

PHYSICAL ACTIVITY AND RISK OF CANCER

A population based cohort study including prostate, testicular, colorectal, lung and breast cancer

Inger Thune

Tromsø 1997



**Institute of Community Medicine
University of Tromsø**



The Norwegian Cancer Society

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All parts of the body which have a function, if used in moderation and exercised in labors in which each is accustomed, become thereby healthy, well developed, and age more slowly, but if unused and left idle they become liable to disease, defective in growth, and age quickly.

Hippocrates, 460-377 BC.

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Tromsø, February 1997

Inger Thune

LIST OF PAPERS

This thesis is based on the following papers, referred to in the text by their Roman numerals

- I. Thune I, Lund E. Physical activity and the risk of prostate and testicular cancer: a cohort study of 53,000 Norwegian men. *Cancer Causes Control* 1994;**5**:549–556
- II. Thune I, Lund E. Physical activity and risk of colorectal cancer in men and women. *Br J Cancer* 1996;**73**:1134–1140
- III. Thune I, Lund E. The influence of physical activity on lung cancer risk. A prospective study of 81,516 men and women. *Int J Cancer* 1997;**70**:57-62
- IV. Thune I, Brenn T, Lund E, Gaard M. Physical activity and risk of breast cancer. *N Engl J Med* 1997;**336**:1269-1275
- V. Thune I, Njølstad I, Løchen M-L, OH Førde. Physical activity improves the metabolic risk profiles in men and women. A seven year follow-up study with repeated assessments of leisure time activity: The Tromsø Study. *Submitted for publication.*

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1 INTRODUCTION

During my childhood I was always instructed that I should be physically active in order to take care of my body, and this continued into my adulthood. The positive psychological and physiological advantages of physical activity were always emphasized. During my time at medical school and later through my contact with cancer patients, I started to wonder whether there is any relationship between physical activity and risk of cancer. In 1987, I came across an article by Frisch and colleagues¹ who had observed a lower prevalence of cancer of the breast and reproductive organs among former college athletes than among non-athletes. On completion of my medical internship, I considered whether there was any possibility of investigating further the relationship between physical activity and cancer risk. At that time, very few studies had elucidated this relationship, in contrast to many studies that were investigating the association between physical activity and cardiovascular diseases.

Strengthened by the belief that physical activity may influence cancer risk I applied for grants from the Norwegian Cancer Society in 1990. During my preparation for the application (Spring 1990), a search on Medline revealed that there were only seven studies world wide that focused on physical activity and cancer risk; this has now increased to 275 articles (Medline - December 1996).

Physical activity

Physical activity has been an important factor in the evolution of the modern human, *Homo sapiens*, 35 000 years ago.^{2,3} The gene pool from which humans currently derive their individual genotypes was formed during a period of over a billion years of evolution.⁴ Through a lifestyle of hunting and gathering, our genetic constitution was selected. The hunter-gatherers' way of life involves endurance activities with peak bouts of strenuous physical activity that involve considerable heat production. These patterns of physical activity continued mainly through the shift to agriculture 10,000 years ago. Such physiological adaptations suggest the importance of endurance activities in our evolutionary past; evaluation of recent preliterate populations confirms that their daily activities would have developed superior aerobic fitness.⁵ The cultural changes, particularly those resulting from the Industrial Revolution, have outpaced any genetic adaptation. Consequently, a sedentary Western lifestyle could be giving modern *Homo sapiens* problems because we still carry genes for a physically active way of life. Therefore, from a genetic standpoint, humans of today are still hunters also reflected by that skeletal muscles constitutes some 40% of the body mass. The capacity to be active, whether defined by total energy expenditure or by intensity of effort, requires a state of physiological fitness. Thus, if we are sedentary and not physically active,

we deteriorate. If we are active, physical activity alters the effects of other influences on our health: our cardiovascular-respiratory physiology, our musculoskeletal strength, our gastrointestinal function, and even our state of mind. Rammazzini⁶ made an extraordinary observation nearly 300 years ago: sitting tailors were more sickly and paler than scurrying messengers. Thus, regular exercise is essential for optimal function of the body.⁷

Physical activity is a complex behaviour; it can be defined as 'any bodily movement produced by skeletal muscles that results in energy expenditure'.⁸ Exercise is a subset of physical activity defined as 'planned, structured, and repetitive bodily movement done to improve or maintain one or more components of physical fitness', whereas physical fitness is 'a set of attributes that people have or achieve that relates to the ability to perform physical activity'.⁸

The general view is that there must be a minimum exercise intensity necessary to stimulate any improvement in physical fitness. This minimum was exemplified recently as amounting to a minimum of exercise intensity of at least 50% of the maximum oxygen uptake or at least 60% of the maximum heart rate.⁹ An alternative to this minimum is the accumulation of total energy expended in exercise over a certain timespan, including not only intensity of exercise.¹⁰ This could mean that even less time than the recommended 30 minutes of moderate physical

activity daily may be enough when either the intensity is high or made up of shorter bouts with accumulated duration, whereas longer duration may be necessary for low exercise intensity to increase physical fitness.¹⁰

An individual's propensity to be physically active may be inherited,¹¹⁻¹² although sociocultural factors are supposed to be of greater importance in physical activity behaviour than genetic factors¹² and the genetic component has been estimated to cover about 30% of the aerobic capacity.¹² Men in the lower social classes have been observed to spend less leisure time being active and to be more active at work than men in the higher classes.¹³

Assessments of physical activity

In any scientific work, it is necessary to have precise, reliable and practical assessment of the variable being studied. More than 30 different techniques have been employed for assessing physical activity in population studies¹⁴ and these have been described in different categories.¹⁴⁻¹⁵

A Direct

1 Questionnaire assessment

The two data collection methods used in relation to physical activity questionnaires¹⁶ are interview¹⁷⁻¹⁸ and self-administration.¹⁹⁻²⁰ The participants report varying degrees of detailed recall of activity, ranging from a recall of physical activity during one day,²¹⁻²² to specific

activity over the last week²³⁻²⁴ to recall of specific activities during both work and leisure time over the past year^{20, 25} or during different periods in life.²⁶ Questionnaires are practical means of measuring physical activity of large populations and are therefore widely used in population studies.²⁷⁻²⁸

Assessment of physical activity by questionnaires in different clinical and epidemiological studies has resulted in both similarities and differences across published systems. As a consequence of differences that limit the comparability of results, Ainsworth et al.²⁹ developed a *compendium of physical activities* for calculating energy expenditure related to one specific activity, to facilitate the coding of physical activities and to promote comparability of coding across studies. All activities are assigned an intensity unit based on their rate of energy expenditure, expressed as METs where one MET is the resting metabolic rate. It is also defined as the energy expenditure for sitting quietly; for the average adult this is approximately 3.5 ml oxygen per kg body weight per min or 1 kilocalorie (kcal) per kg body weight per hour. Only data for adults are included in these calculations. In this way a physical activity index for estimating the total energy in kilocalories expended each week in activity can be obtained, by multiplying the MET score by the duration of workout, body weight in kilograms and frequency per week.

2 Diary surveys

These give detailed information about physical activity on a specific day.¹⁴ This method has seldom been used in epidemiological studies of physical activity, although it has been used in energy balance studies. A diary may be used to compare other estimates of total daily energy expenditure, i.e. through caloric intake,²³ or in validation of a physical activity questionnaire.²³ This method is precise but time-consuming and expensive. In addition, a participant's activity pattern may easily be altered as a result of the recording process.

3 Mechanical and electronic monitoring

Heart rate may reflect both intensity and duration of physical activity. It has therefore been used indirectly as a measure of physical activity in validating surveys of physical activity questionnaires.³⁰⁻³¹

Motion sensors provide measurements of 'movements', primarily through pedometers, and record the acceleration and deceleration of movements. This instrument has mainly been used on a small scale, but recently Sequeria et al.³² observed that it was important for comparison in a questionnaire of a population survey.

4 Behavioural observation

This has been developed by behaviourists, but is impractical in population studies.¹⁴

B Indirect

1 Calorimetry

This is measured directly through the production of heat or indirectly through the consumption of oxygen; these correlate closely with heat production but have little use in assessing physical activity in a general population.¹⁴ Resting energy expenditure, as determined by indirect calorimetry, has, however, been used in validation studies of physical activity questionnaires.³³

2 Caloric intake

This may be an estimate of energy expenditure and, hence, of physical activity, if one assumes that energy balance has been achieved with stable body weight. This method has therefore been used in validation of physical activity questionnaires^{23, 33} However, dietary measures of physical activity are unable to identify the types, frequency, intensity or duration of physical activities.

3 Physiological markers of physical activity

Physical fitness assessment

Maximal oxygen uptake Vigorous physical activity has an influence on cardiorespiratory endurance. Hence, maximum oxygen consumption has frequently been used for estimating physical fitness^{9, 11, 34} and validating self-reported physical activity.³⁵⁻³⁶

Maximal or submaximal work capacity This test can be performed on a bicycle that has an initial workload, with increments being

made after a certain time. Physical fitness is defined as the maximum workload possible.³⁷⁻³⁸

Heart rate Both heart rate measured in a graded exercise test²⁴ and that at rest³⁰ have also been used as a surrogate measure of physical activity; this has the advantage of no recall or reporting bias.

Other assessments

Various others methods have been used to assess cardiorespiratory fitness, including the duration of a graded, submaximal or maximal exercise test.³⁹ In a technique using doubly labelled water,¹⁴ energy expenditure over time can be measured. Participants are given water containing isotopically labelled hydrogen and oxygen atoms to drink. Then the relative proportions of metabolized and non-metabolized water provide an overall estimate of energy expenditure. In field testing, a method in which the participants are asked to walk or run for a certain time or distance, physical performance capacity is converted to aerobic capacity.⁴⁰

4 Sports and recreational participation

This has the possibility of distinguishing between high and low levels of a specific activity; it was used by Frisch and colleagues when they compared non-college athletes with college athletes, in relation to the risk of life-time occurrence of breast and reproductive cancer.¹ Limitations are selection bias and

omission of occupational physical activity, among others.

5 Job classification

Ranking jobs according to levels of physical activity has been used in many studies.⁴¹⁻⁴³ There are several limitations, because of within-job classification, selection bias and omission of leisure time activity. In addition, the variability between job categories is declining as a result of the decline in manual work.

Physiological effects

Energy balance-weight control

Physical activity correlates strongly to weight gain in the general adult population.²⁸ Differences in physical activity represent the largest source of variability in energy requirements, both within and between individuals. The primary components of total daily energy expenditure include resting (basal) metabolism (approximately 50-75%), physical activity (15-40%) and thermic effects of food (< 10%).⁴⁴ Basal metabolism is an almost linear function of lean body mass and is the energy expenditure measured in a resting subject after an overnight fast; it approximates to the minimum energy expenditure necessary for maintenance of critical body functions (i.e. internal work).

Total and specific cancer incidence correlated fairly well with body weight in an ecological study of 24 populations⁴⁵ and increased body mass index has been observed as a

risk factor for colon cancer^{17, 46} and breast cancer.⁴⁷⁻⁴⁸ Weight gain during adult life has been observed to be a predictor of breast cancer risk.⁴⁸⁻⁴⁹

Physical activity also influences the net available energy, and experimental studies have demonstrated that calorie restriction inhibits mammary⁵⁰⁻⁵¹ and colon carcinogenesis⁵² and reduces proliferative activity in rodent mammary glands.⁵³ Consequently, any change related to energy balance can potentially disrupt the steady state of energy and macronutrient balance, inducing weight fluctuations, which are both of importance in carcinogenesis.

Bowel transit time

The normal transit time through the colon shows considerable variation among individuals⁵⁴ and between the sexes.⁵⁵ Physical activity can reduce gastrointestinal transit times.^{54, 56} Consequently, exposure time of the colon mucosa and potential carcinogens in the faecal stream may be decreased by exercise.

Hormonal levels

Levels of cyclic oestrogen and progesterone seem to be related to risk of breast cancer.⁵⁷ Physical activity has been observed to reduce the cumulative exposure to both cyclic oestrogen and progesterone in women⁵⁸⁻⁵⁹ and to influence testosterone level in men.⁶⁰⁻⁶¹ Hard training and moderate leisure activity may both decrease oestradiol and progesterone secretion,⁶²⁻⁶³ as well as inducing anovulation^{59, 63} or causing

secondary amenorrhoea.⁶⁴

High levels of testosterone have been observed to be important in the prostate cancer risk,⁶⁵⁻⁶⁷ and also levels of testosterone within normal endogenous ranges have been associated with an increased risk of prostate cancer.⁶⁷ Athletes have been shown to have lower levels of testosterone⁶⁸⁻⁶⁹ and post-exercise levels of testosterone may be temporarily lower.^{60, 70}

Lung function

Physical activity improves pulmonary capacity. A measure of pulmonary function, the forced expiratory volume in one second adjusted for height (FEV₁/height), correlates positively with strenuous physical activity and duration of exercise.⁷¹⁻⁷² Increased pulmonary ventilation and perfusion could reduce the interaction time and concentration of any carcinogenic factor in the airways.

Trauma

Physical activity may induce injury. Regeneration and increased cell division have been hypothesized as a cause of human cancer.⁷³ Trauma has been associated with increased risk for testicular cancer⁷⁴ and intracranial meningiomas.⁷⁵

Immune response

In 1902, after the Boston Marathon, a leucocytosis had already been found in a small group of runners.⁷⁶ Recent studies have revealed that immune parameters are altered after an acute bout of physical activity⁷⁷⁻⁷⁸ or result from a long-term effect⁷⁹⁻⁸⁰ of

physical activity. Moderate exercise training has been observed to increase serum globulins⁸¹ and enhance natural killer cell activity⁷⁹⁻⁸⁰ in trained subjects. In contrast, over-training may decrease both the resting level of immune function and the responses to acute exercise stress.⁸² Exercise also results simultaneously in many other systemic changes, such as changes in neuroendocrine function and increased blood flow. Does physical activity *de novo* therefore influence the immunity or does it occur only through other physiological parameters? Recent studies may indicate a relationship of physical activity and natural immunity that is important for cancer risk.⁸⁰

Serum lipids

Numerous cross-sectional and interventional studies have observed lower concentrations of total cholesterol and triglycerides⁸³⁻⁸⁵ and higher concentration of high-density lipoprotein-cholesterol (HDL-cholesterol)⁸⁵ in physically active compared with inactive individuals. Triglycerides are known to displace oestradiol from its tight binding to sex hormone-binding globulin (SHBG), thus increasing free oestradiol. SHBG is found in low levels in obese women.⁸⁶

Other factors

One of the most striking effects of increased physical activity, cardiovascular adjustments, may result in an increased capacity for local and central blood flow. This may in turn induce an increase in

transport and removal of possible carcinogenic agents from the blood and tissues. Further high levels of physical activity may increase sensitivity to insulin⁸⁷; recently, this has been suggested as important in carcinogenesis of the colon⁴⁷ and the breast.⁸⁸

Cancer incidences, prognoses and geographical variations

Prostate cancer is the most frequent cancer among men in Norway, contributing 22.7% (2,236 cases) of all cancer cases diagnosed in men in 1993.⁸⁹ Comparing the age-adjusted incidence rate during 1954–58 with that of 1989–93, the incidence rate increased from 26.3 to 46.6 per 100,000 person-years. Among those diagnosed, 27.1% of the cases in 1993 were under the age of 70. The national 5-year survival rate for all ages and stages combined was 58% in 1986–90.⁸⁹ There has been an increase in cause-specific mortality, signifying a genuine increase in incidence over time, not only resulting from increased detection rates.⁹⁰ Geographical differences in incidence rates world wide demonstrate that Asia is a low-risk area, and North America and Scandinavia are high-risk areas.⁹¹ The age-adjusted incidence rate during 1989–93 varies in the geographical regions of Norway studied, with Oslo being the high-risk area with 52.4 per 100,000 person-years and Finnmark the low-risk area with 34.2 per 100,000 person-years. Although

aetiological factors are mainly unknown, these observations point to a potential role for lifestyle factors in the carcinogenesis of prostate cancer.

Testicular cancer was diagnosed in 1.9% of all cancer cases in 1993 ($n = 189$).⁸⁹ There has been a 2.5 times increase in the annual age-adjusted incidence rates in Norway in all age groups, from 3.3 during 1954–58 to 8.2 cases during 1989–93 per 100,000 person-years. In 1986–90, the national 5-year survival rate for all ages and stages combined was 95%.⁸⁹ Both the national⁸⁹ and the worldwide geographical variation,⁹² combined with the increase in incidence over time, indicate that environmental factors could explain these patterns.

Colon cancer was diagnosed among 9.5% of cancer cases in 1993 (n for men = 817, n for women = 980).⁸⁹ The age-adjusted incidence rate increased twofold in both sexes from 1954–58 to 1989–93: from 11.4 to 23.8 per 100,000 person-years in men and from 11.0 to 20.0 per 100,000 person-years in women. Among those diagnosed in 1993, 39.5% and 33.4% of the cases were diagnosed under the age of 70 in men and women, respectively. The age-adjusted incidence rate during 1989–93 varies for the geographical regions studied, because Sogn og Fjordane and Oslo have 2.3 times higher incidence rates than Finnmark for both sexes.⁸⁹ Migration studies and geographical variation world wide points to environmental factors of importance to explain the increase in incidence rates world wide.^{91,93-94} In

1986-90, the national 5-year survival rate for all ages and stages combined was 49% and 53% in men and women, respectively.⁸⁹

Rectal cancer was diagnosed in 5.1% of all new cancer cases in 1993 (n for men = 533, n for women = 429).⁸⁹ The age-adjusted incidence rate increased from 1954-58 to 1989-93: from 7.5 to 14.9 per 100,000 person-years in men and from 5.1 to 10.0 per 100,000 person-years in women. Among those diagnosed in 1993, 42.8% and 41.3% of the cases were diagnosed under the age of 70 in men and women, respectively. In 1986-90, the national 5-year survival rate for all ages and stages combined was 48% and 53%, in men and women, respectively.⁸⁹

Lung cancer was diagnosed in 9.2% of all new cancer cases in 1993 (n for men = 1,216, n for women = 536) and is the second most frequent cancer among men and the third most frequent in women in Norway.⁸⁹ The age-adjusted incidence rate increased from 1954-58 to 1989-93: from 11.8 to 35.3 per 100,000 person-years in men and from 3.1 to 13.7 per 100,000 person-years in women. Among those diagnosed in 1993, 51.8% and 58.4% of the cases were diagnosed under the age of 70 in men and women, respectively. There has recently been an observed shift towards adenocarcinoma as the most frequent histological type.⁹⁵ In 1986-90, the national 5-year survival rate for all ages and stages combined was 9% and 10% in men and women, respectively. The age-adjusted incidence rate during 1989-93 varies

for the geographical regions studied, with Finnmark and Oslo being the high-incidence areas (41.1 and 19.6 per 100,000 person-years in men and women, respectively in Oslo), whereas Oppland is a low incidence area with 28.4 and 8.9 per 100,000 person-years in men and women respectively. World wide, lung cancer is by far the most common cancer of men.⁹¹

Breast cancer is the most frequent cancer among women in Norway, contributing 22.3% of all cancer diagnosed in women in 1993 (n = 2,035).⁸⁹ Comparing the age-adjusted incidence rate from the period 1954-58 with that of 1989-93, it increased from 43.6 to 58.6 per 100,000 person-years. Although 24% of the cases in 1993 were diagnosed under the age of 50, 61.9% of the cases in women were diagnosed under the age of 70. In 1986-90, the national 5-year survival rate for all ages and stages combined was 76% in women. Breast cancer incidence rates varied in the geographical areas studied, with Finnmark being a low-incidence area,⁸⁹ the incidence rates varies among countries with a four- to sevenfold higher incidence rate in Western countries than in Japan.⁸⁹ This points to modifiable factors that are related to lifestyle.

