Is vinclozolin a reproductive hazard to men?

Zober et al examined fertility, hormones, and offspring sex ratio of men exposed to the fungicide vinclozolin. These authors were inclined to dismiss the possibility that it has any deleterious effect on the male reproductive system. I should like to suggest in contrast, that its effects in two respects resemble those of the nematocide dibromochloropropane (DBCP), which lowers sperm count and is associated with a highly significantly lowered offspring sex ratio.1

Zober et al report that although the testosterone concentrations of exposed men were unchanged their follicle stimulating hormone (FSH) concentrations were significantly higher than those of controls (P = 0.004). Similarly, increased FSH and unchanged testosterone were found in men exposed to DBCP.2 Increased gonadotrophin, or lowered testosterone concentrations, or both are characteristic of many illnesses in men.3

The offspring of men exposed to vinclozolin numbered 44 sons and 51 daughters.4 Compared with an expected Caucasian live birth sex ratio (proportion male) of 0.515, these figures yield a chi² of 1.05 (P = 0.15, one way). So although not formally significant, these results are suggestive. I suggest that further study of the possible effects of vinclozolin on the male reproductive system is merited. Its known similarities to an established hazard, DBCP, are disturbing. Especially interesting (and easy to gather) would be the sex ratios of offspring of men exposed to vinclozolin.

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As was pointed out by James, the mean value of the serum FSH measurements was significantly higher in the exposed than in the control group (P = 0.004 both before and after exclusions based on known pre-exposure health conditions) and this would suggest a possible causal relation between exposure to vinclozolin and FSH. For both luteinising hormone (LH) and FSH, non-significant but negative dose-response trends were found relative to current intensity as well as estimates of cumulative dose of vinclozolin exposure (see fig 4 of original paper for scatter plot of FSH and current intensity measure). Also, the man in the exposed group whose FSH level of 14.0 mU/ml was at the upper limit of normal had unilateral testicular atrophy only found to have been present since childbirth. All other readings among exposed men were within the reference range. From toxicological studies on vinclozolin and studies on other agents that block the testosterone receptor such as flutamide, the pattern of gonadotrophins at effective dose rates typically consists of increased testosterone, greatly increased LH, and marginally increased FSH concentrations.5 In the absence of dose-response and any abnormal FSH findings within the restricted exposed group and the lack of similarity between observed gonadotrophin patterns and what would be expected for an antianabolic agent, it is difficult to conclude that our findings are indicative of an effect related to vinclozolin.

Because the medical histories reported in table 3 of our paper did not distinguish between births occurring before and after first exposure, we reviewed the medical and work history data of all study group participants to find the number and sex of children born nine or more months after the date of the first exposure. Within this restricted observation period, there were 31 births, 13 male and 18 female, reported by 21 members of the study group. The number of births per family unit ranged from one to three. There were eight families with more male than female births, two with more female than male births, two with the same sex ratio, and two with male and female births of the same sex. Most probably some people are more sensitive to the psychosocial factors mentioned than others.6

Spurgeon et al did not pay attention to one point. If anxiety and dissatisfaction affect the perception of health, there might be a comparable effect on the perception of environmental factors. If so, negative or anxious feelings may also intensify the reporting of complaints and this would be consistent with our hypothesis that feelings of anxiety or dissatisfaction may lead to increased reporting of complaints about occupational or environmental hazards.7


5 Authors reply—in his letter, James raises questions about two specific health outcome measures reported in our paper, namely, serum follicle stimulating hormone (FSH) and sex of children born to fathers in the study group, and suggests that our findings for vinclozolin are similar to those reported by others for dibromochloropropane (DBCP), a known reproductive toxin in men.

Modifiers of non-specific symptoms in occupational and environmental syndromes

Editor—In a thorough review Spurgeon et al concluded that various occupational or environmental factors may lead to an increase of non-specific symptoms such as headache, tiredness, irritability, and sickness when, for example, people are exposed to various physical agents in the workplace. This sickness may be perceived as health hazards. Dissatisfaction with specific aspects of work may stimulate the occurrence of a similar pattern of health complaints. The authors discussed the role of individual and group factors in the perception of health and the tendency to report symptoms. Most probably some people are more sensitive to the psychosocial factors mentioned than others.

Spurgeon et al did not pay attention to one point. If anxiety and dissatisfaction affect the perception of health, there might be a comparable effect on the perception of environmental factors. If so, negative or anxious feelings may also intensify the reporting of complaints about aspects of work in a non-specific way.

Many studies in the field of occupational epidemiology are at least partly based on self-reported health or work related exposures. It is obvious that the tendencies mentioned may lead to biased results. Any particular problem that crops up in a work situation may generate anxiety or dissatisfaction, which in turn stimulates the tendency to report non-specific symptoms and complaints. Assuming that this tendency is stronger in some people than in others, various associations between exposure and effect can be expected. This is an additional reason to explore how psychosocial factors may colour the reporting of complaints. Two topics should therefore be added to the recommendations for future research as formulated by Spurgeon et al.

(1) Empirical evidence should be searched for our hypothesis that feelings of anxiety or dissatisfaction may lead to increased complaints about (probably non-specific) aspects of work. Research done on
the sick building syndrome suggests that this may apply to both psychosocial and organisational factors as well as physical characteristics of work. 1

(2) A valid method should be developed for the assessment of the psychosocial factors which seem to affect the perception and reporting of symptoms and complaints, to evaluate their influence in a particular study, and—if possible—to correct for their effects.

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Author’s reply—We thank the correspondents for their comments on our paper, and fully agree. The point that psychosocial factors may also affect awareness, and thus reporting of possible hazards in the workplace is well taken and supported by the literature. It was our intention that these aspects should be considered within the psychosocial pathway, particularly in relation to contextual factors, attitudes, beliefs, etc.

