Determinat of the distribution of conduction velocities in workers exposed to lead, zinc, and copper

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ABSTRACT By measuring the distribution of conduction velocities (DCV) in sensory fibres of the median nerve, the effects of asymptomatic increased lead, zinc, and copper absorption on the conduction velocities of slower and faster nerve fibres were examined in 20 gun metal founders with blood lead (BPb) concentrations of 22–59 (mean 39) μg/dl (1.1–2.8 (mean 1.9) μmol/l). The results indicated that the conduction velocities of slower fibres were inversely affected by lead absorption, resulting in a significant reduction in the velocities at the BPb level of 40–59 μg/dl; on the other hand, the conduction velocities of faster fibres and the conventional sensory nerve conduction velocity were presumably positively affected by zinc absorption without a significant reduction in the velocities of faster fibres. It is suggested that assessment of the possible antagonistic effect of zinc is essential when conduction velocities are not decreased in lead exposed workers.

Seppäläinen and coworkers have shown that slower and maximal motor nerve conduction velocities decrease at levels of occupational lead exposure previously considered “safe”—that is, at blood lead (BPb) concentrations below 70 μg/dl (3.4 μmol/l). We have shown that the maximal sensory nerve conduction velocity is also affected in adults with asymptomatic increased lead absorption. These observations, although they have not been made in all published reports, have in the main been confirmed by subsequent investigations.

Peripheral nerve trunks consist of several thousand myelinated nerve fibres with slower and faster conduction velocities in parallel to the fibre diameters. The conventional methods for determining conduction velocities yield only two discrete values that reflect the maximal and slower velocities of fibres in the nerve trunk; the function of the vast majority of fibres cannot be directly assessed. In 1979 three groups of investigators independently introduced new and different transcutaneous techniques to derive the distribution of conduction velocities (DCV) of large myelinated fibres (alpha fibre group) by means of computer analysis. Recently, we have established one of these methods in our laboratory. In the present study our technique is applied to workers with asymptomatic increased lead, zinc, and copper absorption.

SUBJECTS AND METHODS

SUBJECTS

The nature of the procedure in the present study was fully explained to all subjects, and the study was carried out with their informed consent. The 20 subjects, aged 34 to 59 (mean 48), were male gun metal founders employed at a metal factory for two to 17 (mean 10) years; their BPb concentrations ranged from 22 to 59 (mean 39) μg/dl. In nine of the workers (high lead group) the BPb was 40 μg/dl or above; in the remaining workers (low lead group) it was below 40 μg/dl. The plasma zinc (PZn) concentrations ranged from 73 to 111 (mean 88) μg/dl (11.2 to 17.0 (mean 13.5) μmol/l) in 20 gun metal founders; the plasma copper (PCu) concentrations were 64 to 126 (mean 91) μg/dl (10.1 to 19.8 (mean 14.3) μmol/l). These BPb, PZn, and PCu concentrations were significantly higher than those in all (16) male steel foundry workers at the same factory—that is, 8–20 (mean 13) μg/dl for BPb, 54–97 (mean 72) μg/dl for PZn, and 50–100 (mean 73) μg/dl for PCu, respectively (p < 0.05). All gun metal founders drank alcohol equivalent to 0–760 ml of 100% ethanol a week (mean 283); six of them had a history of mental fume fever. None of the subjects had specific signs and symptoms indicative of clinical lead poisoning or

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CAPs at wrist (above) and elbow (below)

Time after stimulation (m/sec)

SFAP calculated

Time after stimulation (m/sec)

Fig 1 Compound action potentials (CAPs) recorded from median nerve at wrist and elbow after stimulation of second finger in a gun metal founder, and calculated single fibre action potential (SFAP, relative amplitude) corresponding to conduction velocity of 67.5 m/sec.

of neurological or other endogenous disorders; none was exposed in their workplace to such chemicals as arsenic and solvents, which might also have neurotoxic effects.