2 AIMS OF THE THESIS

- The overall aim of this thesis was to elucidate any association between physical activity and the subsequent risk of some major types of cancer in the general adult population of Norway: prostate, testicular, colorectal, lung and breast cancer.
- Another aim was to examine whether physical activity at work had a different association to these cancer types than physical activity in leisure time.
- To investigate whether age, sex, body mass index and hormonal status (pre- and postmenopausal), on one side, and lifestyle-related factors such as smoking and dietary factors, on the other, strengthen or reduce these observed associations.
- Could repeated assessments of physical activity induce stronger or weaker risk estimates?
- If any associations between physical activity and risk of cancer were observed, then these associations were investigated to ascertain if these could indicate a causal relationship.
- To study if changes in physical activity or sustained physical activity influence metabolic profiles to any large extent, which would support physical activity as a potent biological mediator of importance in reduction of risk for chronic diseases, e.g. cancer?

3 SUBJECTS AND METHODS

This thesis is based on population-based health surveys, originally used in screening for cardiovascular diseases, which were carried out in five geographical areas in Norway: three counties (Finnmark, Oppland and Sogn og Fjordane) and two cities (Oslo and Tromsø). The participants in this thesis were born between 1922 and 1959. The first surveys were carried out between 1972 and 1978 with repeated surveys between 1977 and 1987 in the three counties and one city.

The study population

Papers I, II, III and IV

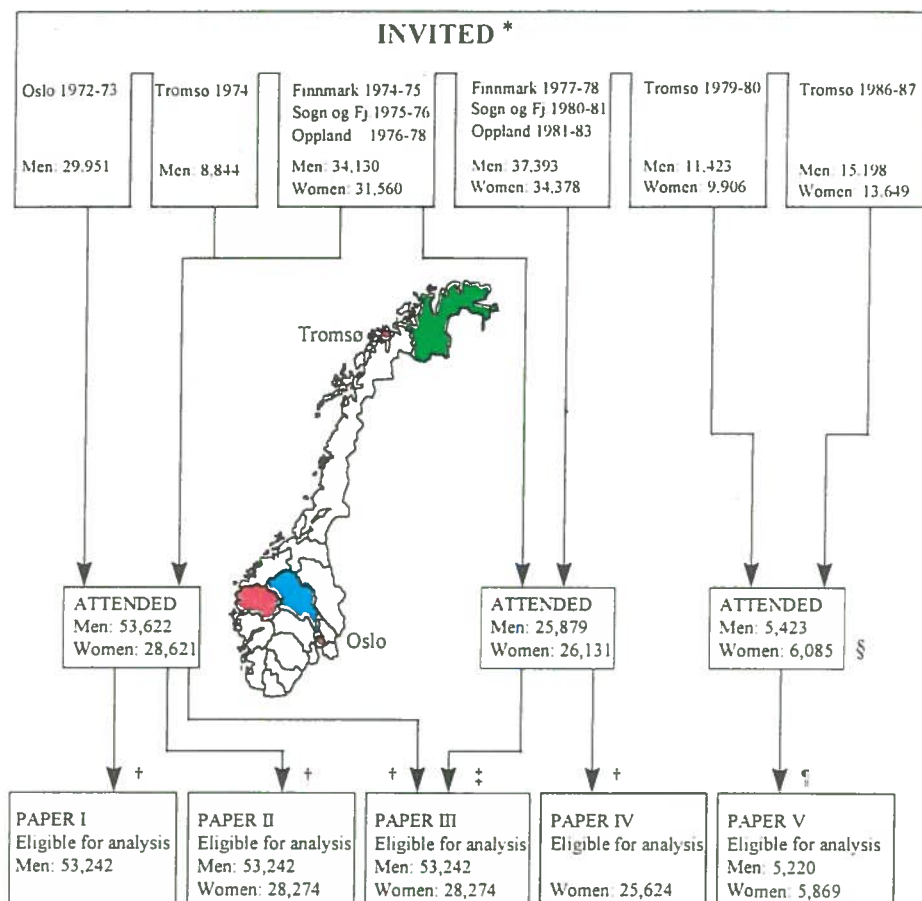
A total of 72,925 men and 31,560 women, who were residents of five geographical areas of Norway (three counties of Finnmark, Oppland and Sogn og Fjordane, and the two cities of Oslo and Tromsø) were invited to participate in a population-based health survey between 1972 and 1978 (Fig 1). In the three counties (Finnmark 1974–75, Sogn og Fjordane 1975–76 and Oppland 1976–78), all men and women aged 40–49 years, and a random sample of people aged 20–34, were invited. In four small municipalities in Finnmark, all men and women aged 20–34 were invited.⁹⁶ In Tromsø in 1974,⁹⁷ all men aged 20–49 were invited, whereas in Oslo 1972–73, men aged

40–49, plus a 7% random sample of men aged 20–39, were invited.²⁵

If attendance is defined as being registered as participating, 53,622 men (attendance rate = 73.5%) and

28,621 women (attendance rate = 90.7%) attended these surveys. Paper I comprises 53,622 men only, whereas papers II and III include both sexes.

Fig. 1 The number of people invited* and eligible for analysis in papers I–V.



* Invited; included persons who appeared without invitation

† Excluded persons with a pre-existing malignancy or got a malignant disease within the first year

‡ Subcohort lung

§ Included persons aged 20–49 years at entry 1979–80 and participating at both surveys (1979–80 and 1986–87)

¶ Excluded persons with previous myocardial infarction, stroke, diabetes and those with missing information about leisure time activity

In the counties of Finnmark (1977), Sogn og Fjordane (1980) and Oppland (1982–83), 3–5 years after the first survey men and women were invited to a similar second health survey.⁹⁸ Men and women who attended both surveys in the three counties (men: $n = 25,879$; women: $n = 26,131$) represent a subcohort in paper III. A food-frequency questionnaire was given to those men and women who attended the second survey, to be completed at home and returned by mail. This food-frequency questionnaire was returned by 26,090 men and 25,892 women. Women participating in both surveys in the three counties provided the study population for paper IV.

Paper V

Paper V included men and women who were residents of Tromsø and who participated in two population surveys carried out in the municipality of Tromsø in the 1979–80 and the 1986–87 surveys.⁹⁹ In the 1979–80 survey, men aged 20–54 years and all women aged 20–49 years were included and this made up a total of 21,329, of whom 16,621 attended (78%). The total number of individuals examined at the 1986–87 surveys was 21,826, 81.3% of the eligible population. Men ($n = 5,423$) and women ($n = 6,085$) aged 20–49 years in 1979–80 and who attended both surveys were included in paper V.

Screening procedures

Surveys 1972-78

The screening procedures for the first surveys were almost identical in the five geographical areas and are described in detail elsewhere.^{25, 96-97} Each person was initially contacted by mail with a cover letter and a one-page questionnaire on the reverse side (Appendix I). The participants were asked to answer the questionnaire at home and bring it to the screening examination, which then included the following elements used in the present thesis.

I A questionnaire which comprised:

A – History of cardiovascular disease, diabetes and treatment for hypertension

B – Symptoms possibly caused by coronary or peripheral atherosclerosis

C – Physical activity during leisure time

D – Smoking habits

E – Conditions at work (physical activity) and stress in social life

F – Ethnic origin (Tromsø and Finnmark)

G – Family history of cardiovascular disease (all counties and Tromsø)

II Measurement of height and weight performed by standardized methods

III A non-fasting blood sample.

The questionnaire was checked by trained nurses, and omissions and logical inconsistencies were corrected

according to a written protocol. While checking the questionnaire, the nurses also asked everyone about the time since the last meal and women about menstrual status and pregnancy.

Follow-up surveys

The Norwegian Counties

Those participants in the counties, Finnmark (1977–78), Sogn og Fjordane (1980–81) and Oppland (1981–83), still resident in the county, together with an additional random sample aged 20–39, were invited to a similar second survey 3–5 years later (1977–83). Each person was initially contacted by mail with a cover letter and a one-page questionnaire (almost identical to the first survey; see Appendix II). Each municipality was surveyed at the same time of year as in the first survey. All attendees were given a food-frequency questionnaire to be completed at home and returned by mail (see Appendix III).

Tromsø

In the municipality of Tromsø, the 1979–80 survey of the procedures were mainly the same⁹⁹ as in the 1974 survey.⁹⁶ The main questionnaire (see Appendix II) covered the same aspects as in 1974 and, in addition, each participant was given a second questionnaire (see Appendix IV) that they were asked to complete at home and return by mail. This questionnaire was a combined food-frequency questionnaire and a questionnaire about previous and present chronic diseases, other than those covered in

the first questionnaire, illness in parents and siblings, or psychosocial conditions. At screening, the main questionnaire was checked for inconsistencies by trained nurses; measurements of height, weight, blood pressure, heart rate and serum lipids have been presented (see paper V).

The 1986–87 survey was set up by the same institutions as those involved in the 1979–80 survey. The main questionnaire (see Appendix V) covered the same aspects as those in 1974 and 1979–80. In addition, each participant was given a second questionnaire (see Appendix V) that they were asked to complete at home and return by mail. At screening, the main questionnaire was checked for inconsistencies by trained nurses; measurements of height, weight, blood pressure, heart rate and serum lipids have been presented (see paper V).

The National Health Screening Service

As the service with the responsibility for tuberculosis screening since 1943, the State Mass Radiography Service, now called the National Health Screening Service (NHSS), organized the main parts of these surveys in the counties, except for the survey in Oslo; they used a mobile unit in collaboration with the County Medical Officers. They were also central in organizing the surveys in Tromsø in collaboration with the University of Tromsø and the Tromsø Health Council.

Reminder

No reminders were sent out in Oslo and, in the counties also no-one received any reminders, although the survey was backed up by the local newspapers, radio and the local health council nurse, all of whom co-operated.⁹⁶ In Tromsø one reminder was sent for each survey.

Ascertainments of variables

The main questionnaire, which also covered physical activity, was filled in at home by all subjects and checked at screening for inconsistencies; this meant that data were complete.

Participants were asked to answer the questions based on an average of performed physical activity over the last year. They marked 'yes' for the level that fitted best.

Leisure time physical activity was divided into:

- 1 Reading, watching TV or other sedentary activity
- 2 Walking, bicycling for at least 4 hours a week
- 3 Participating in recreational athletics for at least 4 hours a week
- 4 Participating in hard training or athletics competitions, regularly, several times a week.

Work activity was divided into:

- 1 Mostly sedentary work, e.g. office work, watchmaker
- 2 Work involving a lot of walking, e.g. shop assistant, light industrial work

3 Work involving a lot of walking and lifting, e.g. postal worker, heavy industrial work

4 Heavy manual labour work, e.g. forestry work, heavy farm work.

Physical fitness was measured in a subpopulation in Tromsø and has been described previously³⁷ (paper V). A graded submaximal or maximal bicycle exercise test, with pedalling frequency of 60 per min, was carried out in a random subgroup in the 1986-87 survey. The initial workload was set at 25 watts(W), with a 25 W increment every minute up to a maximum of 250 W after 10 minutes. The tests were stopped prematurely if exhaustion, or symptoms such as leg pain and angina,³⁷ made it necessary. Physical fitness was defined as the maximum possible work load.

Heart rate is derived from the median pulse-to-pulse interval during the measurement of blood pressure. Three recordings of heart rate were made at 2-minute intervals, and the lowest measurement recorded was used (see paper V).

A semi-quantitative food-frequency questionnaire was designed by a section for dietary research at the University of Oslo; it was used in Finnmark, Oppland and Sogn og Fjordane. The questionnaire requested information on the usual consumption of 80 items. In most of the items, amount or units consumed one each occasion were requested. Among 50 questions, 31 were sufficiently specific to enable an estimation of energy and fats according to the Norwegian Food

Composition table.¹⁰⁰ These comprised milk, potatoes, bread, spreads (cheese, meat, jam, salads), fats on bread, fat in cooking, a range of meat and fish meals, cakes, eggs, oranges, porridge, cod-liver oil and vitamin pills. The energy and fat intake for each woman was derived as the sum of all foods consumed.¹⁰¹

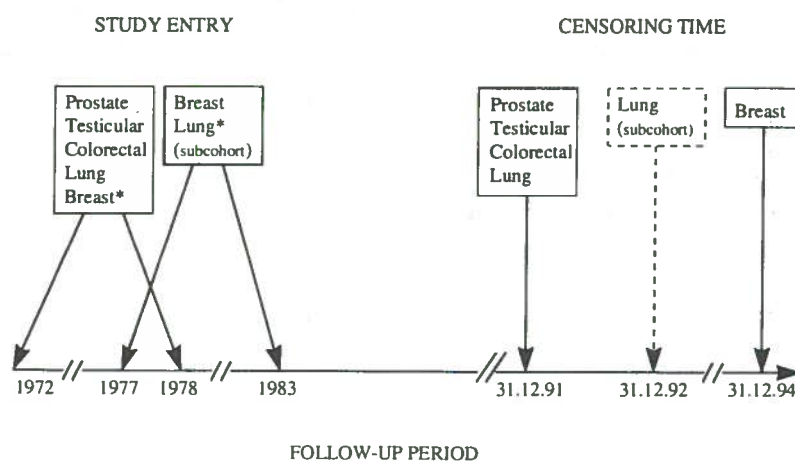
In Tromsø, participants answered a somewhat shorter food-frequency questionnaire comprising information about both type and quantity of bread, fruit and vegetables, type of fat used at table, the type of milk normally used, the amount of fat on each slice of bread, the number of glasses of milk and cups of coffee,

and the consumption of alcohol (see Appendices IV and V).

The questionnaire used in the Oslo study had pre-set groups for number of cigarettes smoked per day and packs of pipe tobacco smoked per week (see Appendix I). Otherwise, people were asked to give their average daily consumption rather than specify a pre-set group (see Appendix I).

Height and weight were measured in light clothing with no shoes. Height was measured to the nearest centimetre and weight to the nearest half kilogram, on regularly calibrated scales. A non-fasting venous blood sample was taken for analysis.^{25, 96-99}

Fig 2. The follow up period with study entry (assessment of physical activity) and censoring time of cancer (prostate, testicular, colorectal, lung and breast)



*analysing the effect of repeated assessment of leisure time physical activity

The Central Population Register at Statistics Norway has been recording the reproductive history of every woman, including date of first liveborn child, which was used in paper IV. The national 11-digit personal identification number enabled linkage to Statistics Norway.

Identification of cases

In Norway there is mandatory reporting of cancer by all physicians, hospital departments and via copies of pathological reports to the Cancer Registry of Norway. This ensures almost complete registration of incident cases of cancer.¹⁰² In addition, four times a year, all death certificates mentioning cancer are forwarded to the Cancer Registry of Norway. The national 11-digit personal identification number has enabled this linkage, resulting in close to 100% histological verification of every incident case of cancer of the prostate, testes, colon, rectum, lung and breast in the present studies.

Identification of death and emigration

In Norway all deaths are by law recorded through certification from physicians; they are further recorded at Statistics Norway. All emigrations are also recorded here, the national 11-digit personal identification number enabling linkage to Statistics Norway. Information about death and emigrations were used in papers I-IV when calculating the observation years at risk in these papers.

Statistical methods

All analyses were sex specific. Baseline characteristics were age adjusted and compared across levels of leisure and categories of work activity by analyses of co-variance.

Person-years at risk of developing cancer (Fig. 2) of the prostate, testes, colon, rectum, lung and breast were calculated as the number of years from entry into the study until the time of withdrawal (year of diagnosis of cancer, time of death or end of follow-up on 31 December 1991 for prostate/testicular/colon/rectum/lung cancer, 1992 for the subcohort of lung cancer, or 1994 for breast cancer, whichever event was the earliest).

Cox's proportional hazards analysis was used to investigate the associations of leisure, work or total physical activity with the risk of certain types of cancer in the cohort (papers I-IV). Incident cases of prostate, testicular, colon, rectal, lung or breast cancer were defined as outcome events. Confidence intervals (95%) were estimated. The fit of the models was examined by plotting the hazards; the results indicated that the application of the models was appropriate.

Analysis of co-variance was used when comparing differences and changes in serum lipids and BMI in leisure time sedentary and active attendees over the 7 years of follow-up (paper V).

All tests of significance were two sided and the significance level was chosen at 5%. These analyses were performed with the SAS Statistical package version 6.11.¹⁰³

4 MAIN RESULTS

Paper I: *Physical activity and the risk of prostate and testicular cancer: a cohort study of 53,000 Norwegian men*

A reduced, adjusted, relative risk (RR) of prostate cancer was observed among men who walked during occupational hours and performed either moderate recreational activity (RR = 0.61; 95%CI = 0.36–1.01) or regular training (RR = 0.45; 95%CI = 0.20–1.01), relative to sedentary men (p for trend = 0.03). In contrast, no association was observed between physical activity and the risk of testicular cancer. However, the number of cases of testicular cancer were small, limiting the statistical power of the study.

Paper II: *Physical activity and risk of colorectal cancer in men and women*

Among women, total physical activity (work and leisure combined) was inversely associated with colon cancer risk with a dose–response relationship (p for trend = 0.04). The reduction in colon cancer risk in women was particularly related to recreational physical activity. This was particularly evident with cancer of the proximal colon showing a greater reduction when recreationally active women were compared with sedentary women (RR = 0.51; 95%CI = 0.28–0.93). Among men aged 45 years or over at entry to the study, an inverse dose–response effect was observed between total physical activity and colon

cancer risk (p for trend = 0.04). This association was not observed for those men younger than 45 years at entry.

Furthermore, there was a reduced risk among recreationally active, non-obese (< 23.6 kg/m²) women compared with sedentary non-obese (< 23.6 kg/m²) women (RR = 0.45; 95%CI = 0.25–0.82). This study also suggests an effect modification by body mass index (BMI) among men because occupationally active men aged over 45 years at entry, in the lowest tertile of BMI (< 23.3 kg/m²), had a reduction in colon cancer risk (RR = 0.50; 95%CI = 0.26–0.97). No association was observed between physical activity and rectal cancer in men or women.

Paper III: *The influence of physical activity on lung cancer risk: a prospective study of 81,516 men and women*

Leisure but not work activity was inversely related to lung cancer risk in men after adjustments had been made (p for trend = 0.01). Men who exercised at least 4 hours a week had a lower risk than men who did not exercise (RR = 0.71; 95%CI = 0.52–0.97). Reduced risk of lung cancer was particularly marked for small cell carcinoma (RR = 0.59; 95%CI = 0.38–0.94), less for adenocarcinoma (RR = 0.65; 95%CI = 0.41–1.05), with no association seen for squamous cell carcinoma. When physical activity was assessed twice in a subcohort after 3–5 years, the risk of lung cancer was particularly reduced among men

who were most active at both assessments (RR = 0.39; 95%CI = 0.18–0.85). The small number of incident cases, combined with the narrow range of physical activity reported, may have limited our ability to detect an association between physical activity and lung cancer in women.

Paper IV: *Physical activity and risk of breast cancer*

Leisure time activity reduced the overall risk of breast cancer, after adjustments for age, body mass index (BMI), height, parity and geographical region; relative risk declined to 0.93 (0.71–1.22) and further to 0.63 (0.42–0.95) across increasing levels of leisure activity (p for trend = 0.04). Reduced risk was more pronounced in regularly exercising premenopausal relative to postmenopausal women and in younger (< 45 years at study entry) regularly exercising women (RR = 0.38; 95%CI = 0.19–0.79) relative to older (\geq 45 years) women when compared with the respective sedentary groups. Reduced risk was particularly demonstrated in lean (lowest tertile, BMI < 22.8 kg/m²) women who exercised at least 4 hours/week (RR = 0.28; 95%CI = 0.11–0.70). Repeated assessment of leisure time activity further confirmed this protective effect among non-obese, premenopausal women. Reduced risk was also observed in relation to work activity, especially for those who did heavy manual work compared with those whose work was sedentary (RR = 0.48; 95%CI = 0.25–

0.92); there was a more pronounced effect among premenopausal than postmenopausal women.

