabadi, E. (1987). The effect of diazepam on the responsiveness of human eccrine sweat glands to carbachol: Influence of ambient temperature. *Eur. J. clin. Pharmac.*, **31**, 661–665.
- Bini, G., Hagarth, K. E., Hynniken, P. & Wallin, B. G. (1980). Regional similarities and differences in thermoregulatory vaso- and sudomotor tone. J. Physiol. (Lond.), 306, 553-565.
- Ishikawa, S., Oono, S. & Hikita, H. (1977). Drugs affecting the iris. In *Drugs and ocular tissues*, ed. Kikstein, S., pp. 288–382. Basel: Karger.

- Kobinger, W. (1978). Central alpha-adrenergic system as target for hypotensive drugs. *Rev. Physiol. Biochem. Pharmac.*, 81, 39–100.
- Morley, M. J., Bradshaw, C. M. & Szabadi, E. (1991). Effects of clonidine and yohimbine on the pupillary light reflex and carbachol-evoked sweating in healthy volunteers. *Br. J. clin. Pharmac.*, **31**, 99–101.
- Smith, S. A. (1988). Pupillary function in autonomic failure. In Autonomic failure (2nd edition), ed Bannister, R., pp. 393–412. Oxford: Oxford University Press.
- Szabadi, E. & Bradshaw, C. M. (1988). Biological markers in anxiety states. In *Recent advances in clinical psychiatry*, *Vol. 6.*, ed. Granville-Grossman, K., pp. 69–99. Edinburgh: Churchill-Livingstone.

(Received 7 November 1991, accepted 2 January 1992)