Control subjects for electrophysiological studies were 20 male "healthy" adults neither occupationally exposed to lead, zinc, or copper nor having neurological disorders; ages ranged from 31 to 58 (mean 45). There were no significant differences in age, height, skin temperature, and alcohol and tobacco consumption between either of the high and low lead groups and the 20 control subjects at the time of electrophysiological studies (Wilcoxon's rank sum test, p > 0.05).

ELECTROPHYSIOLOGICAL STUDIES

The DCV was measured by the same method reported previously by us (a modified method of Barker et al). The study was conducted in a warm laboratory (28–32°C) by the use of a Medelec MS-92 two channel electromyograph; skin temperature was maintained in the range of 32–35°C for all subjects. The right median nerve was stimulated repeatedly with stainless ring cathode and anode electrodes, tied around the proximal and distal interphalangeal joints of the second finger respectively, with a 70 to 100 V square wave pulse of 0.2 msec duration. The compound action potential (CAP) was picked up with surface electrodes at the wrist and the elbow. The responses were averaged 200 to 1000 times until no changes were discerned in the recorded CAP on the oscilloscope or electromyograph. Typical records of the CAP are illustrated in fig 1 together with the single fibre action potential calculated. The CAP was digitised with a sampling interval of 0.04 msec and stored for DCV analysis on the discs of the Hitachi MB6891 programmable electric calculator.

The DCV was calculated by the double conduction distance method between the velocities of 30.0 and 77.5 m/sec with an interval of 2.5 m/sec from the two CAPs recorded at the wrist and the elbow. The calculation was continued until the estimated numbers of nerve fibres with varying conduction velocities converged to the values that did not change significantly.

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Fig 2 Distribution of conduction velocities in 20 gun metal founders and 20 control subjects (averages and standard errors).
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The DCVs in 20 gun metal founders and in 20 control subjects are illustrated in fig 3. The V10 velocity and SCVs were significantly slowed in the high lead group of gun metal founders (fig 3). There were no significant differences in other parameters of DCV between the high lead group and control subjects; similarly, in all parameters between the low-lead group and control subjects (p > 0.05).

The V5, V10, and V20 velocities were inversely correlated with indicators of lead absorption (MPb and BPb) as well as PCu in 20 gun metal founders (fig 4 and table); the Spearman's rank correlations for these relations were also significant (−0.615 for V5 velocity and MPb, p < 0.01; −0.518 for V10 velocity and BPb, p < 0.05). PCu was positively correlated with MPb (r = 0.536, p < 0.05). On the other hand, the V80, V90, and Vpeak velocities and SCVs were positively correlated with indicators of zinc absorption (UZn and EZn) (fig 4 and table). The SCV (finger to elbow) was significantly correlated with the V70, V80, and V90 velocities (r = 0.447, 0.579, and 0.646, respectively, p < 0.05).

**Factors significantly correlated with the distribution of conduction velocities and with sensory nerve conduction velocity in 20 gun metal founders†**

<table>
<thead>
<tr>
<th>Conduction velocity</th>
<th>Factors (Pearson's product moment correlation, r, in parentheses)</th>
</tr>
</thead>
<tbody>
<tr>
<td>V5</td>
<td>MPb (−0.595**); PCu (−0.450*)</td>
</tr>
<tr>
<td>V10</td>
<td>BPb (−0.459*); MPb (−0.542*); PCu (−0.497*)</td>
</tr>
<tr>
<td>V20</td>
<td>MPb (−0.440*); PCu (−0.459*)</td>
</tr>
<tr>
<td>V80</td>
<td>UZn (0.459*)</td>
</tr>
<tr>
<td>V90</td>
<td>UZn (0.482*)</td>
</tr>
<tr>
<td>Vpeak</td>
<td>UZn (0.514*)</td>
</tr>
<tr>
<td>SCV (finger to wrist)</td>
<td>EZn (0.452*)</td>
</tr>
<tr>
<td>SCV (finger to elbow)</td>
<td>UZn (0.455*)</td>
</tr>
</tbody>
</table>