Paper V: *Physical activity improves the metabolic risk profiles in men and women: a 7 year follow-up study with repeated assessments of leisure time activity – the Tromsø Study*

Both sustained levels and change in level of leisure time physical activity were found to influence body mass index and serum lipids in a dose-response fashion, in both sexes, after adjustments for potential confounders. The differences of BMI and serum lipids between sustained sedentary and sustained exercising groups were consistently more pronounced after 7 years than at baseline, especially in the oldest age group. Men reporting sustained hard training, compared with sustained sedentary men, had the following: a lower concentration of Total-C (5.65 mmol/l vs 6.21 mmol/l) and triglycerides (1.34 mmol/l vs 1.85 mmol/l), a lower Total-C:HDL-C ratio by 19.0%, a lower BMI (23.9 kg/m² vs 25.7 kg/m²) and higher HDL-C concentration (1.52 mmol/l vs 1.36 mmol/l). Women reporting sustained regular or hard training, compared with sustained sedentary women, had the following: a lower concentration of Total-C (5.70 mmol/l vs 5.90 mmol/l) and triglycerides (1.03 mmol/l vs 1.18 mmol/l), a lower Total-C:HDL-C ratio by 7.5%, a lower BMI (23.1 kg/m² vs 23.6 kg/m²) and higher HDL-C concentration (1.73 mmol/l vs 1.66 mmol/l). An

increase in leisure time activity over 7 years improved the metabolic profiles, whereas a decrease worsened them in both sexes. Heart rate and physical fitness (women) were used to validate

physical activity; we observed a lower heart rate with increasing level of leisure time activity, and physical fitness increased with reported increase in activity level.

5 GENERAL DISCUSSION

This very large population-based cohort study included both leisure and occupational physical activity in men and women. This gives a comprehensive consideration of total physical activity for each individual. Although prospective cohort studies have many preferences compared with case-control studies, there are some limitations. Basic questions to consider are the accuracy of the physical activity questionnaire used, in addition to the degree of association between risk for site-specific cancer and physical activity; the extent to which the observed associations may result from bias, confounding or chance, and the extent to which they may be described as causal.

Most cancers are rare diseases and even with such a large cohort the number of some site-specific cancer cases in subgroups may be small, limiting the statistical power. As a result, the possibility of observing any associations between physical activity and testicular cancer in men and lung cancer in women could not be elucidated properly.

Assessment of physical activity

Self-reported questionnaires are an appropriate and practical method for the assessment of physical activity in large populations.¹⁵ The physical activity questionnaire used in the present thesis was originally developed for male subjects,³⁵ then adapted for both men and women. The observations that heart rate

decreased and physical fitness increased with increasing leisure time activity level (see paper V) support the fact that the questionnaire used is adequate for ranking the individuals to levels of leisure time physical activity when related to intensity. The ranking of levels of leisure time also seems to be valid in relation to energy expenditure (Table I), strengthening the comparability with other studies.²⁹

Table 1 Energy expenditure estimates related to levels of leisure time activity during 4 hours/week in a participant weighing 80 kg

Leisure time activity	Energy expenditure (kcal)
Sedentary	320
Moderate	1280
Regular exercise	2560
Hard exercise	3200

Sedentary: inactive = 1 MET value/4 hours per week (1 MET x 80 kg body weight) x (x 4 (4 @ 60 min)) = 320 kcal.

Moderate: walking/bicycling = 4 MET value/4 hours per week; (4 MET X 80 kg body weight) x (x 4 (4 @ 60 min)) = 1280 kcal.

Regular exercise: skiing = 8 MET value/4 hours per week; (8 MET x 80 kg body weight) x (x 4 (4 @ 60 min)) = 2560 kcal.

Hard exercise: running = 10 MET value/ 4 hours per week; (10 MET x 80 kg body weight) x (x 4 (4 @ 60 min)) = 3200 kcal.

The questionnaire used has been observed to discriminate sedentary men who were former athletes from their still active counterparts with respect to maximal oxygen uptake.³⁵ A correlation coefficient of 0.39 was

observed between leisure time physical activity and fitness in women.³⁷ These two observations support the finding that intensity of leisure time activity assessed is valid in relation to the reported leisure activity levels. Further, the association between changes in levels of physical activity and changes in BMI and serum lipids also supports the reproducibility of the present questionnaire (see paper V).

Comparing eight physical activity questionnaires, Albanes and colleagues³³ observed the questionnaires to be adequate for ranking individuals related to energy expenditure, as observed in the present study. Others have observed questionnaires to be more appropriate for assessing heavy intensity physical activity as well as treadmill performance, whereas questionnaires related to light or moderate activity were less valid.³¹ The present activity questionnaire also seems to cover higher intensity better than moderate activity (see paper V).

The large proportion of women reporting moderate regular leisure activity in contrast to men may indicate that also this questionnaire may have failed to include all activities relevant for women.¹⁰⁴

A person may under- or overestimate the level of performed physical activity. Recall bias related to underestimates of sedentary activities and overestimates of aerobic activities have been observed.¹⁰⁵ We cannot exclude such a 'wish' or recall bias, but such random misclassification is anticipated as being equal in cases and

non-cases as a result of the prospective design. Consequently, a non-differential misclassification may contribute to a reduced statistical significance by reducing the magnitude of the relationship in the present thesis; it cannot therefore explain the observed significant risk estimates. However, recall over a time period may provide less accurate recall. A three-month recall questionnaire can, however, give results similar to a detailed 7-day diary²³ and a questionnaire may yield a reasonable estimate of the past year.¹⁰⁶

The influence of error on the results from physical activity also depends on how closely this variable has been tracked over many years. One of the advantages of a prospective design, in contrast to case-control studies, is that long-term effects of a given exposure can be provided.¹⁰⁷ However, any exposure variable such as the activity level at baseline may change during the follow-up, as observed in the present thesis (papers III, IV and V). Repeated assessment allows consideration of the influence on risk estimates of these changes, adjustment for changes in the level of physical activity and observation of the influence of long-term effects of physical activity (papers III, IV and V). However, combining two assessments of physical activity may also increase the precision. By comparing the observed results with those of other studies and metabolic profiles, we can elucidate whether the combination of two assessments increased the precision and/or whether

continued physical activity has a different influence on risk estimates than short time activity, as observed by others.¹⁰⁸ Sustained higher activity level with repeated assessment showed more marked differences between sedentary and active men and women on metabolic risk profiles than one assessment (see paper V). Changes in leisure activity level influence the metabolic risk profiles in both sexes. This may indicate that consistently higher levels of leisure activity are important to reduce the risk for e.g. breast cancer and that the observed effect by combining two assessments not only can be explained by increase in the precision of our physical activity measurement.

Total physical activity can be analysed if both leisure and occupational activity are taken into account. However, variations of within-job classifications may have limited the distribution of occupational activity, for example, the large number of housewives (70% at baseline) in our cohort. Most of these housewives (76%) categorized themselves as group 2 (a lot of walking) during occupational hours. Consequently, we cannot exclude a misclassification among women related to occupational physical activity. In addition, a change in occupational physical activity may have been introduced in both sexes as a result of increases in mechanization and declines in manual work that occurred during follow-up.

In comparing physical fitness in a subgroup with the self-reported occupational physical activity level used, Løchen and Rasmussen³⁷

observed a decrease in physical fitness with increase in level of occupational activity for both sexes. This may be explained by the fact that occupational activity levels, as assessed in the present surveys, do not improve aerobic capacity also observed by others.¹⁰⁹ However, even short bouts of occupational activity need energy as indicated in paper IV, because occupationally active women have a higher level of daily energy intake which is important in carcinogenesis. ✓

Could the observed results be explained by bias?

The collection of data, analysis and interpretation of the results may deviate from the truth because of systematic variations. **Bias is not a property of the underlying population and is generally not a major problem in cohort studies.**¹¹⁰

In population-based cohort studies, with attendance rates above 80%, selection bias is generally regarded as unlikely. However, the attendance rate in the present study showed geographical variations, about 60% in Oslo to about 90% in Sogn og Fjordane. Holme et al.¹¹¹ observed that male non-attendees in Oslo varied by social class, with the highest attendance rate among the middle class and the lowest among the lowest social class. Accordingly, low leisure time physical activity is more likely among non-attendees because men in lower social classes are more frequently sedentary during leisure time. However, repeated assessment

of physical activity makes the reference category their own controls and reduces the effect of this type of selection bias (papers III, IV and V).

Furthermore, combining leisure and occupational sedentary physical activity reflects both high and low social classes.¹³ Consequently, a selection bias resulting from attendance rate or social class is unlikely to explain the observed associations, especially when both occupational and recreational physical activity are taken into account. Analyses with repeated assessments of physical activity may have introduced selection bias among those with persistent sedentary and persistent active leisure time at the two assessments. However, even a change in activity over 7 years of follow-up had an influence on the metabolic profiles in both sexes (see paper V). This supports an effect from sustained activity, in addition to improved precision of the physical activity variable.

One of the advantages of prospective studies is the possibility of obtaining data on both the exposure group and the outcome group at baseline, before cancer is diagnosed. However, any preclinical disease that reduces the possibility of performing physical activity will influence the reported activity level (*information bias*). Therefore, exclusion of preclinical diseases was carried out. In addition, those who emigrated, had a pre-existing malignancy or were diagnosed with a malignant disease within the first year of attending the cohort were excluded from the

analysis for all cancer types studied. Hence, both in validating the physical activity assessment used (see paper V) and in the studies related to risk of cancer (see papers I-IV), the possibility of any undiagnosed cancer influencing the reported level of physical activity is less likely. However, any other preclinical or chronic disease, such as ischaemic heart disease, was not excluded in the final analysis related to risk of cancer. When excluding people (< 1%) with reported diabetes or ischaemic heart disease at baseline, no changes in risk estimates were observed (papers I-IV).

In Norway there is compulsory reporting by hospital departments, pathology laboratories and death certificates; there is also an almost 100% histological verification of most cancer cases. This indicates no influence on risk estimates from inadequate reporting of cancer cases (*diagnostic bias*), which has often been a problem in prospective studies.¹⁰ In addition, the unique 11-digit identification number in Norway validates the diagnosis among cohort members who have died in the cohort, who have emigrated or who are currently alive.

Physically active people are more likely to have contact with physicians than those who are physically inactive.¹¹² This difference would create an increased likelihood of early diagnosis of cancer in physically active men and women. If such a diagnostic bias occurred in this study, the observed estimates are likely to be an underestimate of the true strength of

the association between physical activity and risk of cancer.

Appropriate adjustments for confounders?

It is important to be cautious about whether unadjusted or residual effects of certain variables affect the risk of cancer. A confounding variable must be an independent risk factor for the disease as well as being associated with the exposure under study. In addition, if a variable is included in an intermediate step between an exposure and disease, that variable is not a true confounder.¹¹³

No dietary data were available for the analysis related to prostate, testicular, colon, rectal and lung cancer. High intake of dietary fat has been observed to increase the risk of prostate cancer¹¹⁴⁻¹¹⁵ and increased intake of vegetables and fruit is associated with reduced risk of lung cancer in men and women.¹¹⁶⁻¹¹⁷ In addition, higher intake of total fat has been observed in sedentary men relative to physically active men.¹¹⁸ We observed a higher intake of fruit and vegetables in participants who were active in their leisure time compared with those who were sedentary (see paper V). Consequently, dietary factors may be a potential confounder in relation to physical activity and risk of prostate and lung cancer.

A recent Norwegian prospective study, performed in part in the same study population, did not observe any association between meat, fish, fat,

energy, fibre or calcium intake and risk of colon cancer.¹¹⁹ Other studies have concluded that physical activity and dietary factors are independent risk factors for colon cancer.^{118,120} Consequently, it is less likely that dietary intake is an important confounder in relation to the association between physical activity and risk of colon cancer.

Could any unadjusted potential confounding factor, with a strong relationship to both physical activity and risk of lung cancer, explain the protective effect of the results observed? Men in lower social classes have been observed to be less active at leisure but more active at work than men in the higher social classes.¹³ However, in a recent review of differences in cancer incidence among socioeconomic groups the authors concluded that it is still unclear whether the reported associations in studies can be attributed to lifestyle related riskfactors for cancer such as smoking and nutritional habits¹²¹ also observed in some studies.^{121,122} As the population under study also consists of a relatively homogeneous social group, any residual confounding from other factors related to socioeconomic status is unlikely to be substantial. The observation that increased total physical activity reduced the overall lung cancer risk for those who were sedentary at leisure ('low social class') and work ('high social class') was used as a reference to reduce the possibility that social class is a major confounder.

There is a strong causal relationship between smoking and risk of lung cancer.¹²³ In addition, people

active in their leisure time were less often current smokers, and more often never and ex-smokers than those who were sedentary (see paper III). However, small differences were observed related to smoking habits in different categories of occupational activity. Moreover, careful adjustments for smoking habits were performed in analyses related to lung cancer. This reduces the possibility that smoking habits could influence the association observed when related to total physical activity and risk of lung cancer. Smoking-related lung cancer risk by cell type is strongest for squamous cell carcinoma and small cell carcinoma.¹²⁴⁻¹²⁵ The lack of a protective effect of physical activity on squamous cell carcinoma reduces a residual effect of smoking, and supports the reduced risk of lung cancer resulting from physical activity. The age-adjusted risk estimates related to physical activity and lung cancer risk changed, but not much and significant inverse associations were still observed after performing multivariate adjustments for current, past, amount and duration of smoking.

Smoking habits were also adjusted for in multivariate analysis in relation to other types of cancer: prostate, testes, colon, rectum and breast. The relative risk estimates adjusted for smoking were, in general, similar to age-adjusted risk estimates and excluded smoking as a substantial confounder.

The two cities and three counties studied have some different incidence rates of the cancer types studied,⁸⁹ and the participants were also enrolled

into the study in different years. Stratified analysis by geographical areas was chosen to take care of this problem, and the relative risks, adjusted for geographical region and age at entry, were in general similar to the age-adjusted relative risk estimates. This reduced the possibility of place of residence being a substantial confounder for the association between physical activity and risk of cancer types studied.

Numerous studies have observed that body weight and weight gain during adulthood are influenced by performed physical activity (see paper V). Thus, body mass index represents an intermediate factor between diet and cancer, and not a true confounder. Comparable consideration could be given to age at menarche, as this may be influenced by physical activity – physical activity reduces weight, promotes later ovarian maturation and consequently promotes a later onset of menstruation.^{64, 126}

In relation to prostate and testicular cancer, we were unable to control for some other potential risk factors which may act as confounders such as sexually transmitted disease and sexual habits, early life events (e.g. prematurity)¹²⁷⁻¹²⁸ and cryptorchidism in relation to testicular cancer.¹²⁹⁻¹³⁰

Is there any causality between physical activity and cancer?

Careful evaluations should be drawn before any conclusions are made about causality. Hill¹³¹ suggested nine criteria for causality: strength, consistency, specificity, temporality,

biological gradient, plausibility, coherence, experimental evidence and analogy. These criteria have been critically evaluated and found to be of importance for cause-and-effect associations.¹¹³

Strength of the association

Strong associations between physical activity and risk of site-specific cancer studied indicate a greater likelihood of a causal relationship. Although the observed statistical associations between physical activity and prostate and colon cancer (in men and women) may be rather weak, it should not exclude a causal relationship. However, an undetected bias could explain the modest observed associations for prostate and colon cancer (papers I and II). In papers III (lung cancer) and IV (breast cancer), the observed associations are stronger, especially when related to repeated assessment of leisure time activity.

Biological gradient: dose-response relationship

A genuine association is expected to increase with increasing level and duration of exposure. A dose-response is an important indication of causality, while the lack of such a relationship argues against causality.¹⁰⁷ A dose-response relationship between total physical activity and risk of prostate cancer was observed among those who walked a lot during occupational hours. A dose-response relationship was also observed between total physical activity and risk of colon

cancer among women and men aged over 45 years at entry. Moreover, this pattern was observed between leisure activity and lung cancer for both one single and repeated assessments of leisure activity. An association was especially marked between occupational and leisure time physical activity and risk of breast cancer. The strongest relationship was observed between leisure time activity and cancer risk in leaner premenopausal women through repeated assessment of activity and among younger postmenopausal women. We also observed a dose-response relationship between changes in levels and sustained levels of leisure time activity and serum lipids and BMI in both sexes.

Repeated assessment of physical activity may increase precision and this could explain the observed stronger dose-response relationship between leisure activity and risk of lung and breast cancer because loss of accuracy of physical activity would diminish any dose-response relationship.

Temporality

The prospective design satisfies the criteria if physical activity precedes the development of cancer. Different times after assessment of physical activity were eliminated to assess the effect of latent disease, which may have affected physical activity level in diseased individuals. Relative risk estimates and tests for linear trend were essentially unchanged for men and women, after excluding an additional 2 or 4 years of follow-up

for those cancer types studied.

Between single shot and continuous lifetime exposures, one can clearly consider a wide range of types of intermittent or limited exposure. We considered a continuous exposure of physical activity and concentrated on what appear to be two features of particular importance: the age at which exposure starts and the time period until the end of follow-up. Therefore, an age split is carried out to see if these differences interact on risk estimates. We suggest a protective effect of physical activity on prostate cancer risk only among those over 60 years, but our results are borderline and confidence intervals overlap with those younger than 60 years.

Migrant studies indicated that environmental factors operating in young adulthood may be determinants for breast cancer, whereas factors operating throughout life seem to be important for colon cancer.⁹⁴ The observed association for colon cancer in older, relative to younger, men supports colon cancer being more sensitive to lifetime physical activity. More convincing are the observations of a stronger association between physical activity and risk of breast cancer among premenopausal and younger postmenopausal women, which may indicate that breast cells are more sensitive during the early period of life when both the number of stem cells and cell differentiation are greater.⁵⁷

Specificity and consistency

Specificity of risk to subgroups indicates a causal relationship and this was observed between leisure time activity and lung cancer, especially small cell carcinoma. Moreover, the increased risk of breast cancer in younger women, relative to older ones, supports specificity of risk to age groups.

A single study of the effect of physical activity on cancer risk seldom provides convincing evidence, but it might trigger interest and create new hypotheses. However, several studies have observed a protective effect of occupational and/or leisure time physical activity^{22,132-134} and physical fitness¹³⁵ on the risk of prostate cancer. The variance observed in some studies^{42,136} may be explained by methodological bias because sedentary jobs are a poor indicator¹³⁵ and college athletics is too remote in time to be important in carcinogenesis of the prostate.⁴²

The observation¹²⁹⁻¹³⁰ that lack of exercise and a sedentary lifestyle increase the risk of testicular cancer in contrast to our study. These results need to be confirmed in other studies.

The evidence for a protective influence of physical activity on colon cancer risk has consistently been observed in numerous studies in different countries world wide, related to both occupational^{41,137-138} and leisure time activity.^{17,41,47, 108,137-141}

The lack of association observed between physical activity and rectal cancer is supported from many others.^{108, 137}

The associations between physical activity and lung cancer risk has not been studied much, but are supported with a few previous observations.^{22,142} However, no other studies have analysed the association between physical activity and lung cancer risk by histological subtypes.

Since Frisch and colleagues¹ observed lower prevalence of breast cancer among former college athletes, a reduction in breast cancer risk has been observed among women resulting from leisure time^{18,22, 26,143} as well as occupational physical activity.¹⁴⁴⁻¹⁴⁵ A few others have observed inconsistent results^{27,42} and these observations may be explained by the fact that college athletics is too remote an event,⁴² whereas in the Framingham study most of the women were older.²⁷

Coherence and biological knowledge

The association between physical activity and the cancer types studied is more plausible if the effect of physical activity supports currently known pathophysiological mechanisms.

Overall physical activity has numerous health benefits, such as improvements in the concentration of serum lipids,⁸⁵ weight control (paper V)²⁸ and reduction in insulin levels.¹⁴⁶ High circulating levels of insulin may promote the growth of tumours,¹⁴⁷ and recently hypothesized its importance in carcinogenesis of the colon⁴⁷ and breast.⁸⁸

Another, important physiological effect of physical activity is related to energy balance and utilization of calories. Tannenbaum¹⁴⁸ was the first

to observe that caloric restriction inhibits the development of spontaneous tumours (mammary and lung) in animals. Kritchevsky⁵² observed that exercise halved tumour incidence in rats fed *ad libitum* and was equivalent to 25% caloric restriction. Caloric restriction reduces cellular proliferation and mitotic activity.¹⁴⁹ Energy restriction can also lead to enhanced DNA repair.¹⁵⁰

The study of carcinogenesis through animal experiments confirms this as a multi-step process (initiation, promotion, progression) driven by carcinogen-induced genetic and epigenetic damage in suspicious cells. Thus, it is of importance to consider at what stage in the carcinogenic process physical activity may act. As recent studies have observed, adult weight gain is a risk factor for breast cancer,^{117,151} so physical activity may therefore act as a late stage factor. However, influence on stem cells and mitotic activity through caloric restriction indicates that physical activity may influence initiation. Physical activity may therefore modify the carcinogenesis at different stages. In addition, both *amount* and *duration* of physical activity have been observed to be important in relation to their influence on carcinogenesis.