We also endorse the view that there is a need for a valid method to assess the influence of psychosocial factors on symptom reporting in relation to any particular hazard. This is particularly important in the workplace where the results of any such assessment should help to determine the focus of any prevention and control strategies. In many cases this focus may turn out to be psychosocial rather than, for example ergonomic, physical, or chemical.

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9th International conference on occupational respiratory diseases, 13-16 October 1997. Kyoto International Conference Hall, Japan

The Conference is organised by the Japanese National Organising Committee for the Ninth International Conference on Occupational Respiratory Diseases, in collaboration with the Ministry of Labour of Japan, the International Labour Office and the Japan Industrial Safety and Health Association.

This Conference provides an excellent opportunity for scientists, health practitioners, hygienists, engineers, management workers, and legislators to share experiences and ideas on the management and prevention of occupational respiratory diseases and to set priorities for the next century. The success of the meeting will depend upon contributions of papers and exhibits by leaders in the several disciplines of occupational health. We particularly encourage submission of abstracts by junior scientists, as well as from senior investigators.

Themes:
- Epidemiology of occupational respiratory diseases
- Health surveillance of workers exposed to respiratory hazards
- Aetiology, pathogenesis, diagnosis and treatment of occupational respiratory diseases
- Health hazard assessment by environmental and exposure monitoring
- Control measures against health hazards at the workplace
- Respiratory protective equipment
- Information, education and training on occupational respiratory diseases

Working Groups:
- ILO international classification of radiographs of pneumoconioses
- Global action on elimination of silicosis
- Occupational respiratory allergies
- Natural and synthetic fibres
- Relationship between occupational respiratory diseases and lung cancer

Further information from: International Communications Specialists, Kashi Building, 2-14-9 Nihombashi, Chuo-ku, Tokyo 103, Japan.

International symposium on environment, lifestyle, and fertility, 1-5 December 1997, Scandinavian Congress Center, Aarhus, Denmark

The symposium is organised within the framework of Asclepios (a European concerted action on occupational hazards to male reproductive capacity).

Two European concerted actions have been launched with the objective of mapping the European occurrence of infertility and identifying occupational hazards to male reproductive function. Dozens of national studies in Europe, the United States, and throughout the world within reproductive epidemiology have provided additional knowledge to add to agreed recommendations for future research. There is a need to identify environmental risk factors as well as suspected factors with zone or limited significance for infertility and there is a need to identify studies and methods to be recommended and not recommended.

The symposium will include the following themes of the art reviews:
- The use of time to pregnancy in a demographic and epidemiological perspective: Alfred Spira
- Design and bias issues related to studies of subfertility: Jan Olven
- Validity of time to pregnancy data in men and women: Michael Joffe
- Semen quality as marker of fecundity in epidemiological studies: Jens Peter Bonde
- Endocrine markers of male fecundity: Richard Sharpe
- Impact of lifestyle and social factors: Nel Roedelev
- Occupational and environmental impact on fertility: Steen Schrader

The symposium is open for free communication as well as posters and the poster session with plenary review will also be organised. The symposium language is English. Deadline for registration and submission of papers is 1 September 1997.

Further information from: Asclepios, Pia Pont/Jane Boilesen, Department of Occupational Medicine, The Steno Center of Public Health, Aarhus University Hospital, Noerrebrogaard 37-39, DK-8000 Aarhus C, Denmark. Phone: +45 8949 4294; Fax: +45 8949 4260; E-mail: akh gp22s skamlop kaa.dk

NOTICES

17th Annual New England epidemiology summer programme (5 and 10 day courses) June 9-July 4 1997. Tufts University, USA

The New England Epidemiology Institute summer programme at Tufts University’s Medford campus includes methodological, statistical, and substantive courses. This programme is intended for those seeking an introduction to modern epidemiological concepts as well as those desiring a review of recent developments in epidemiological thinking.

Twenty five and 10 day courses cover the following: introduction to epidemiology, epidemiologic basis for causal inference, biostatistics for epidemiologists, clinical research, theory and practice of epidemiology, pharmacoepidemiology, meta-analysis, epidemiology in public health practice, clinical trials, survival analysis in epidemiology, decision and cost effectiveness analysis in health care, the biology and epidemiology of cancer, health care use and outcomes research, regression modeling in epidemiology, perinatal epidemiology, occupational and environmental epidemiology, molecular epidemiology and the use of biomarkers, ethnics and epidemiology, and writing for publication in a scientific journal. Invited faculty include excellent teachers and prominent researchers from leading universities. Registrants may receive graduate-degree credit or continuing education credits from Tufts University, continuing medical education (AMA category 1) through Tufts University Medical School, and certification maintenance from the American Industrial Hygiene Association.

Further information from: The New England Epidemiology Institute, Department PA-OBM, One Newton Executive Park, Newton Lower Falls, MA 02162-1450, USA. Phone: (617) 244-1200; Fax: (617) 244-9669; E-Mail: epidemiol@axol.com; World Wide Web Page: http://www.epidemiology.com

15th Asian conference on occupational health 31 August-3 September 1997. Kuala Lumpur, Malaysia

The theme of the conference is preparing for occupational health and safety needs of the next decade.

For more information on registration and presentation of papers, free communications, and posters at the conference please contact: Conference Secretariat, Society of Occupational & Environmental Medicine of MMEA, 4th floor, MMA House, 124 Jalan Pahang, 53000 Kuala Lumpur, Malaysia. Tel: (60) (3) 4431740/4431743; Fax: (60) (3) 4481817.

INTERNATIONAL SYMPOSIUM ON
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1-5 December 1997, Scandinavian Congress Center, Aarhus, Denmark

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