* p < 0.05; ** p < 0.01.
† Abbreviations as in text.
Fig 4  Relations between distribution of conduction velocities (V5, V10, V90, and Vpeak velocities) or sensory nerve conduction velocity (SCV) and indicators of lead and zinc absorption (MPb, BPb, UZn, and EZn) in 20 gun metal founders. * indicates Pearson's product moment correlation. One μg/dl of BPb corresponds to 0.05 μmol/l; 1 mg/24 h of MPb to 4.8 μmol/24 h; 1 mg/24 h of UZn to 15.3 μmol/24 h; and 1 mg/dl of EZn to 0.15 mmol/l.
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Discussion

This is probably the first report to disclose the significant effects of lead and zinc on the DCV; it is suggested that DCV analysis is useful for evaluation of subclinical lead neuropathy. Sax et al23 applied another technique of DCV analysis (method of Kovacs et al19) to four lead exposed subjects, but failed to find a consistent pattern of the changes. This is probably due to the small sample size or a difference of their technique from ours.

The V5, V10, and V20 velocities were significantly correlated with two or three (15 or 23%) of all the 13 indicators of lead, zinc, or copper absorption examined; these indicators were consistent (MPb, PCu (and BPb)). Furthermore, the significance level for the V5 velocity was substantially high (p < 0-01). These results, together with a significant slowing of the V10 velocity, suggest that slower fibre conduction velocities of the sensory median nerve were inversely affected by lead, resulting in a conduction delay at the BPb level of 40–50 μg/dl. Significant correlations of BPb and MPb with conventional motor and sensory nerve conduction velocities have been shown in some studies.2 8 10 11 The inverse relation of PCu to the V5, V10, and V20 velocities is considered to reflect lead effects on the slower fibre velocities, as PCu was positively correlated with MPb.

Seppäläinen et al have shown that the conduction velocity of the slower motor fibres of the ulnar nerve (CVSF) is particularly sensitive to lead.1 24 It appears that both motor and sensory slower fibres are sensitive to lead. Histological studies are needed to clarify the changes underlying these physiological alterations.

Faster fibre conduction velocities (V80 and V90 velocities), on the other hand, were positively correlated with UZn; similarly, SCVs were correlated with EZn and UZn. It is thus suggested that zinc antagonised a lead induced conduction delay in these nerve fibres. These data are consistent with another observation by us that zinc possibly antagonised the effects of lead on the conventional motor and sensory conduction velocities of the radial nerve in lead exposed workers.25 As no indicator of lead and copper absorption was significantly correlated with indicators of zinc absorption in the present study (p > 0-05), the effects of zinc on the nerve conduction velocities are considered to be independent of the effects of those metals. Alternatively, the correlations between zinc absorption and nerve conduction velocities might have been random results; therefore, epidemiological and clinical studies with a greater number of subjects are required.

The extent to which faster fibre conduction velocities were affected by lead remains unclarified in this study. Further studies with lead workers free from zinc exposure might disclose the relative strength of lead effects on the faster and slower fibres. These studies by the use of DCV analysis might terminate one of the long lasting controversies over the effects of lead on peripheral nerve conduction velocity—that is, whether or not the conduction velocity of the slower nerve fibres is more sensitive to lead than that of the fastest fibre.1 9 26 Similarly, we have no convincing explanation to offer for the difference in the strength of lead effects between faster fibre conduction velocities and SCVs.

A few investigators have failed to find significant reductions in the conventional nerve conduction velocities in workers with asymptomatic increased lead absorption.3 6 In at least one of their reports,3 however, there was simultaneous exposure to lead and zinc. It is thus suggested that investigators estimate the possible antagonistic effect of zinc when no reduction in nerve conduction velocities is found in workers exposed to lead.

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