Carcinogenesis of the prostate seems to involve testosterone.⁶⁶⁻⁶⁷ Hence, a role for physical activity in its development is plausible because physical activity may reduce testosterone levels.⁶⁸⁻⁶⁹ The normal functioning of the large bowel includes a certain level of physical activity, and exercise has shown to

reduce stool transit time through the large bowel. Consequently, exposure time of the colon mucosa and potential carcinogens in the faecal stream may be decreased by exercise.

Physical activity is also known to improve pulmonary capacity, and increased pulmonary ventilation and perfusion may reduce the interaction time and concentration of any carcinogenic factors in the airways. Moreover, pulmonary function has been observed as a predictor for risk of lung cancer.¹⁵²

The specific relationship between physical activity and breast cancer may be related either to the general influence on energy balance and/or more specifically to a reduction in the cumulative exposure of both cyclic oestrogen and progesterone in physically active women relative to sedentary women.

Finally, a physically active lifestyle counteracts adverse health habits and improves the quality of life; psychological attitudes threaten the quality of living and these have recently been suggested as important for risk of cancer.¹⁵²⁻¹⁵³

A randomized controlled trial is the best way of showing that changes in physical activity change the risk of prostate, testicular, colon, rectal, lung and breast cancer. However, experimental studies in humans meet ethical and critical problems. Consequently, some hypothesis have first to be investigated in experimental animal studies. Animal studies have observed inhibition of mammary carcinogenesis by voluntary treadmill exercises¹⁵⁴ and reduction in incidence

and multiplicity of colon adenocarcinomas has been observed in voluntary exercised rats.¹⁵⁵ This gives experimental evidence for a relationship between physical activity and carcinogenesis.

6 IMPLICATIONS AND FURTHER RESEARCH

Cultural changes have outpaced any genetic adaptation. Thus, from a genetic point of view we are still hunters and future research should focus on the suggestion that physical inactivity may act as a possible carcinogenic hazard. Even if physical activity has only a small protective effect, the prevalence of inactivity in industrialized societies is great. The attributable risk associated with inactivity may be quite high and be of importance for cancer prevention. Although numerous studies have been performed during the past five years many important questions remain and point to further studies.

- Studies in which physical activity is analysed as a dynamic process from childhood throughout the whole life are needed. Hence, repeated assessments of physical activity in cohort studies should be done. In this way, the importance of age-specific physical activity and life-time physical activity between populations and time periods could be further elaborated.

- There should be improved comparability of the physical activity assessments between different study populations and at different ages. Moreover, performed physical activity

reflecting type, intensity, duration and amount of performed physical activity is related to both age and gender. Consequently, these differences should be further focused in order to improve assessments of physical activity in epidemiological studies.

- Given the complexity of physical activity, epidemiological research should, in collaboration with clinical and laboratory medicine, focus on energy balance and biomarkers associated with different levels of physical activity and changes in physical activity, using experience from these areas of medicine.

7 CONCLUSIONS

The present work has demonstrated that sustained high levels and change from sedentary to higher levels of leisure time physical activity in adult men and women improved metabolic risk profiles; reduced weight gain and serum concentration of total cholesterol, triglycerides and increased serum concentration of high-density lipoprotein cholesterol. This confirms that the physical activity assessments used have importance for metabolic profiles at a magnitude that can influence the risk of chronic diseases.

- The inverse association between both recreational and occupational physical activity and the risk of prostate cancer may act through a combination of improved energy balance and modified hormonal influence. The evidence is still sparse but more knowledge may point to a

protective role of physical activity.

- The small number of cases of testicular cancer limits the statistical power of the study and any causality remain unsolved.

- In spite of a modest association observed the overwhelming results from comparable studies support that physical activity is protective against colon but not rectal cancer in both genders.

- Our results indicate a negative dose-response association between recreational physical activity and lung cancer in men. This association was especially observed between leisure physical activity and small cell carcinoma of the lung. It is hypothesized that improved pulmonary function is sufficient to counteract carcinogenesis in groups with exposure to carcinogenic agents. However, more studies are needed before we can conclude if this is a cause-effect association.

- Our study suggests an inverse association between both leisure and work activity and the overall risk of breast cancer. Further, the leisure time effect indicates a more marked effect among premenopausal and younger postmenopausal, lean women. This is in agreement with most other studies supporting a protective role for physical activity in the prevention of breast cancer. Possible mechanisms may be hormone- and energy-dependent effects.

8 REFERENCES

1. Frisch RE, Wyshak G, Albright NL, et al. Lower prevalence of breast cancer and cancers of the reproductive system among former college athletes compared to non-athletes. *Br J Cancer* 1985;52:885-91.
2. Eaton CB, Konner M, Shostak M. Stone agers in the fast line: chronic degenerative disease in evolutionary perspective. *Am J Med* 1984;84:739-49.
3. Park RJ. How active were early population ? In; *Physical activity in early and modern populations*. ed. R Malina, H Eckert, Champaign: Human Kinetics 114pp 1988;1:13-21.
4. Stanley SM: Chronospecies' longevities, the origin of genera, and the punctuational model of evolution. *Paleobiology* 1978;4:26-40.
5. Cooper KH: *The aerobics way*. New York: Bantam Books, 1977; 257-66.
6. Ramazzini B. *Diseases of workers* (1700) (Latin). Translated by Wright. New York: Hafner, 1964.
7. Åstrand P-O. From exercise physiology to preventive medicine. *Ann Clin Res* 1988;20:10-17.
8. Caspersen CJ, Powell KE, Christenson GM. Physical activity, exercise, and physical fitness: definitions and distinctions for health-related research. *Public Health Rep* 1985;100:126-31.
9. American College of Sports Medicine: The recommended quantity and quality of exercise for developing and maintaining fitness in healthy adults. *Med Sci Sports Exerc* 1990;22:265-74.
10. NIH Consensus Conference. Physical activity and health. NIH consensus development panel on physical activity and cardiovascular Health. *JAMA* 1996;276:241-6.
11. Bouchard C, Lesage R, Lortie G, et al. Aerobic performance in brothers, dizygotic and monozygotic twins. *Med Sci Sports Exerc* 1986;18:639-46.
12. Pérusse L, Tremblay A, Leblanc C, Bouchard C. Genetic and environmental influences on level of habitual physical activity and exercise participation. *Am J Epidemiol* 1989; 129:1012-22.
13. Holme I, Helgeland A, Hjermann I, Leren P, Lund-Larsen PG. Physical activity at work and at leisure in relation to coronary risk factors and social class. A 4-year mortality follow-up. The Oslo study. *Acta Med Scand* 1981;209:277-83.
14. Laporte RE, Montoye HJ, Caspersen CJ. Assessment of physical activity in epidemiologic research: problems and prospects. *Public Health Rep* 1985;100:131-46.

15. Paffenbarger Jr. RS, Blair SN, Lee I-M; Hyde RT. Measurement of physical activity to assess health effects in free-living populations. *Med Sci Sports Exerc* 1993;**25**:60-70.
16. Washburn RA, Montoye HJ. The assessment of physical activity by questionnaire. *Am J Epidemiol* 1986;**123**:563-76.
17. Ballard-Barbash R, Schatzkin A, Albanes D, et al. Physical activity and risk of large bowel cancer in the Framingham Study. *Cancer Res* 1990;**50**:3610-13.
18. Bernstein L, Henderson BE, Hanisch R, Sullivan-Halley J, Ross RK. Physical exercise and reduced risk of breast cancer in young women. *J Natl Cancer Inst* 1994;**86**:1403-8.
19. Paffenbarger Jr. RS, Hyde RT, Wing AL, Lee I-M, Jung DL, Kampert JB. The association of changes in physical activity level and other lifestyle characteristics with mortality among men. *N Engl J Med* 1993;**328**:538-45.
20. Lissner L, Bengtsson C, Björklund C, Wedel H. Physical activity levels and changes in relation to longevity. *Am J Epidemiol* 1996;**143**:54-62.
21. Lakka TA, Salonen JT. Physical activity and serum lipids: a cross-sectional population study in eastern Finnish men. *Am J Epidemiol* 1992;**136**:806-18.
22. Albanes D, Blair A, Taylor PR. Physical activity and risk of cancer in the NHANES I population. *Am J Public Health* 1989;**79**:744-50.
23. Arroll B, Jackson R, Beaglehole R. Validation of a three-month physical activity recall questionnaire with a seven-day food intake and physical activity diary. *Epidemiology* 1991;**2**:296-9.
24. Slattery ML, Jacobs Jr. DR. Physical fitness and cardiovascular disease mortality. The US railroad study. *Am J Epidemiol* 1988;**127**:571-80.
25. Leren P, Askevold EM, Foss OP, et al. The Oslo Study. Cardiovascular disease in middle-aged and young men. *Acta Med Scand suppl* 1975;**588**:1-38.
26. D'Avanzo, Nanni O, La Vecchia C et al. Physical activity and breast cancer risk. *Cancer Epidemiol, Biomarkers Prevention* 1996;**5**:155-60.
27. Dorgan JF, Brown C, Barrett M, et al. Physical activity and risk of breast cancer in the Framingham Heart Study. *Am J Epidemiol* 1994;**139**:662-9.
28. Williamson DF, Madans J, Anda RF, Kleinman JC, Kahn HS, Byers T. Recreational physical activity and ten-year weight change in a US national cohort. *Int J Obes* 1993;**17**:279-86.

29. Ainsworth BE, Haskell WL, Leon AS, et al. Compendium of physical activities: classification of energy costs of human physical activities. *Med Sci Sports Exerc* 1993;**25**:71-80.
30. Severson RK, Nomura AM, Grove JS, Stemmermann GN. A prospective analysis of physical activity and cancer. *Am J Epidemiol* 1989;**130**:522-9.
31. Jacobs DR, Ainsworth BE, Hartman TJ, Leon AS. A simultaneous evaluation of 10 commonly used physical activity questionnaires. *Med Sci Sport Exrc* 1993;**25**:81-91.
32. Sequeira MM, Rickenbach M, Wietlisbach V, Tullen B, Schutz Y. Physical activity assessment using a pedometer and its comparison with a questionnaire in a large population survey. *Am J Epidemiol* 1995;**142**:989-99.
33. Albanes D, Conway JM, Taylor PR, Moe PW, Judd J. Validation and comparison of eight physical activity questionnaires. *Epidemiology* 1990;**1**:65-71.
34. Marrugat J, Elosua R, Covas M-I, Molina L, Rubiés-Prat J and the MARATHOM investigators. Amount and intensity of physical activity, physical fitness, and serum lipids in men. *Am J Epidemiol* 1996;**143**:562-9.
35. Saltin B, Grimby G. Physiological analysis of middle-aged and old former athletes. Comparison with still active athletes of the same ages. *Circulation* 1968;**38**:104-15.
36. Siconolfi SF, Lasater TM, Snow RCK, Carleton RA. Self-reported physical activity compared with maximal oxygen uptake. *Am J Epidemiol* 1985;**122**:101-5.
37. Løchen M-L, Rasmussen K. The Tromsø study: physical fitness, self reported physical activity, and their relationship to other coronary risk factors. *J Epidemiol Community Health* 1992;**26**:103-7.
38. Sandvik L, Erikssen J, Thaulow E, Eriksen G, Mundal R, Rodahl. Physical fitness as a predictor of mortality among healthy, middle-aged Norwegian men. *N Engl J Med* 1993;**328**:533-7.
39. Blair SN, Kohl HW, 3d, Paffenbarger Jr. RS, Clark DG, Cooper KH, Gibbons LW. Physical fitness and all-cause mortality. A prospective study of healthy men and women. *JAMA* 1989;**262**:2395-2401.
40. Oja P, Laukkanen R, Pasanen M, Vuori I. A new fitness test for cardiovascular epidemiology and exercise promotion. *Ann Med* 1989;**21**:249-50.
41. Garabrant DH, Peters JM, Mack TM, Bernstein L. Job activity and colon cancer risk. *Am J Epidemiol* 1984;**119**:1005-14.

42. Paffenbarger Jr. RS, Hyde RT, Wing AL. Physical activity and incidence of cancer in diverse populations: a preliminary report. *Am J Clin Nutr* 1987;**45**:312-7.
43. Brownson RC, Zahm SH, Chang JC, Blair A. Occupational risk of colon cancer. An analysis by anatomic subsite. *Am J Epidemiol* 1989;**130**:675-87.
44. Danforth E. Diet and obesity. *Am J Clin Nutr* 1985;**41**:1132-45.
45. Albanes D, Taylor PR. International differences in body height and weight and their relationship to cancer incidence. *Nutr Cancer* 1990;**14**:69-77.
46. Giovannucci E, Ascherio A, Rimm EB, Colditz GA, Stampfer MJ, Willett WC. Physical activity, obesity, and risk for colon cancer and adenoma in men. *Ann Intern Med* 1995;**122**:327-34.
47. Tretli S. Height and weight in relation to breast cancer morbidity and mortality. A prospective study of 570,000 women in Norway. *Int J Cancer* 1989;**44**:23-30.
48. Barnes-Josiah D, Potter JD, Sellers A, Himes JH. Early body size and subsequent weight gain as predictors of breast cancer incidence (Iowa, United States). *Cancer Causes Control* 1995;**6**:112-8.
49. Ziegler RG, Hoover RN, Nomura AM, et al. Relative weight, weight change, height, and breast cancer risk in Asian-American women. *J Natl Cancer Inst* 1996;**88**:650-60.
50. Cohen LA, Choi K, Wang C-X. Influence of dietary fat, caloric restriction, and voluntary exercise on *N*-Nitrosomethylurea-induced mammary tumorigenesis in rats. *Cancer Res* 1988;**48**:4276-83.
51. Boissonneault GA, Elson CE, Pariza MW. Net energy effects of dietary fat on chemically induced mammary carcinogenesis in F344 rats. *J Natl Cancer Inst* 1986;**76**:335-8.
52. Kritchevsky D. Caloric restriction and experimental carcinogenesis. Exercise, calories, fat and cancer. *Adv Exp Med Biol* 1992;**322**:131-7.
53. Welsch CW, DeHoog JV, O'Connor DH, Scheffield LG. Influence of dietary fat levels on development and hormone responsiveness of the mouse mammary gland. *Cancer Res* 1985;**45**:6147-54.
54. Koffler KH, Menkes A, Redmond RA, Whitehead WE, Pratley RE, Hurley BF. Strength training accelerates gastrointestinal transit in middle-aged and older men. *Med Sci Sports Exerc* 1992;**24**:415-19.
55. Lampe JW, Fredstrom SB, Slavin JL, Potter JD. Sex differences in colonic function: a randomised trial. *Gut* 1993; **34**:531-6.

56. Cordain L, Latin RW, Behnke JJ. The effects of an aerobic running program on bowel transit time. *J Sports Med Phys Fitness* 1986;26:101-4.
57. Pike MC, Spicer DV, Dahmouh L, Press MF. Estrogens, progestogens, normal breast cell proliferation and breast cancer risk. *Epidemiol Rev* 1993;15:17-35.
58. Bernstein L, Ross RK, Lobo RA, Hanisch R, Krailo MD, Henderson BE. The effects of moderate physical activity on menstrual cycle patterns in adolescence: implications for breast cancer prevention. *Br J Cancer* 1987;55:681-5.
59. Prior JC, Vigna YM, Schechter MT, Burgess AE. Spinal bone loss and ovulatory disturbances. *N Engl J Med* 1990;323:1221-7.
60. Aakvaag A, Sand T, Opstad PK, Fonnum F. Hormonal changes in serum in young men during prolonged physical strain. *Eur J Appl Physiol* 1978;39:283-91.
61. Wheeler GD, Wall SR, Belcastro AN, Cumming DC. Reduced serum testosterone and prolactin levels in male distance runners. *JAMA* 1984;252:514-16.
62. Ellison PT, Lager C. Moderate recreational running is associated with lowered salivary progesterone profiles in women. *Am J Obstet Gynecol* 1986;154:1000-3.
63. Broocks A, Pirke K, Schweiger U et al. Cyclic ovarian function in recreational athletes. *J Appl Physiol* 1990;68:2003-86.
64. Frisch RE, Gotz-Welbergen AV, McArthur JW, et al. Delayed menarche and amenorrhea of college athletes in relation to age at onset of training. *JAMA* 1981;246:1559-63.
65. Zumoff B, Levin J, Strain GW, et al. Abnormal levels of plasma hormones in men with prostate cancer: evidence toward a "two-disease" theory. *Prostate* 1982;3:579-88.
66. Ross R, Bernstein L, Judd H, Hanisch R, Pike M, Henderson B. Serum testosterone levels in healthy young black and white men. *J Natl Cancer Inst* 1986;76:45-8.
67. Gann PH, Hennekens CH, Ma J, Longcope C, Stampfer MJ. Prospective study of sex hormone levels and risk of prostate cancer. *J Natl Cancer Inst* 1996;88:1118-26.
68. Morville R, Pesquies PC, Guezennec CY, Serrurier BD, Guignard M. Plasma variations in testicular and adrenal androgens during prolonged physical exercise in man. *Ann Endocrinol Paris* 1979;40:501-10.
69. Hackney AC, Sinning WE, Bruot BC. Reproductive hormonal profiles of endurance-trained and untrained males. *Med Sci Sports Exec* 1988;20:60-5.

70. Remes K, Kuoppasalmi K, Adlercreutz H. Effect of physical exercise and sleep deprivation on plasma androgen levels: modifying effect of physical fitness. *Int J Sports Med* 1985;6:131-5.
71. Kuller LH, Ockene J, Meilahn E, Svendsen KH. Relation of forced expiratory volume in one second (FEV1) to lung cancer mortality in the Multiple Risk Factor Intervention Trial (MRFIT). *Am J Epidemiol* 1990;132:265-74.
72. Higgins M, Keller JB, Wagenknecht LE, et al. Pulmonary function and cardiovascular risk factor relationships in black and in white young men and women. The CARDIA Study. *Chest* 1991;99:315-22.
73. Preston-Martin S, Pike MC, Ross RK, Jones PA, Henderson BE. Increased cell division as a cause of human cancer. *Cancer Res* 1990;50:7415-21.
74. Brown LM, Pottern LM, Hoover RN. Testicular cancer in young men: the search for causes of the epidemic increase in the United States. *J Epidemiol Community Health* 1987;41:349-54.
75. Preston-Martin S, Mack W, Henderson B. Risk factors for gliomas and meningiomas in males in Los Angeles County. *Cancer Res* 1989;49:6137-43.
76. Larrabee RC. Leukocytosis after violent exercise. *J Med Res* 1902;7:76-82.
77. Yu DTY, Clements PJ, Pearson CM. Effect of corticosteroids on exercise-induced lymphocytosis. *Clin Exp Immunol* 1977;28:326-31.
78. Nieman DC, Nehlsen-Canarella SL, Donohue KM, et al. The effects of acute moderate exercise on leucocyte and lymphocyte subpopulations. *Med Sci Sports Exerc* 1991;23:578-85.
79. Nieman DC, Buckley KS, Henson DA, et al. Immune function in marathon runners versus sedentary controls. *Med Sci Sports Exerc* 1995;27:986-92.
80. Shephard RJ, Shek PN. Exercise, aging and immune function. *Int J Sports Med* 1995;16:1-6.
81. Nehlsen-Canarella SL, Nieman DC, Balk-Lamberton AJ, et al. The effects of moderate exercise training on immune response. *Med Sci Sports Exerc* 1991;23:64-70.
82. Klarlund Pedersen B, Ullum H. NK cell response to physical activity: possible mechanisms of action. *Med Sci Sports Exerc* 1994;26:140-6.
83. Kiens B, Lithell. Lipoprotein metabolism influenced by training-induced changes in human skeletal muscle. *J Clin Invest* 1989; 83:558-64.

84. Hellenius M-L, de Faire U, Berglund B, Hamsten A, Krakau I. Diet and exercise are equally effective in reducing risk for cardiovascular disease. Results of a randomized controlled study in men with slightly to moderately raised cardiovascular risk factors. *Atherosclerosis* 1993;**103**:81-91.
85. Williams PT. High-density lipoprotein cholesterol and other risk factors for coronary heart disease in female runners. *N Engl J Med* 1996;**334**:1298-1303.
86. Haffner SM, Katz MS, Dunn JF. Increased upper body and overall adiposity is associated with decreased sex hormone binding globulin in postmenopausal women. *Int J Obes* 1991;**15**:471-78.
87. Krotkiewski M, Björntorp P. Muscle tissue in obesity with different distribution of adipose tissue, effects of physical training. *Int J Obes* 1986;**10**:331-41.
88. Bruning PF, Bonfrer JMG, Hart AAM, Van Noord PAH, De Jong-Bakker M, Nooijen WJ. Insulin resistance and breast-cancer risk. *Int J Cancer* 1992;**52**:511-16.
89. The Cancer Registry of Norway. *Cancer in Norway, 1993*. Oslo: The Norwegian Cancer Registry, Norway, 1996.
90. Harvei S, Tretli S, Langmark F. Cancer of the prostate in Norway 1957-1991- A descriptive study. *Eur J Cancer* 1996;**32A**:111-7.
91. Parkin DM, Pisani P, Ferlay J. Estimates of the worldwide incidence of eighteen major cancers in 1985. *Int J Cancer* 1993;**54**:594-606.
92. Adami HO, Bergstrom R, Mohner M, et al. Testicular cancer in nine northern European countries. *Int J Cancer* 1994;**59**:33-8.
93. Haenszel W. Cancer mortality among foreign born in the United States. *J Natl Cancer Inst* 1961;**26**:32-137.
94. Buell P, Dunn JE. Cancer mortality among Japanese Issei and Nisei of California. *Cancer* 1965;**18**:656-64.
95. Travis WD, Travis LB, Devesa SS. Lung cancer. *Cancer* 1995;**75**:191-202.
96. Bjartveit K, Foss OP, Gjervig T, Lund-Larsen PG. The cardiovascular disease study in Norwegian counties. Background and organization. *Acta Med Scand suppl* 1979;**634**:1-70.
97. Thelle DS, Førde OH, Try K, Lehman EH. The Tromsø Heart Study. Methods and main results of the cross-sectional study. *Acta Med Scand* 1976;**200**:107-18.
98. National Health Screening Service. *The cardiovascular disease study in Norwegian counties: Results from second screening*:Oslo, 1988.

99. Bønaa KH. *Relationship between hemodynamics and blood lipids in population surveys, and effects of n-3 fatty acids*. Thesis. Tromsø: Institute of Community Medicine, University of Tromsø, 1992.
100. Blaker B, Rimestad AH. Food composition table by the Norwegian National Nutrition Council, 6th edition:Oslo,1991 (In Norwegian).
101. Gaard M, Sandvin O, Løken EB. *Calculation of nutrient intake based on dietary data from 2nd cardiovascular disease survey in Finnmark, Sogn og Fjordane, and Oppland 1977-1983*. Technical Report 1. The Cancer Registry of Norway:Oslo, 1996.
102. Lund E. Pilot Study for the evaluation of completeness of reporting to the Cancer Registry. In *Incidence of Cancer in Norway 1978; 1981*;11-15.
103. SAS Institute I. *SAS/STAT Guide for Personal Computers*, Version 6 Edition. SAS Institute, Inc. 1992.
104. Ainsworth BE, Richardson M, Jacobs Jr. DR, Leon AS. Gender differences in physical activity. *WSPAJ* 1993;2:1-16.
105. Klesges RC, Eck LH, Mellon MW, Fulliton W, Somes GW, Hanson CL. The accuracy of selfreport of physical activity. *Med Sci Sports Exerc* 1990;22:690-7.
106. Aaron DJ, Kriska AM, Dearwater SR, Cauley JA, Metz KF, LaPorte RE. Reproducibility and validity of an epidemiologic questionnaire to assess past year physical activity in adolescents. *Am J Epidemiol* 1995;142:191-201.
107. Breslow NE, Day NE. *Statistical Methods in cancer Research*, Vol I. The analysis of case-control studies. Lyon International Agency for Research on Cancer, 1980.
108. Lee I-M, Paffenbarger Jr. RS, Hsieh C-C. Physical activity and risk of developing colorectal cancer among college alumni. *J Natl Cancer Inst* 1991; 83:1423-9.
109. Farrell PA, Maksud MG, Pollock, et al. A comparison of plasma cholesterol, triglycerides, and high density lipoprotein-cholesterol in speed skaters, weightlifters and non-athletes. *Eur J Appl Physiol* 1982;48: 77-82.
110. Breslow NE, Day NE. *Statistical Methods in Cancer Research*. The design and analysis of cohort studies, Vol II. Lyon International Agency for Research on Cancer,1987.
111. Holme I, Helgeland A, Hjermann I, Leren P. The Oslo study: Social indicators, risk factors and mortality. In: *Medical aspects of mortality statistics*. Skandia International Symposia, Stockholm 1981;165-77.

112. Fylkesnes K, Johnson R, Førde OH. The Tromsø study: factors affecting patient-initiated and provider-initiated use of health care services. *Sociology of Health & Illness* 1992;14:275-91.
113. Rothman KJ. *Modern epidemiology*. Boston/Toronto: Little Brown & Company, 1986.
114. Talamini R, La Vecchia C, Decarli A, Negri E, Franceschi S. Nutrition, social factors and prostatic cancer in a Northern Italian population. *Br J Cancer* 1986; 53:817-21.
115. Giovannucci E, Rimm EB, Colditz GA, et al. A prospective study of dietary fat and risk of prostate. *J Natl Cancer Inst* 1993;85:1571-9.
116. Bjelke E. Dietary vitamin A and human lung cancer. *Int J Cancer* 1975;15:561-5
117. Ziegler RG, Mayne ST, Swanson CA. Nutrition and lung cancer. *Cancer Causes Control* 1996;7:157-77.
118. Whittemore AS, Wu-Williams AH, Lee M, et al. Diet, physical activity, and colorectal cancer among Chinese in North America and China. *J Natl Cancer Inst* 1990;82:915-26.
119. Gaard M, Tretli S, Løken EB. Dietary factors and the risk of colon cancer: a prospective study of 50,535 young Norwegian men and women. *Eur J Cancer Prev* (in press).
120. Giovannucci E, Rimm EB, Stampfer MJ, Colditz GA, Ascherio A, Willett WC. Intake of fat, meat, and fiber in relation to risk of colon cancer in men. *Cancer Res* 1994; 54:2390-7.
121. van Loon AJM, Burg J, Goldbohm RA, van den Brandt PA. Differences in cancer incidence and mortality among socio-economic groups. *Scand J Soc Med* 1995;23: 110-20.
122. Prener A, Hsieh C-C, Engholm G, Trichopoulos D, Jensen OM. Birth order and risk of testicular cancer. *Cancer Causes Control* 1992;3:265-72.
123. Doll R, Peto R. Cigarette smoking and bronchial carcinoma: dose and time relationships among regular smokers and lifelong non-smokers. *J Epidemiol Community Health* 1978;32:303-13.
124. Brownson RC, Chang JC, Davis JR. Gender and histologic type variations in smoking-related risk of lung cancer. *Epidemiology* 1992;3:61-4.
125. McDuffie HH, Klaassen DJ, Dosman JA. Determinants of cell type in patients with cancer of the lungs. *Chest* 1990;98:1187-93.
126. Meyer F, Moisan J, Marcoux D, Bouchard C. Dietary and physical determinants of menarche. *Epidemiology* 1990;1:377-81.

127. Akre O, Ekblom A, Hsieh C-C, Trichopoulos D, Adami HO. Testicular nonseminoma and seminoma in relation to perinatal characteristics. *J Natl Cancer Inst* 1996;**88**:883-9.
128. Ekblom A, Hsieh C-C, Lipworth et al. Perinatal characteristics in relation to incidence of and mortality from prostate cancer. *BMJ* 1996;**313**:337-41.
129. United Kingdom Testicular Cancer Study Group. Aetiology of testicular cancer: association with congenital abnormalities, age at puberty, infertility, and exercise *BMJ* 1994;**308**:1393-8.
130. Gallagher RP, Huchcroft S, Phillips N, et al. Physical activity, medical history, and risk of testicular cancer (Alberta and British Columbia, Canada). *Cancer Causes Control* 1995;**6**:398-406.
131. Hill AB. The environment and disease: association or causation? *Proceed Royal Soc Med* 1965;**58**:295-300.
132. Vena JE, Graham S, Zielezny M, Brasure J, Swanson MK. Occupational exercise and risk of cancer. *Am J Clin Nutr* 1987;**45**:318-27.
133. Brownson RC, Chang JC, Davis JR, Smith CA. Physical activity on the job and cancer in Missouri. *Am J Public Health* 1991;**81**:639-42.
134. Lee I-M, Paffenbarger Jr. RS, Hsieh C-C. Physical activity and risk of prostatic cancer among college alumni. *Am J Epidemiol* 1992;**135**:169-79.
135. Oliveria SA, Kohl III HW, Trichopoulos D, Blair SN. The association between cardiorespiratory fitness and prostate cancer. *Med Sci Sports Exerc* 1996;**28**:97-104.
136. Marchand LL, Kolonel LN, Yoshizawa CN. Lifetime occupational physical activity and prostate cancer risk [see comments]. *Am J Epidemiol* 1991;**133**:103-11.
137. Gerhardsson M, Norell SE, Kiviranta H, Pedersen NL, Ahlbom A. Sedentary jobs and colon cancer. *Am J Epidemiol* 1986;**123**:775-80.
138. Chow WH, Dosemeci M, Zheng W, et al. Physical activity and occupational risk of colon cancer in Shanghai, China. *Int J Epidemiol* 1993;**22**:23-9.
139. Wu AH, Paganini-Hill A, Ross RK, Henderson BE. Alcohol, physical activity and other risk factors for colorectal cancer: A prospective study. *Br J Cancer* 1987; **55**:687-94.
140. Slattery ML, Schumacher MC, Smith KR, West DW, Abd-Elghany N. Physical activity, diet, and risk of colon cancer in Utah. *Am J Epidemiol* 1988;**128**:989-99.
141. Gerhardsson M, Floderus B, Norell SE. Physical activity and colon

- cancer risk. *Int J Epidemiol* 1988;17:743-6.
142. Sellers TA, Potter JD, Folsom AR. Association of incident lung cancer with family history of female reproductive cancers: The Iowa Women's Health Study. *Genet Epidemiol* 1991;8:199-208.
143. Friedenreich CM, Rohan TE. Physical activity and risk of breast cancer. *Eur J Cancer Prev* 1995;4:145-51.
144. Zheng W, Shu XO, McLaughlin JK, Chow W-H, Gao YT, Blot WJ. Occupational physical activity and the incidence of cancer of the breast, corpus uteri, and ovary in Shanghai. *Cancer* 1993;71:3620-4.
145. Dosemeci M, Hayes RB, Vetter R et al. Occupational physical activity, socioeconomic status, and risk of 15 cancer sites in Turkey. *Cancer Causes Control* 1993;4:313-23.
146. Folsom AR, Jacobs DR, Wagenknecht LE, et al. Increase in fasting insulin and glucose over seven years with increasing weight and inactivity of young adults. The CARDIA study. *Am J Epidemiol* 1996;144:235-46.
147. Papa V, Reese CC, Brunetti A, Vigneri R, Siiteri PK, Goldfine ID. Progestins increase insulin receptor content and insulin stimulation of growth in human breast carcinoma cells. *Cancer Res* 1990;50:7858-62.
148. Tannenbaum A. The dependence of tumor formation on the composition of the calorie-restricted diet as well as on the degree of restriction. *Cancer Res* 1945;5:616-25.
149. Koga A, Kimura S. Influence of restricted diet on the cell cycle in the crypt of mouse small intestine. *J Nutr Sci Vitaminol*, Tokyo 1980;26:33-8
150. Lipman JM, Turturro A, Hart RW. The influence of dietary restriction on DNA repair in rodents: A preliminary study. *Mech Ageing Dev* 1989;48:135.
151. Ballard-Barbash R, Schatzkin A, Taylor PR, Kahle LL. Association of change in body mass with breast cancer. *Cancer Res* 1990;50:2152-5.
152. Nomura A, Stemmermann GN, Chyou PH, Marcus EB, Buist AS. Prospective study of pulmonary function and lung cancer. *Am Rev Respir Dis* 1991;144:307-11.
153. Baltrusch HJ, Stangel W, Titze I. Stress, cancer and immunity. New developments in biopsychosocial and psychoneuroimmunologic research. *Acta Neurol Napoli* 1991;13:315-27.
154. Knecht P, Raitasalo R, Heliövaara M, et al. Elevated lung cancer risk among persons with depressed mood. *Am J Epidemiol* 1996;144:1096-1103.

155. Thompson HJ, Westerlind KC, Snedden JR, Briggs S, Singh M. Inhibition of mammary carcinogenesis by treadmill exercise. *J Natl Cancer Inst* 1995;**87**:453-5.

156. Reddy BS, Sugie S, Lowenfels A. Effect of voluntary exercise on azoxymethane-induced colon carcinogenesis in male F344 rats. *Cancer Res* 1988;**48**:7079-81.

Paper I

Physical activity and the risk of prostate and testicular cancer: a cohort study of 53,000 Norwegian men

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The associations between recreational and occupational physical activity and the subsequent risk of prostate and testicular cancer were examined in a population-based cohort study of 53,242 men in Norway. Age at study entry was 19 to 50 years. Information on physical activity was based on questionnaire responses and a brief clinical examination. A total of 220 prostate and 47 testicular cancer cases were recorded in the Cancer Registry of Norway during a mean follow-up time of 16.3 years. We found a nonsignificant, reduced, adjusted relative risk (RR) of prostate cancer with increased level of physical activity at work and among those men with the greatest recreational physical activity. When occupational and recreational physical activity were combined, a reduced adjusted risk of prostate cancer was observed among men who walked during occupational hours and performed either moderate recreational activity (RR = 0.61, 95 percent confidence interval [CI] = 0.36 to 1.01) or regular recreational training (RR = 0.45, CI = 0.20 to 1.01) relative to sedentary men (test for trend, $P = 0.03$). Physically active men who were older than 60 years of age at diagnosis showed a reduced adjusted RR of borderline significance, while no association was observed for younger men. No evidence was found for any association between physical activity and testicular cancer regardless of physical activity at work and recreation. *Cancer Causes and Control* 1994, 5, 549 - 556

Key words: Cohort study, Norway, physical activity, prostate cancer, testicular cancer.

Introduction

Prostate cancer is the most frequent cancer among males in Norway, contributing 21 percent of all cancer-cases diagnosed in males in 1991; whereas testicular cancer was diagnosed in only two percent of all cancer cases in the same year.¹ Both cancer sites have shown increasing incidence rates during the last decades.¹ Carcinogenesis of the prostate and testis is still poorly understood. Since growth and tissue differentiation of

both organs is related mainly to the influence of sex hormones, it has been demonstrated that these factors also may have importance for tumor development of the prostate and testis.²⁻⁹

Physical activity is associated with both energy balance and hormones. Athletes have been shown to have lower basal levels of circulating testosterone than do untrained men.¹⁰⁻¹² In addition, increased physical

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activity can decrease body fat, thereby reducing the extragonadal production of estrogens. The effect of physical activity is, therefore, of considerable interest in the study of both prostate and testicular cancer in terms of understanding the carcinogenesis of the two organs and in cancer prevention.

Previous studies have reported decreased risk of prostate cancer,¹³⁻¹⁶ as well as increased risk or no effect in males with high physical activity.¹⁷⁻²¹ Brownson *et al*¹⁴ observed a decreased risk of testicular cancer with increased physical activity. Other studies have focused mainly on trauma in sports leading to increased risk of testicular cancer.²¹⁻²²

The aim of this study is to focus on the relationship between different levels of self-reported, occupational and recreational physical activity, and the subsequent risk of prostate and testicular cancer after adjusting for some possible risk factors. Since prostate cancer is especially frequent among older males, we were interested to see whether physical activity had different influences on prostate cancer risk in younger relative to older males.

Materials and methods

Cohort

This population-based cohort was initiated as a prospective study of risk factors for cardiovascular disease. Men and women from three counties in Norway (Oppland, Sogn and Fjordane, and Finnmark) and two cities (Oslo and Tromsø) were invited to participate in a health-screening program from 1972 to 1978 organized by the National Health Screening Service. In Tromsø, all men aged 20-49 years were invited, while in Oslo, men aged 40-49 were invited plus a seven-percent random sample of men aged 20-39. In the three counties of Oppland, Sogn and Fjordane, and Finnmark, all men and women aged 35-49 and a 10-percent random sample of persons aged 20-34 years were invited. In four small municipalities in Finnmark, all men and women aged 20-34 were invited. A total of 104,485 males and females were invited, and 53,622 males (73.5 percent) and 28,621 females (90.7 percent) attended the screening.

The screening procedure used was similar in the five geographic areas. Each person was invited by mail, with a cover letter and one-page questionnaire enclosed. The participants were asked to answer the questionnaire at home and bring it with them to the clinical examination. The clinical examination comprised checking the questionnaire for inconsistency, measurements of weight, height, and blood pressure, and collection of blood samples. Heart rate and other

measures of physical fitness were not assessed during the clinical examination.

The questionnaire covered the following: physical activity in recreational and occupational hours during the last year; history of chronic diseases, especially cardiovascular symptoms and diseases; and smoking habits and stress in daily life.

Self-reported physical-activity categories during recreational hours were graded from 1-4 according to which of the following categories best described the participant's usual level of recreational physical activity: R1 = reading, watching TV, or other sedentary activities; R2 = walking, bicycling, or physical activities for at least four hours a week; R3 = exercise to keep fit, participating in recreational athletics, etc., for at least four hours a week; R4 = regular hard training or exercise for competition several times a week. In the analysis, the categories R3 and R4 were merged due to few subjects in category R4 ($n = 1,316$).

Self-reported physical-activity categories during occupational hours were: O1 = mostly sedentary work; O2 = work with much walking; O3 = work with much lifting and walking; O4 = heavy manual work.

Identification of cases

The national 11-digit, personal identification number enabled a linkage to the Cancer Registry of Norway. This allowed for identification of every incident case of invasive prostate and testicular cancer that occurred in the cohort from the time of examination until the end of follow-up (31 December 1991). Those who emigrated or had a pre-existing malignancy or were diagnosed with a malignant disease within the first year after attending the cohort ($n = 380$) were excluded from the analyses. This reduced the possibility for any undiagnosed cancer to influence the level of physical activity. The 53,242 men eligible for analysis were then followed-up through the Norwegian Central Bureau of Statistics to identify deaths in the cohort to the end of 1991. Prostate and testicular cancers were coded according to an extended version of ICD-7.²³

Data analysis

Cox proportional-hazards regression technique was used to analyze the simultaneous effect of physical activity and covariates on prostate and testicular cancer incidence in the cohort. Person-years at risk of developing prostate or testicular cancer were calculated as the number of years from the time of entry until the time of withdrawal (year of diagnosis, time of death, or end of follow-up). Ten percent (5,092 men) died during follow-up.

As no cases of prostate cancer were observed among

men younger than age 35 years at entry, this age-group was excluded from the analyses of prostate cancer ($n = 9,557$). These analyses, therefore, were based on a limited number of men concerning the risk of prostate cancer ($n = 43,685$), but not of testicular cancer ($n = 53,242$). All analyses were adjusted for age at entry, geographic regions, and obesity (using body mass index [BMI] of weight/height²). Because of missing data, the number of subjects included in the individual analyses varies slightly.

We divided the cohort in two categories of age at diagnosis in order to analyze a possible different influence of physical activity on the prostate cancer risk in younger (aged under 60 years) relative to older (60 or more years) males. Mean age at diagnosis for prostate cancer was 60.3 years and few prostate-cancer cases were older than 65 years. Other cutoff points were considered without extended information. Age at entry was adjusted for as a continuous variable in this model. The analyses were performed with the Proc Phreg procedure available in the SAS statistical package.²⁴

Results

A total of 220 prostate and 47 testicular cancer-cases were diagnosed during a mean follow-up time of 16.3 years (Table 1). All testicular cancers and 99 percent of prostate cancers were histologically verified. We found strongly increasing incidence-rates of prostate cancer with increasing age at entry in contrast to decreasing incidence rates with increasing age at entry of testicular cancer. Due to the young age at entry and the relatively short follow-up, the median age at diagnosis for prostate cancer was 61.2 years (42.9-68.5 years) while testicular cancer was diagnosed in men aged 25.5-65.7 years old with a median age at diagnosis of 45.8 years.

In univariate analysis, only age and BMI were significant risk factors for prostate cancer, while a negative association was observed between age and risk of testicular cancer (Table 2). None of the variables in Table 2 gave any significant deviation from linearity by adding a second-order term (results not shown).

When calculating the crude incidence rate of prostate cancer in different occupational activity categories,

Table 1. Number of persons, person-years (PY) at risk, and prostate and testicular cancer by age at entry, Norway

Age at entry (yrs)	Number of persons	Mean follow-up (yrs)	Prostate cancer		Testicular cancer	
			No.	Incidence per 100,000 PYs	No.	Incidence per 100,000 PYs
19-24	2,648	16.9	0	0.0	6	13.4
25-29	3,459	17.0	0	0.0	7	11.9
30-34	3,450	16.7	0	0.0	3	5.2
35-39	9,436	15.8	7	4.7	10	6.7
40-44	15,930	16.5	48	18.3	12	4.6
45-49	17,629	16.1	152	53.6	9	3.2
50-54	690	15.8	13	119.1	0	0.0
Total	53,242	16.3	220	25.4	47	5.4

Table 2. Age-adjusted relative risk (RR) of prostate and testicular cancer in relation to possible risk factors; Cox proportional hazards model; Norway

Variable	Prostate cancer				Testicular cancer			
	Cases	No. of persons	RR ^a	(CI) ^b	Cases	No. of persons	RR ^a	(CI) ^b
Age at entry (per yr)	220	43,685	1.27	(1.21-1.33)	47	53,242	0.95	(0.92-0.98)
BMI (kg/m ²) per unit	217	42,851	1.25	(1.05-1.50)	46	53,242	1.32	(0.82-2.14)
Cholesterol (mmol/liter) per unit	220	43,685	0.98	(0.88-1.08)	47	53,242	1.07	(0.87-1.32)
Triglycerides (mmol/liter) per unit	220	43,685	1.05	(0.99-1.11)	47	53,242	1.05	(0.92-1.20)
Glucose (mmol/litre) per unit	220	43,685	0.98	(0.91-1.06)	47	53,242	1.08	(0.97-1.22)
Height (cm) per every 10 cm	217	42,859	0.99	(0.82-1.19)	46	52,199	1.12	(0.75-1.67)
Smoking per 10 cigarettes	211	42,067	1.08	(0.90-1.30)	46	51,532	1.20	(0.85-1.70)
Married/separated (cf never married)	220	43,685	1.63	(0.96-2.75)	47	53,242	0.72	(0.35-1.45)

^a Adjusted for age at entry.

^b CI = 95% confidence interval.

Table 3. Person-years (PY), crude incidence rates, and adjusted relative risk (RR) of prostate and testicular cancer with 95% confidence interval (CI) associated with occupational and recreational physical activity; Cox proportional hazard model; Norway

Physical activity (PhA)	PYs	Cancer cases	Crude incidence rate per 100,000 PYs	RR ^a	(CI) ^b
Prostate cancer					
Occupational PhA					
Sedentary	250,772	92	36.7	1.00	(Ref)
Walking	185,938	49	26.4	0.77	(0.54-1.09)
Lifting and walking	149,369	46	30.8	0.99	(0.69-1.42)
Heavy manual	111,533	25	22.4	0.81	(0.50-1.30)
Recreational PhA					
Sedentary	136,324	44	32.3	1.00	(Ref)
Moderately active	395,406	132	33.4	1.03	(0.73-1.45)
Regular training	169,371	41	24.2	0.87	(0.57-1.34)
Testicular cancer					
Occupational PhA					
Sedentary	309,845	13	4.2	1.00	(Ref)
Walking	225,925	6	2.7	0.60	(0.23-1.57)
Lifting and walking	189,190	13	6.9	1.38	(0.63-3.01)
Heavy manual	130,126	13	10.0	1.95	(0.86-4.41)
Recreational PhA					
Sedentary	170,718	8	4.7	1.00	(Ref)
Moderately active	471,125	26	5.5	1.22	(0.55-2.69)
Regular training	217,200	12	5.5	1.01	(0.41-2.49)

^a Adjusted for age at entry, geographic region, and body mass index (BMI).

^b CI = 95% confidence interval.

there was a lower incidence rate among those walking, lifting and walking combined, and, in particular, among those doing heavy manual work, compared with the sedentary group (Table 3). We observed a 25 percent lower crude incidence-rate of prostate cancer among those doing regular physical training during recreational hours compared with the sedentary and the moderately active group (Table 3). The crude incidence rate of testicular cancer was about two times higher among men with heavy manual work compared with men with sedentary work (Table 3).

We observed a nonsignificant, but consistent reduction in relative risk (RR) of prostate cancer after adjusting for age at entry, BMI, and geographic region among those walking or doing heavy manual work compared with the sedentary working-group (Table 3). A nonsignificant reduction in RR of prostate cancer also was observed among recreational-active men compared with the recreational-sedentary group (Table 3). Recreational activity did not seem to influence the adjusted RR of testicular cancer to any significant extent. However, close to twofold increased RR among men with heavy manual work was observed.

In order to study total physical activity, recreational (R) and occupational (O) activity were combined. We observed that, for prostate cancer, all physically non-

sedentary groups except for one group (RR = 0.94) were at a lower risk when the overall sedentary men both at work and leisure was the reference group (Table 4). Among men walking during occupational hours combined with a low or moderate recreational physical activity, we observed a borderline significant reduced RR (RR = 0.61, 95 percent confidence interval [CI] = 0.36-1.01). Even lower relative risk (RR = 0.45, CI = 0.20-1.01) was observed in the group of men walking at work combined with high recreational physical-activity (test for trend $P = 0.03$). Performing analysis separately for R1 and R2 in the combined moderate-activity risk groups did not give other information than increased confidence intervals.

Men older than 60 years of age at diagnosis, who were occupationally and recreationally physically-active, had a 39 percent reduced adjusted RR of prostate cancer, while younger males only had 11 percent reduced adjusted risk compared with the most sedentary group (borderline significance) (Table 5). Including interaction terms as age-activity and age-BMI in the models, did not influence the risk estimates.

Men with heavy manual work were at about 85-95 percent increased risk of testicular cancer. However, due to small numbers, none of the estimates was statistically significant (Table 6).

Table 4. Adjusted relative risk (RR) of prostate cancer with 95% confidence interval (CI) of combined occupational and recreational physical activity among men aged 35-50 years at entry; Cox proportional hazard model; Norway

Physical activity ^a		Cases	RR ^b	(CI)	Trend test P value
Occupational (O)	Recreational (R)				
Sedentary (O1)	Sedentary (R1)	23	1.0	(Ref)	
Sedentary (O1)	Moderate (R2)	53	0.78	(0.47-1.27)	
Sedentary (O1)	Active (R3)	16	0.64	(0.34-1.22)	0.17
Walking (O2)	Sedentary/moderate (R1 + R2)	41	0.61	(0.36-1.01)	
Walking (O2)	Active (R3)	8	0.45	(0.20-1.01)	0.03
Lifting/walking (O3)	Sedentary/moderate (R1 + R2)	37	0.81	(0.48-1.37)	
Lifting/walking (O3)	Active (R3)	9	0.77	(0.35-1.71)	0.44
Heavy manual (O4)	Sedentary/moderate (R1 + R2)	17	0.71	(0.34-1.48)	
Heavy manual (O4)	Active (R3)	8	0.94	(0.38-2.36)	0.77

^a R1 = Recreational sedentary; R2 = Recreational moderate active; R3 = Recreational training.

^b Adjusted for age at entry, geographic region, and body mass index (BMI).

Table 5. Adjusted relative risk (RR) of prostate cancer in different age-groups at diagnosis according to combined occupational and recreational physical activity; Cox proportional hazard model; Norway

Age at diagnosis (yrs)	Cases	Person-years at risk	Sedentary ^a RR ^a	Moderate/active ^c	
				RR ^a	(CI) ^d
< 60	84	603,518	1.0	0.89	(0.40-1.94)
60 +	128	85,755	1.0	0.61	(0.36-1.04)

^a Sedentary during both occupation (O1) and recreation (R1).

^b Adjusted for age at entry, geographic region, and BMI.

^c Moderate/active = all other groups than sedentary.

^d CI = 95% confidence interval.

Discussion

This study suggests that recreational activity in particular, but also occupational activity, may reduce the risk of prostate cancer, especially among the young elderly. When occupational and recreational physical-

activity were combined, a reduced risk (nonsignificant) of prostate cancer was observed in all active groups except one group relative to the sedentary group. A significant dose-response association between physical activity and risk of prostate cancer was observed among men walking during occupational hours, with increasing level of physical activity at leisure. No obvious, significant, statistical effect of physical activity on the risk of testicular cancer was observed though a close to twofold increased risk was observed in the group with heavy manual work.

The strength of this study is the population-based approach and the large sample size of the cohort including more than 50,000 men. In addition, both recreational and occupational physical-activities are taken into account giving a comprehensive consideration of total physical activity of each individual. In this way, we were able to analyze a greater range of the exposure variable.

Table 6. Adjusted relative risk (RR) of testicular cancer with 95% confidence interval (CI) of combined occupational and recreational physical activity among men aged 19-50 years at entry; Cox proportional hazard model; Norway

Physical activity ^a		Cases	RR ^b	(CI)	Trend test P value
Occupational (O) +	Recreational (R) ^a				
Sedentary (O1)	Sedentary/moderate (R1 + R2)	10	1.0	(Ref)	
Sedentary (O1)	Active (R3)	3	0.77	(0.21-2.84)	
Walking (O2)	Sedentary/moderate (R1 + R2)	4	0.52	(0.16-1.66)	
Walking (O2)	Active (R3)	2	0.66	(0.14-3.09)	0.37
Lifting/walking (O3)	Sedentary/moderate (R1 + R2)	10	1.44	(0.59-3.55)	
Lifting/walking (O3)	Active (R3)	3	1.19	(0.31-4.47)	0.80
Heavy manual (O4)	Sedentary/moderate (R1 + R2)	9	1.85	(0.67-5.09)	
Heavy manual (O4)	Active (R3)	4	1.95	(0.55-6.94)	0.23

^a R1 = Recreational sedentary; R2 = Recreational moderate active; R3 = Recreational training.

^b Adjusted for age at entry, geographic region, and body mass index (BMI).

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Also of importance is that the reporting and recording of all new cases of prostate and testicular cancer in Norway is based on a very strict system. This includes compulsory reporting by hospital departments, pathology laboratories, and death certificates, thus achieving a very high case-ascertainment in addition to the close to 100 percent histologic verification of the cancer cases.

However, due to the relatively young age at entry and the short follow-up, the numbers of cases of prostate cancer and especially testicular cancer are small, thereby limiting the statistical power of the study.

The questionnaire used in this study has been validated extensively for the accurate assessment of physical activity in several studies.²⁵⁻²⁹ Based on the same questionnaire, Løchen *et al.*²⁹ demonstrated that physical fitness increased with activity in leisure time. Physical fitness was measured as physical working capacity in a graded bicycle exercise. There is good reason to believe that the allocation of the individuals to the respective groups reflects real differences in the level of physical activity in the study population.

Nondifferential misclassification may occur when a person describes himself as more physically active than he is. The risk estimates become closer to 1.0, contributing to a reduced statistical significance for prostate cancer. In addition, a stronger effect probably would be achieved if the evaluation of physical activity had been repeated. This was demonstrated by Lee *et al.*¹⁵ who observed a far greater effect of physical activity when two activity assessments with 10 years' interval were combined.

Socioeconomic factors have been observed to influence the risk of both prostate and testicular cancer in some studies, but not in others.^{9,30-33} No geographic or urban-rural differences were observed in our study, and adjustments for marital status did not influence the association between physical activity and risk of either prostate or testicular cancer. However, due to different age at study entry, and variation of incidence rate in the different geographic regions (results not shown), we chose to adjust for geographic region in the multivariate analysis. In addition, partly based on the same population, Holme *et al.*²⁸ observed that higher social classes were overrepresented among men who reported sedentary work activity and that those who were sedentary at leisure were overrepresented in lower social classes. Therefore, the reference group used (the overall sedentary men when combined with recreational and occupational activity) reflects both high and low social classes. It is less likely, therefore, that social class explains the observed association.

Confounding by diet may be important, as dietary fat has been found to increase the risk of prostate can-

cer.^{8,32} Unfortunately, dietary information was not available in this study. Obesity, measured as BMI, did not contribute to our results when this variable was adjusted for in the analysis. However, BMI independently had a positive association with the risk of prostate cancer.

Prostate cancer

The present results are in agreement of those of Albanes *et al.*¹³ and Lee *et al.*¹⁵ who observed an increased risk of prostate cancer among inactive men. One reason why those men who reported walking (O2) during occupational hours combined with high recreational activity were at lowest risk (RR = 0.45) in this study may be that they perform more dynamic activity relative to the other occupational groups. The heavy manual-labor group may perform more static activity which may not influence prostate cancer risk in the same way.

The biologic implication of these findings may be related to different mechanisms. The incidence of prostate cancer is high in countries with a Western lifestyle. Obesity as the result of high caloric intake, sedentary work, and low leisure activity are important elements of this lifestyle. The bearing of physical activity on the carcinogenesis of the prostate thus may reflect the energy imbalance with decreased metabolism resulting in an increase in adipose tissue. Lew *et al.*³⁴ found that overweight men had 30 percent increased mortality rates of prostate cancer. The increase in BMI significantly contributed to increased risk of prostate cancer with about 25 percent in univariate analysis in this study. However, in our study, the effect of physical activity on the risk of prostate cancer was independent of BMI, as this variable was adjusted for in the analysis. It is reasonable to believe that physical activity acts through additional or other mechanisms than those related merely to energy imbalance.

Any influence of physical activity on the level of sex-hormones is of special interest, with testosterone and its metabolite being particularly important. A decrease in these androgens immediately after prolonged exercise has been observed in males,^{12,35} and several studies^{11,36-38} have recorded lower basal levels of testosterone among trained men relative to untrained men. Black men in United States, who have a high risk of prostate cancer, have 15 percent higher levels of testosterone in serum than White men.³ This strengthens the importance of testosterone in the pathogenesis of the disease and could be the mediator for the effect of physical activity.

We observed that the preventive effect of physical activity was higher among those older than 60 years than among younger men. This is in agreement with

Lee *et al.*,¹⁵ who observed that the protective effect of total physical activity was limited to those older than 70 years of age. Lee *et al.* also argued that long-term maintenance of high levels of physical activity may be necessary for reducing the risk of prostate cancer. The fact that most men were relatively young at the end of the follow-up may explain why a stronger association with physical activity was not detected.

In contrast, Marchand *et al.*²⁰ observed an opposite effect as they found a negative association between years spent in sedentary occupations among men aged 70 years or older, while the results for younger men were less clear. They evaluated only occupational activity, not recreational activity, which may explain the divergent results. Physical activity has probably only a weak or moderate influence on the carcinogenesis of prostate cancer. Thus, physical activity may be of greater importance among the elderly (in whom environmental and endogenous hormonal factors may be more prevailing) relative to younger men (in whom the importance of genetic predisposition may be greater and less likely to be influenced by physical activity). Further, Zumoff *et al.*³⁹ observed that young prostate-cancer patients differed markedly in their endogenous hormonal pattern from patients aged 65 or older. The level of testosterone in serum in males declines by increasing age.^{40,41} If physical activity contributes to a further suppression of the age-dependent, decreasing, testosterone-curve, this also may explain why the effect of physical activity is observed among the elderly only.

Testicular cancer

Physical activity has been suggested to act as a persistent trauma to the scrotum and thus increase the risk of testicular cancer.^{21-22,42} This effect has been found for horse riding and bicycling.²¹ We had no such information available and it may explain why no statistical association between physical activity and risk of testicular cancer was found in this study. It is reasonable to suggest that a true biologic effect of some types of physical activities on cancer risk probably will disappear if the effect is of minor strength or the particular activity is insufficiently represented in the study population. Another explanation for the lack of association may be the late assessment of physical activity relative to the highest cancer-risk period in this study. The assessment of physical activity was performed in males at age 20 to 49 years which also is the period with the highest incidence rate of testicular cancer. Thus, physical activity may have a decreased influence on testicular cancer risk, as the carcinogenic process may already have emerged.

The small number of testicular cancer cases limits the statistical power of the study. Also of importance may be a too-small range of variation in the assessment of physical activity which is not sensitive enough to produce statistical associations. A positive association of physical activity on the risk of testicular cancer has been demonstrated by Brownson *et al.*¹⁴ An influence of physical activity on the risk of testicular cancer may be present also in this study although none of the risk estimates were of statistical significance. Indication for this may be a close to twofold increased risk observed among individuals performing heavy manual work.

Conclusion

Our study suggests an inverse association between both recreational and occupational physical-activity and the risk of prostate cancer. This effect may be related to energy imbalance, but more likely to be a hormonal influence on the carcinogenesis of the prostate. In contrast, no obvious statistical association between physical activity and the risk of testicular cancer was observed. Further studies are needed where the assessment of both physical activity and fitness are taken into account combined with hormonal and dietary status and metabolic measurements.

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References

1. Cancer Registry of Norway. *Incidence of Cancer in Norway, 1991*. Oslo, Norway: Cancer Registry, 1993.
2. Noble RL. The development of prostatic adenocarcinoma in Nb rats following prolonged sex hormone administration. *Cancer Res* 1977; 37: 1929-33.
3. Ross R, Bernstein L, Judd H, Hanisch R, Pike M, Henderson B. Serum testosterone levels in healthy young black and white men. *JNCI* 1986; 76: 45-8.
4. Barrett-Connor E, Garland C, McPhillips JB, Khaw KT, Wingard DL. A prospective, population-based study of androstenedione, estrogens and prostatic cancer. *Cancer Res* 1990; 50: 169-73.
5. Hsing AW, Comstock GW. Serological precursors of cancer: serum hormones and risk of subsequent prostate cancer. *Cancer Epidemiol Biomarkers Prev* 1993; 2: 27-32.
6. Boyle P, Zaridze DG. Risk factors for prostate and testicular cancer. *Eur J Cancer* 1993; 7: 1048-55.
7. Moss AR, Osmond D, Bachetti P, Torti PM, Gurgin V. Hormonal risk factors in testicular cancer. A case-control study. *Am J Epidemiol* 1986; 124: 39-52.

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8. Gershman S, Stolley P. A case-control study of testicular cancer using Connecticut tumour registry data. *Int J Epidemiol* 1988; 17: 738-42.
9. Prener A, Hsieh C-c, Engholm G, Trichopoulos D, Jensen OM. Birth order and risk of testicular cancer. *Cancer Causes Control* 1992; 3: 265-72.
10. Adlercreutz H, Harkonen M, Kuoppasalmi K, et al. Effect of training on plasma anabolic and catabolic steroid hormones and their response during physical exercise. *Int J Sports Med* 1986; 7: 27-8.
11. Hackney AC, Sinning WE, Bruot BC. Reproductive hormonal profiles of endurance-trained and untrained males. *Med Sci Sports Exerc* 1988; 20: 60-5.
12. Remes K, Kuoppasalmi K, Adlercreutz H. Effect of physical exercise and sleep deprivation on plasma androgen levels: Modifying effect of physical fitness. *Int J Sports Med* 1985; 6: 131-5.
13. Albanes D, Blair A, Taylor PR. Physical activity and risk of cancer in the NHANES I population. *Am J Public Health* 1989; 79: 744-50.
14. Brownson RC, Chang JC, Davies JR, Smith CA. Physical activity on the job and cancer in Missouri. *Am J Public Health* 1991; 81: 639-42.
15. Lee IM, Paffenbarger RS, Hsieh C-c. Physical activity and risk of prostatic cancer among college alumni. *Am J Epidemiol* 1992; 135: 169-79.
16. Hsing AW, McLaughlin JK, Zheng W, Gao YT, Blot WJ. Occupation, physical activity, and risk of prostate cancer in Shanghai, People's Republic of China. *Cancer Causes Control* 1994; 5: 136-40.
17. Poldenak AP. College athletics, body size and cancer mortality. *Cancer* 1976; 38: 382-7.
18. Paffenbarger RS, Hyde RT, Wing AL. Physical activity and incidence of cancer in diverse populations: a preliminary report. *Am J Clin Nutr* 1987; 45: 12-7.
19. Marchand LL, Kolonel LN, Yoshizawa CN. Lifetime occupational physical activity and prostate cancer risk. *Am J Epidemiol* 1991; 133: 103-11.
20. Severson RK, Nomura AMY, Grove JS, Stemmermann GN. A prospective analysis of physical activity and cancer. *Am J Epidemiol* 1989; 130: 522-9.
21. Coldman AJ, Elwood JM, Gallagher RP. Sports activities and risk of testicular cancer. *Br J Cancer* 1982; 46: 749-56.
22. Haughey BP, Graham S, Brasure J, Zielezny M, Sufrin G, Burnett WS. The epidemiology of testicular cancer in upstate New York. *Am J Epidemiol* 1989; 130: 25-36.
23. World Health Organization. *International Classification of Diseases, Seventh Revision*. Geneva, Switzerland: WHO, 1957.
24. SAS Institute, Inc. *SAS/STAT Guide for Personal Computers. Version 6 Edition*. Cary, NC (USA): SAS Institute, Inc., 1992.
25. Saltin B, Grimby G. Physiological analysis of middle-aged and old former athletes. *Circulation* 1968; 38: 1104-15.
26. Wilhelmsen L, Tibblin G, Aurell M, Bjure J, Ekström-Jodal B, Grimby G. Physical activity, physical fitness and risk of myocardial infarction. *Adv Cardiol* 1976; 18: 217-30.
27. Bjartveit K, Foss OP, Gjervig T. The cardiovascular disease study in Norwegian counties. Results from first screening. *Acta Med Scand* 1979; 634.
28. Holme I, Helgeland A, Hjermann I, Leren P, Lund-Larsen PG. Physical activity at work and at leisure in relation to coronary risk factors and social class. A 4-year mortality follow-up. The Oslo Study. *Acta Med Scand* 1981; 209: 277-83.
29. Løchen ML, Rasmussen K. The Tromsø study: physical fitness, self-reported physical activity, and their relationship to other coronary risk factors. *J Epidemiol Commun Health* 1992; 26: 103-7.
30. Yu H, Randall EH, Wynder EL. Case-control study of prostate cancer and socioeconomic factors. *Prostate* 1988; 13: 317-25.
31. Swerdlow AJ, Douglas AJ, Huttley SRA, Smith PG. Cancer of the testis, socioeconomic status, and occupation. *Br J Ind Med* 1991; 48: 670-4.
32. Talamini R, La Vecchia C, DeCarli A, et al. Nutrition, social factors and prostatic cancer in a Northern Italian population. *Br J Cancer* 1986; 53: 817-21.
33. Fincham SM, Hill GB, Hanson J, Wijayasinghe C. Epidemiology of prostatic cancer: A case-control study. *Prostate* 1990; 17: 189-206.
34. Lew EA, Garfinkel L. Variations in mortality by weight among 750,000 men and women. *J Chron Dis* 1979; 32: 563-76.
35. Aakvaag A, Sand T, Opstad PK, Fonnum F. Hormonal changes in young men during prolonged physical strain. *Eur J Appl Physiol* 1978; 39: 283-91.
36. Morville R, Pesquies PC, Guezennec CY, Serrurier BD, Guignard M. Plasma variations in testicular and adrenal androgens during prolonged physical exercise in man. *Annales d'Endocrinologie* 1979; 40: 501-10.
37. Wheeler GD, Wall SR, Belcastro AN, Cumming DC. Reduced serum testosterone and prolactin levels in male distance runners. *JAMA* 1984; 252: 514-6.
38. Strauss RH, Lanese RR, Malarkey WB. Weight loss in amateur wrestlers and its effect on serum testosterone levels. *JAMA* 1985; 254: 3337-8.
39. Zumoff B, Levine J, Starin GW, et al. Abnormal levels of plasma hormones in men with prostate cancer: Evidence toward a "two-disease" theory. *Prostate* 1982; 3: 579-88.
40. Pirke K, Dorr P. Age related changes and interrelationships between plasma testosterone, oestradiol and testosteronebinding globulin in normal adult males. *Acta Endocrinol* 1973; 73: 792-800.
41. Dai WS, Kuller LH, Laporte RE, Gutai JP, Falvo-Gerard L, Caggiula A. The epidemiology of plasma testosterone levels in middle-aged men. *Am J Epidemiol* 1981; 114: 804-16.
42. Brown LM, Pottern LM, Hoover RN. Testicular cancer in young men: the search for causes of the epidemic increase in United States. *J Epidemiol Commun Health* 1987; 41: 349-54.

Paper II

Physical activity and risk of colorectal cancer in men and women

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Summary We examined the association between self-reported occupational and recreational physical activity and the subsequent risk of colorectal cancer in a population-based cohort in Norway. During a mean follow-up time of 16.3 years for males and 15.5 years for females, 236 and 99 colon cancers and 170 and 58 rectal cancers were observed in males and females, respectively, among 53 242 males and 28 274 females who attended the screening between 1972 and 1978. Physical activity at a level equivalent to walking or bicycling for at least four hours a week during leisure-time was associated with decreased risk of colon cancer among females when compared with the sedentary group (RR = 0.62, 95% CI 0.40–0.97). Reduced risk of colon cancer was particularly marked in the proximal colon (RR = 0.51, 95% CI 0.28–0.93). This effect was not observed for occupational physical activity alone, probably due to a narrow range of self-reported physical activity at work among females. However, by combining occupational and recreational physical activity we observed an inverse dose–response effect as increasing total activity significantly reduced colon cancer risk (P for trend = 0.04). Among males 45 years or older at entry to the study, an inverse dose–response effect was observed between total physical activity and colon cancer risk (P for trend = 0.04). We also found in males a stronger preventive effect for physical activity in the proximal as compared to distal colon. In addition, we found a borderline significant decrease in colon cancer risk for occupational physical activity in males 45 years or older when compared to the sedentary group (RR = 0.74, 95% CI 0.53–1.04). All results were adjusted for age, body mass index, serum cholesterol and geographic region. No association between physical activity and rectal cancer was observed in males or females. The protective effect of physical activity on colon cancer risk is discussed in regard to energy balance, dietary factors, age, social class, body mass index and gastrointestinal transit time.

Keywords: physical activity; colorectal cancer; cohort study; gender differences; subsites

Cancer of the large intestine is one of the most common neoplasms in western countries (Muir *et al.*, 1987; Engeland *et al.*, 1993). Recently, the role of exercise in the aetiology of colon carcinogenesis has drawn particular interest. A growing number of epidemiological studies have reported a protective effect of occupational physical activity on colon cancer risk (Garabrant *et al.*, 1984; Gerhardsson *et al.*, 1986; Brownson *et al.*, 1989; Peters *et al.*, 1989; Arbman *et al.*, 1993; Chow *et al.*, 1993; Fraser and Pearce, 1993). Others have observed that recreational physical activity protects against colon cancer (Wu *et al.*, 1987; Slattery *et al.*, 1988; Gerhardsson *et al.*, 1988; Severson *et al.*, 1989; Ballard-Barbash *et al.*, 1990; Lee *et al.*, 1991; Markowitz *et al.*, 1992; Giovannucci *et al.*, 1995). In contrast, the association between physical activity and risk of rectal cancer is more inconsistent (Vena *et al.*, 1985; Gerhardsson *et al.*, 1986; Fraser and Pearce, 1993).

However, few studies have analysed the association between physical activity and colon cancer risk in females or have taken gender differences, age and subsites into consideration. In addition, patients with proximal colon cancer are older than patients with distal colon and rectal cancer, and women make up a higher percentage of patients with cancer in the proximal colon (Møller Jensen, 1984; Halvorsen, 1986; Flesher *et al.*, 1989). Furthermore, physiological differences in the proximal and the distal colon may reflect different susceptibility to neoplastic transformation (Bufill, 1990; Dubrow *et al.*, 1993).

We therefore investigated the association between self-reported physical activity both during leisure and work and the subsequent risk of colorectal cancer in a population-based, prospective study among both sexes. We further examined whether physical activity had a different effect according to age-, gender- and site-specific colorectal cancer risk.

Material and methods

Between 1972 and 1978, 104 485 males and females from five geographical areas in Norway—Oslo, Oppland, Sogn and Fjordane, Tromsø and Finnmark—were invited to participate in a population-based health survey of risk factors for cardiovascular disease. In Tromsø, all men aged 20–49 years were invited, while in Oslo men aged 40–49 were invited plus a 7% random sample of men aged 20–39. In the three counties of Oppland, Sogn and Fjordane and Finnmark all men and women aged 35–49 and a 10% random sample of persons aged 20–34 years were invited. In four small municipalities in Finnmark all men and women aged 20–34 were invited: a total of 104 485, of whom 53 622 males (73.5%) and 28 621 females (90.7%) attended the screening.

The screening procedures were similar in the five areas. Each person was invited by mail, with a covering letter and one-page questionnaire enclosed. The participants were asked to answer the questionnaire at home and bring it to the clinical examination. The clinical examination consisted of checking the questionnaire for inconsistency, measurements of weight, height and blood pressure, and the collection of blood samples. Heart rate and other measures of physical fitness were not assessed.

The questionnaire covered the following: physical activity (PhA) during recreational (R) and occupational (O) hours in the last year; history of chronic diseases especially cardiovascular symptoms and diseases, smoking habits and stress in daily life.

Self-reported physical activity categories during recreational hours were graded from 1 to 4 according to which of the following categories best described the participant's usual level of physical activity: R1 = reading, watching TV or other sedentary activities; R2 = walking, bicycling or physical activities for at least four hours a week; R3 = exercise to keep fit, participating in recreational athletics etc. for at least four hours a week; R4 = regular hard training or participation in competitive sports several times a week.

Self-reported physical activity during occupational hours

was divided into four categories; O1 = mostly sedentary work; O2 = work with much walking; O3 = work with much lifting and walking; O4 = heavy manual work.

The national 11-digit personal identification number enabled a linkage to the Cancer Registry of Norway. This allowed for identification of every incident case of colorectal cancer that occurred in the cohort from the time of examination until the end of follow-up (31 December 1991). Colorectal cancers were coded according to ICD7. In some analyses, cancers in the colon were categorised as occurring in the proximal colon (153.0+153.1), or the distal colon (153.2+153.3+153.4). Cases identified only incidentally at autopsy were not included. Histological confirmation was obtained in 95% of the cases and among these 96.7% were adenocarcinomas and eight cases (2.5%) were classified as malignant carcinoid tumours.

In addition all 53 622 men and 28 621 women were followed up through the Norwegian Central Bureau of Statistics to identify deaths in the cohort up to the end of 1991. Those who emigrated or had a pre-existing malignancy or developed a malignancy within the first year of the study (males, $n=380$; females, $n=347$) were excluded from the analyses. This reduced the possibility for any undiagnosed cancer to influence the level of physical activity. The present cohort study is restricted to males and females aged 20–69 years in the follow-up period. Included for analysis were 53 242 males (867 822 person-years) and 28 274 females (437 785 person-years).

Cox's proportional hazards regression techniques were used to analyse the simultaneous effects of physical activity and possible confounders on colon and rectal cancer incidence in the cohort. In these analyses, the categories R3 and R4 of recreational physical activity were merged due to small numbers in category R4 (males, $n=316$; females, $n=62$). Observation years at risk of developing colon or rectal cancer were calculated as the number of years from 1 year after study entry until the time of withdrawal (year of diagnosis of cancer, time of death or end of follow-up in December 1991, which ever was earliest). In the sex-specific analyses, we adjusted for attained age (continuous variable), geographical regions and obesity at time of measurements. As a measure of obesity, we used the body mass index (BMI) ($\text{weight height}^{-2}$).

To study the influence of total physical activity on colon cancer risk, occupational (O) and recreational (R) physical activity were combined. As a reference group (R1/O1+O2), we used sedentary leisure (R1) and both sedentary (O1) and moderate (O2) activities at work in order to increase the number of persons in the reference group.

We examined models stratified by age at entry (<45 years, ≥ 45 years) and BMI (median split and tertiles) to analyse if there was any effect modification by age and BMI. Other cut-off points for age were considered without extended information. These analyses were performed with the Proc

Phreg procedure in the SAS statistical package (SAS Institute, 1992). Owing to missing data, the number of subjects included in the individual analyses varies slightly.

Results

A total of 236 colon and 170 rectal cancers among males and 99 colon and 58 rectal cancers among females were diagnosed in the study population during a mean follow-up time of 16.3 years and 15.5 years in males and females respectively. Median age at diagnosis for colon cancer was 58.1 years in males and 54.6 years in females. For rectal cancer the median age at diagnosis was 57.3 years and 55.4 years in males and females respectively. Of all cases of colorectal cancer, cases of proximal colon cancer were less frequent among males (23.4%) than among females (30.5%), whereas the proportion of distal colon cancer was reversed between the two sexes (31.8% vs 28.7%).

The grade of physical activity was differently distributed in males and females. Two-thirds of the females and 76% of the housewives reported frequent walking (O2) during occupational hours in contrast to only one-quarter among males (Table 1). Fewer females than males reported sedentary work (O1). Gender differences were also observed during leisure time as only 10% of females reported regular training (R3+R4) in contrast to 25.4% of males.

Age at entry was a significant risk factor in univariate analyses for both colon and rectal cancer in both sexes (Table 11). A positive association was observed between body mass index (BMI) and colon cancer risk in males, but not in females. None of the variables in Table 11 significantly deviated from linearity when a second-order term was introduced (results not shown).

Total physical activity (occupational and recreational combined) showed an overall negative dose-response relationship with colon cancer risk among females (P for trend = 0.04), but not in males (Table 11).

We analysed colon cancer risk in relation to a possible age effect of total physical activity by dividing the sex-specific cohort into those younger and older than 45 years at study-entry. Among males 45 years or older at study entry (median age at diagnosis = 60.0 years), we observed a negative dose-response relationship between total physical activity and colon cancer risk (P for trend = 0.04), which was not observed among males younger than 45 years at study entry (median age at diagnosis = 52.1 years) (Table 1V). In addition, a borderline significant reduction on total colon cancer risk was observed among occupationally physically active males (O2, O3, O4) 45 years or older at study entry compared with the sedentary ones (RR = 0.74, 95% CI 0.53–1.04) (results not shown in Table 1V). No similar age effect was observed in females.

Table 1 Self-reported physical activity during occupational (O) and recreational (R) hours among males and females aged 20–49 years at study entry

Physical activity (PhA)	Males		Total		Females Housewives		Non-housewives	
	Number	%	Number	%	Number	%	Number	%
Occupational PhA								
Sedentary (O1)	18 737	35.4	3 232	11.5	690	3.4	2 542	31.4
Walking (O2)	13 990	26.4	19 192	68.2	15 221	76.0	3 971	49.0
Lifting and walking (O3)	11 804	22.3	4 462	15.9	3 049	15.2	1 413	17.4
Heavy manual (O4)	8 414	15.9	1 237	4.4	1 065	5.3	172	2.1
Recreational PhA								
Sedentary (R1)	10 640	20.0	6 336	22.4	4 625	23.0	1 711	21.1
Moderately active (R2)	29 040	54.6	19 100	67.6	13 453	66.8	5 647	69.6
Regular training (R3)	12 206	22.9	2 757	9.8	2 033	10.1	724	8.9
Regular hard training (R4)	1 316	2.5	62	0.2	31	0.2	31	0.4

Table II Age-adjusted relative risk (RR) of colorectal cancer with 95% confidence interval (CI) in relation to possible risk factors. Cox's proportional hazards model

Variable	Colon cancer			Rectal cancer		
	No. of cases	RR ^a	95% CI	No. of cases	RR ^a	95% CI
Age at entry (year)	236	1.13	(1.10-1.16)	170	1.11	(1.07-1.15)
BMI (1 g cm ⁻³)	230	1.25	(1.01-1.55)	169	0.99	(0.87-1.12)
Cholesterol (1 mmol l ⁻¹)	236	1.02	(0.93-1.12)	170	1.00	(0.89-1.12)
Triglycerides (1 mmol l ⁻¹)	236	1.03	(0.96-1.10)	170	1.01	(0.92-1.11)
Height (10 cm)	224	1.13	(0.94-1.35)	169	1.02	(0.82-1.25)
Smoking (10 cigarettes)	229	0.96	(0.80-1.16)	164	1.06	(0.86-1.30)
Married/separated (ever vs never)	236	1.24	(0.80-1.91)	170	2.83	(1.39-5.78)

^a Adjusted for age at entry. Number of cases varied slightly due to missing values for some variables.

Table III Adjusted relative risk (RR) of colorectal cancer with 95% confidence interval (CI) according to total physical activity (occupational (O) and recreational (R) combined) among males and females; Cox's proportional hazards model

Total Physical activity	Total colon cancer			Proximal colon cancer			Distal colon cancer			Rectal cancer		
	No. of cases	RR ^a	95% CI	Trend test P value	No. of cases	RR ^a	95% CI	Trend test P value	No. of cases	RR ^a	95% CI	Trend test P value
Males												
Sedentary ^b	26	1.00			10	1.00			14	1.00		
Moderate ^c	95	1.18	(0.76-1.82)	0.49	36	1.16	(0.57-2.34)	0.64	56	1.29	(0.72-2.33)	0.53
Active ^d	107	0.97	(0.63-1.50)		48	0.96	(0.47-1.93)		57	0.99	(0.55-1.80)	
Females												
Sedentary ^b	22	1.00			10	1.00			11	1.00		
Moderate ^c	17	0.97	(0.33-2.77)	0.04	10	1.22	(0.51-2.94)	0.10	7	0.84	(0.32-2.17)	0.96
Active ^d	59	0.63	(0.39-1.04)		27	0.62	(0.30-1.28)		27	0.61	(0.30-1.23)	0.15

^a Adjusted for age at entry, geographic region and body mass index (BMI). ^b Sedentary (R1+O1-2). ^c Moderate (R1+O3-4, O1+R3-4). ^d Active (O2-4+R2-4).

When occupational activity was examined separately, each sex showed a consistent negative adjusted reduced risk for colon cancer in the active groups compared with the sedentary group, but in neither case was this significant (Table V). Recreational activity showed no consistent reduced trend in either sex, but females with moderate recreational physical activity (R2) had an almost 40% significant reduction in the risk of total colon cancer (RR=0.62, 95% CI 0.40–0.97). No consistent associations were observed between total physical activity, occupational or recreational physical activity and risk of rectal cancer in males or females (Table III and V).

When taking subsite into consideration, we performed site-specific analyses of the relationship between total physical activity and proximal and distal colon cancer. A negative trend for both proximal (P for trend=0.10) and distal cancers (P for trend=0.15) was observed in females, though this was not significant (Table III). A negative trend was observed only for proximal cancers in males older than 45 years at entry (P for trend=0.08) (results not presented in Table). Further the reduction of colon cancer risk among the recreational physically active females was particularly marked in the proximal colon (RR=0.51, 95% CI 0.28–0.93) (Table VI). No corresponding subsite differences were observed in males when taking only recreational activity into consideration.

Further, we examined models stratified by BMI (median split) to analyse if there was any effect modification related to body weight (Table VI). Among females an inverse recreational physical activity–colon cancer association was

stronger among leaner females (RR = 0.45, 95% CI 0.25–0.82) compared with more obese females. Among males an inverse physical activity–colon cancer association was strongest in older and leaner males. This was observed especially by dividing BMI into tertiles, as occupationally active males 45 years or older belonging to the lowest tertile (BMI < 2.33 g cm⁻²) had the greatest reduction in total colon cancer risk (RR=0.50, 95% CI 0.26–0.97) (results not shown in Table VI).

To examine if the effect of physical activity on colorectal cancer differed between males and females we performed combined analyses both for total colon and for subsites. Here, we observed no significant effect of gender alone or when introducing an interaction term of gender and physical activity on colorectal cancer risk in any of the analyses (results not shown).

Discussion

In the present study an inverse dose–response relationship between total physical activity and colon cancer risk was observed in females. In males this inverse dose–response relationship was found only for those 45 years or older at study entry. An almost 40% reduction in risk of colon cancer among the moderately leisure time active compared with sedentary females was demonstrated. This reduction in cancer risk in females from recreational physical activity was particularly related to proximal colon with an almost 50%

Table IV Adjusted relative risk (RR) of colon cancer with 95% confidence interval (CI) according to total physical activity (occupational (O) and recreational (R) combined) stratified by age at entry among males and females; Cox's proportional hazards model

Total Physical activity	No. of cases	Males			Trends test P value	No. of cases	Females		Trend test P value
		RR ^a	95% CI	95% CI			RR ^a	95% CI	
<45 years at entry									
Sedentary ^b	5	1.00				11	1.00		
Moderate ^c	30	2.02	(0.78–5.21)			8	0.96	(0.39–2.40)	
Active ^d	49	2.23	(0.88–5.66)	0.13		30	0.62	(0.31–1.23)	0.13
≥45 years at entry									
Sedentary ^b	21	1.00				11	1.00		
Moderate ^c	65	0.96	(0.59–1.58)			9	0.99	(0.41–2.39)	
Active ^d	58	0.66	(0.40–1.10)	0.04		29	0.66	(0.33–1.33)	0.19

^a Adjusted for age at entry, geographic region and body mass index (BMI). ^b Sedentary (R1+O1–2). ^c Moderate (R1+O3–4, O1+R3–4). ^d Active (O2–4+R2–4).

Table V Adjusted relative risk (RR) of colorectal cancer with 95% confidence interval (CI) related to categories of occupational (O) and recreational (R) physical activity among males and females; Cox's proportional hazards model

Physical activity (PhA)	No. of cases	Colon cancer			Trend test P value	No. of cases	Rectal cancer		Trend test P value
		RR ^a	95% CI	95% CI			RR ^b	95% CI	
Males									
Occupational PhA									
Sedentary (O1)	92	1.00				71	1.00		
Walking (O2)	62	0.92	(0.67–1.28)			43	0.90	(0.61–1.31)	
Lifting/Heavy manual (O3+O4)	74	0.82	(0.59–1.13)	0.22		54	1.00	(0.69–1.45)	0.95
Recreational PhA									
Sedentary (R1)	41	1.00				29	1.00		
Moderately active (R2)	125	1.05	(0.74–1.50)			106	1.25	(0.83–1.89)	
Regular training (R3+R4)	64	1.33	(0.90–1.98)	0.13		34	0.98	(0.60–1.61)	0.85
Females									
Occupational PhA									
Sedentary (O1)	12	1.00				6	1.00		
Walking (O2)	66	0.82	(0.44–1.51)			37	0.95	(0.40–2.26)	
Lifting/Heavy manual (O3+O4)	20	0.69	(0.34–1.42)	0.32		12	0.88	(0.33–2.36)	0.78
Recreational PhA									
Sedentary (R1)	30	1.00				9	1.00		
Moderately active (R2)	57	0.62	(0.40–0.97)			40	1.51	(0.73–3.11)	
Regular training (R3+R4)	12	0.84	(0.43–1.65)	0.25		6	1.49	(0.53–4.22)	0.35

^a Adjusted for age at entry, geographic region and body mass index (BMI). ^b Adjusted for age at entry, geographic region, body mass index (BMI) and civil status.

Table VI Adjusted relative risk^a of colon cancer with 95% confidence intervals (in parentheses) related to occupational (O) and recreational (R) physical activity stratified by subsites and body mass index (BMI) in males and females; Cox's proportional hazards model

	No. of cases	Occupational physical activity		No. of cases	Recreational physical activity		
		Sedentary (O1)	Active (O2-4)		Sedentary (R1)	Active (R2-4)	
Males							
Subsites							
Proximal	89	1.00	0.89 (0.57-1.18)	90	1.00	1.05 (0.62-1.78)	
Distal	127	1.00	0.82 (0.57-1.18)	128	1.00	1.19 (0.75-1.89)	
BMI (g cm ⁻²)							
< 2.44	89	1.00	0.87 (0.56-1.35)	89	1.00	1.36 (0.74-2.51)	
≥ 2.44	139	1.00	0.85 (0.60-1.21)	141	1.00	1.05 (0.69-1.58)	
Females							
Subsites							
Proximal	47	1.00	1.14 (0.41-3.18)	48	1.00	0.51 (0.28-0.93)	
Distal	45	1.00	0.52 (0.24-1.11)	45	1.00	0.80 (0.41-1.56)	
BMI (g cm ⁻²)							
< 2.36	48	1.00	1.43 (0.51-3.98)	48	1.00	0.45 (0.25-0.82)	
≥ 2.36	50	1.00	0.50 (0.23-1.06)	51	1.00	0.93 (0.49-1.74)	

^a Adjusted for age at entry, geographic region and body mass index (BMI).

reduction among active females. No association between physical activity and rectal cancer was observed in males or females.

The strength of this study beyond its prospective design, large size, broad population base and inclusion of both sexes, is a nearly complete cancer case ascertainment. Compulsory reporting by hospital departments and pathological laboratories for all new cases of cancer in Norway as well as death certificates results in very high case ascertainment. This is in addition to an almost 100% histological verification of colon cancer cases.

The accuracy of the self-reported physical activity questions used in the present analysis has been validated in several studies (Wilhelmsen *et al.*, 1976; Bjartveit *et al.*, 1981; Holme *et al.*, 1981; Lochen and Rasmussen 1992). Lochen and Rasmussen (1992) demonstrated that physical fitness among males increased with physical activity in leisure time. However, there are some limitations in using a single brief questionnaire reporting physical activity during one year without repeated assessments of physical activity and measurements of energy expenditure or dietary information. The large proportion (70%) of housewives in our cohort may have limited our ability to detect any effect of occupational activity on colon cancer risk among females. A greater variability in physical activity during leisure time rather than at work may in part explain why leisure time activity in females significantly reduced risk of colon cancer and occupational activity did not. In addition, the participants had to choose between only four occupational categories and four recreational levels of physical activity and we may therefore have underestimated the strength of physical activity for those most active.

The present findings support and extend previous results showing that physical activity is inversely related to colon cancer risk in humans (Garabrant *et al.*, 1984; Gerhardsson *et al.*, 1986; Brownson *et al.*, 1989; Peters *et al.*, 1989; Arbmán *et al.*, 1993; Chow *et al.*, 1993; Fraser and Pearce, 1993; Wu *et al.*, 1987; Slattery *et al.*, 1988; Gerhardsson *et al.*, 1988; Severson *et al.*, 1989; Ballard-Barbash *et al.*, 1990; Lee *et al.*, 1991; Markowitz *et al.*, 1992; Giovannucci *et al.*, 1995) and animals (Andrianopoulos *et al.*, 1987; Reddy *et al.*, 1988).

We did not find an overall protective effect of total physical activity on colon cancer in males. This may be owing to the young age at entry and the fact that the number of cases of colon cancer are relatively small among the youngest males, thereby limiting the statistical power. The observation that only males 45 years or older at study entry had a protective effect of physical activity on colon cancer risk is consistent with similar findings in previous studies which support that age may be an effect modifier for colon cancer (Albanes *et al.*, 1989; Ballard-Barbash *et al.*,

1990; Slattery *et al.*, 1994). The observed 50% reduction in colon cancer risk among occupationally active, older and leaner males compared with sedentary males is in agreement with findings in the Framingham study in which the strongest inverse physical activity-large bowel cancer association was found among older (> 50 years) and leaner males (Ballard-Barbash *et al.*, 1990). In contrast, no such age effect was found among females in the present nor in the Framingham study. An interpretation may be a somewhat different age distribution at diagnosis in females relative to males (median age at diagnosis; males, 58.1 years; females 54.6 years). Power may also be greater for males owing to the much greater number of cancer cases compared with females in both studies, thus making any age effect easier to discover in males. Consequently, physical activity as a protective factor in colon cancer risk may be of greater importance among the elderly relative to younger subjects in whom the importance of genetic predisposition may be greater. Biological mechanisms related to an age effect from physical activity on colon cancer risk have been proposed to act through improvements of the immune system among physically active elderly subjects (Shepard and Shek, 1995) or that physical activity, acting over a longer period of time in older people, is particularly important (Lee *et al.*, 1991). In spite of no significant gender differences from physical activity on colon cancer risk observed in the present study, previous studies suggest sex differences as men and women show differences under controlled experimental conditions in gastrointestinal transit time, stool bulk and bile acid production (Stephen *et al.*, 1986; Lampe *et al.*, 1993).

The inverse association between physical activity and colon cancer risk observed in the present study could be confounded. Physically active individuals may have had a diet with less saturated fat and more fibre than the inactive ones. Unfortunately, no dietary data were available for this analysis. However, other studies that examined dietary differences have concluded that physical activity and dietary factors are independent risk factors for colon cancer (Slattery *et al.*, 1988; Peters *et al.*, 1989; Gerhardsson de Verrier *et al.*, 1990; Whittemore *et al.*, 1990; Giovannucci *et al.*, 1994).

Holme *et al.* (1981), partly examining the same male cohort as followed in the present study, observed that higher social classes dominated among males who reported sedentary work, while males who were sedentary at leisure time more often represent lower social classes. Therefore, the reference group used in total physical activity reflect both high and low social classes. Social class did not influence colon cancer risk in a comparable society (Suadicani *et al.*, 1993). It is less likely, therefore, that social class explains the observed association between total physical activity and colon cancer risk.

Another observation of interest was the protective effect

on proximal colon cancer incidence among physically leisure-time active females. This observation was in part also demonstrated in males 45 years or older at study entry with a particular reduction in proximal colon cancer incidence as a result of total physical activity. A possible explanation could be that exercise affects gut mobility more extensively in the proximal relative to distal colon. However, previous studies which have taken site-specific colon cancer risk into consideration have been inconsistent (Fraser and Pearce, 1993; Peters *et al.*, 1989; Gerhardtsson de Verdier *et al.*, 1990; Vena *et al.*, 1985; Gerhardtsson *et al.*, 1986; Brownson *et al.*, 1989).

Our study suggests that body size may modify the effect of physical activity as we observed leaner active males and females to be at a decreased colon cancer risk compared with obese subjects. This agrees with previous reports (Albanes *et al.*, 1989; Ballard-Barbash *et al.*, 1990; Giovannucci *et al.*, 1995). Body mass index (BMI) as a significant risk factor for colon cancer among males is consistent with previous studies (Marchand *et al.*, 1992; Ballard-Barbash *et al.*, 1990). The results have been more inconsistent in females (Albanes *et al.*, 1989; Ballard-Barbash *et al.*, 1990; Whittemore *et al.*, 1990). However, no direct mechanism has been suggested for the colon cancer-obesity association, but the association may indirectly be an effect of both diet and physical activity.

Several potential biological mechanisms may contribute to an observed protective effect of physical activity on colon cancer risk, including constipation which is often improved by physical activity. Walking (Holdstock *et al.*, 1970), running (Cordain *et al.*, 1986) and strength training (Koffler *et al.*, 1992), have generally been found to reduce GI transit times although not in one study (Bingham and Cummings, 1989). Contact between the colon mucosa and potential carcinogens in the faecal stream may be decreased by exercise because of shortened transit time. The fact that physical

activity does not seem to lower the risk of rectal cancer accords with this 'transit time theory' as the rectum is only intermittently filled with faeces and colon peristalsis has less influence on the faecal transit time in the rectum. A decrease in the ratio of secondary to primary bile acids has been observed in obese patients after treatment with subcaloric diet and graded physical activity (Kadyrova and Shakieva, 1986). This effect of physical activity may be of importance since a high excretion of bile acids may increase the risk of colon cancer. Exercise can also elevate the production of some prostaglandins that, in turn, may influence colon cancer risk (Demers *et al.*, 1981). Physical activity may also increase colonic bloodflow so that faecal mutagens are transported away from the mucous membrane.

In conclusion, our study supports a protective effect of total physical activity on colon cancer, but not rectal cancer, in both males and females. In males this protective effect of physical activity is of greatest importance among the elderly. The stronger protective effect of physical activity on proximal rather than distal colon cancer risk supports the assumption that physical activity affects gut mobility more extensively in the proximal relative to distal colon. Further studies are needed in which repeated measurements of duration and intensity of physical activity besides energy balance, dietary factors, age, subsites and gender differences are taken into account.

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References

- ALBANES D, BLAIR AA AND TAYLOR PR. (1989). Physical activity and risk of cancer in the NHANES I population. *Am. J. Public Health*, **79**, 744-750.
- ANDRIANPOPULOS G, NELSON RL, BOMBECK CT AND SOUZA G. (1987). The influence of physical activity in 1,2-dimethylhydrazine induced colon carcinogenesis in the rat. *Anticancer Res.*, **7**, 849-852.
- ARBMAN G, AXELSON O, FREDRIKSSON M, NILSSON E AND SJÖDAHL R. (1993). Do occupational factors influence the risk of colon and rectal cancer in different ways? *Cancer*, **72**, 2543-2549.
- BALLARD-BARBASH R, SCHATZKIN A, ALBANES D, SCHIFFMAN MH, KREGER BE, KANNEL WB, ANDERSON KM AND HELSEL WE. (1990). Physical activity and risk of large bowel cancer in the Framingham Study. *Cancer Res.*, **50**, 3610-3613.
- BINGHAM SA AND CUMMINGS JH. (1989). Effect of exercise and physical fitness on large intestinal function. *Gastroenterology*, **97**, 1389-1399.
- BJARTVEIT K, FOSS OP AND GJERVIG T. (1981). The cardiovascular disease study in Norwegian counties. Results from first screening. *Acta Med. Scand.*, **209**, 277-283.
- BROWNSON RC, ZAHM SH, CHANG JC AND BLAIR AA. (1989). Occupational risk of colon cancer. An analysis by anatomic subsite. *Am. J. Epidemiol.*, **130**, 675-687.
- BUFILL JA. (1990). Colorectal cancer: Evidence for distinct genetic categories based on proximal or distal tumor location. *Ann. Int. Med.*, **113**, 779-788.
- CHOW WH, DOSEMECI M, ZHENG W, VETTER R, MCLAUGHLIN JK, GAO YT AND BLOT WJ. (1993). Physical activity and occupational risk of colon cancer in Shanghai, China. *Int. J. Epidemiol.*, **22**, 23-29.
- CORDAIN L, LATIN RW AND BEHNKE JJ. (1986). The effects of an aerobic running program on bowel transit time. *J. Sports Med.*, **26**, 101-104.
- DEMERS LM, HARRISON TS, HALBERT DR AND SANTEN RJ. (1981). Effect of prolonged exercise on plasma prostaglandin levels. *Prostaglandins Med.*, **6**, 413-418.
- DUBROW R, BERNSTEIN J AND HOLFORD TR. (1993). Age-period-cohort modelling of large-bowel-cancer incidence by anatomic sub-site and sex in Connecticut. *Int. J. Cancer*, **53**, 907-913.
- ENGELAND A, HALDORSEN T, TRETLI S, HAKULINEN T, HÖRTE LG, LUOSTARINEN T, MAGNUS K, SCHAU G, SIGVALDASON H, STORM HH, TULENIUS H AND VAITTINEN P. (1993). Prediction of cancer incidence in the Nordic countries up to the years 2000 and 2010. *APMIS*, **38**(suppl.), 101.
- FLESHNER P, SLATER G AND AUFSES AH. (1989). Age and sex distribution of patients with colorectal cancer. *Dis. Colon Rectum*, **32**, 107-111.
- FRASER G AND PEARCE N. (1993). Occupational physical activity and risk of cancer of the colon and the rectum in New Zealand males. *Cancer Causes Control*, **4**, 45-50.
- GARABRANT DH, PETERS JM, MACK TM AND BERNSTEIN L. (1984). Job activity and colon cancer risk. *Am. J. Epidemiol.*, **119**, 1005-1014.
- GERHARDSSON M, NORELL SE, KIVIRANTA H, PEDERSEN NL AND AHLBOM A. (1986). Sedentary jobs and colon cancer. *Am. J. Epidemiol.*, **123**, 775-780.
- GERHARDSSON M, FLODERUS B AND NORELL SE. (1988). Physical activity and colon cancer risk. *Int. J. Epidemiol.*, **17**, 743-746.
- GERHARDSSON DE VERDIER M, STEINECK G, HAGMAN U, RIEGER Å AND NORELL SE. (1990). Physical activity and colon cancer: A case-referent study in Stockholm. *Int. J. Cancer*, **46**, 985-989.
- GIOVANNUCCI E, RIMM EB, STAMPFER MJ, COLDITZ GA, ASCHERIO A AND WILLETT WC. (1994). Intake of fat, meat, and fibre in relation to risk of colon cancer in men. *Cancer Res.*, **54**, 2390-2397.
- GIOVANNUCCI E, ASCHERIO A, RIMM EB, COLDITZ GA, STAMPFER MJ AND WILLETT WC. (1995). Physical activity, obesity, and risk for colon cancer and adenoma in men. *Ann. Int. Med.*, **122**, 327-334.
- HALVORSEN TB. (1986). Site distribution of colorectal adenocarcinomas. A retrospective study of 853 tumours. *Scand. J. Gastroenterol.*, **21**, 973-978.
- HOLDSTOCK DJ, MISIEWICZ JJ, SMITH T AND ROWLANDS EN. (1970). Propulsion (mass movements) in the human colon and its relationship to meals and somatic activity. *Gut.*, **11**, 91-99.

- HOLME I, HELGELAND A, HJERMANN I, LEREN P AND LUND-LARSEN PG. (1981). Physical activity at work and at leisure in relation to coronary risk factors and social class. A 4-year mortality follow-up. The Oslo Study. *Acta Med. Scand.*, **209**, 277-283.
- KADYROVA RK AND SHAKIEVA RA. (1986). Dynamics of changes in the lipid composition of bile in patients with alimentary obesity during treatment. *Ter. Arkh. (Moscow)*, **58**, 79-82.
- KOFFLER KH, MENKES A, REDMOND RA, WHITEHEAD WE, PRATLEY RE AND HURLEY BF. (1992). Strength training accelerates gastrointestinal transit in middle-aged and older men. *Med. Sci. Sports and Exercise*, **24**, 415-419.
- LAMPE JW, FREDSTROM SB, SLAVIN JL AND POTTER JD. (1993). Sex differences in colonic function: a randomized trial. *Gut*, **34**, 531-536.
- LEE I-MIN, PAFFENBARGER JR. RS AND HSIEH CC. (1991). Physical activity and risk of developing colorectal cancer among college alumni. *J. Natl Cancer Inst.*, **83**, 1324-1329.
- LØCHEN M-L AND RASMUSSEN K. (1992). The Tromsø study: physical fitness, self reported physical activity, and their relationship to other coronary risk factors. *J. Epidemiol. Commun. Health*, **26**, 103-107.
- MARCHAND LL, WILKENS LR AND MI M-P. (1992). Obesity in youth and middle age and risk of colorectal cancer in men. *Cancer Causes Control*, **3**, 349-354.
- MARKOWITZ S, MORABIA A, GARIBALDI K AND WYNDER E. (1992). Effect of occupational and recreational activity on the risk of colorectal cancer among males: a case-control study. *Int. J. Epidemiol.*, **21**, 1057-1062.
- MØLLER JENSEN OM. (1984). Different age and sex relationship for cancer of subsites of the large bowel. *Br. J. Cancer*, **50**, 825-829.
- MUIR C, WATERHOUSE J, MACK T, POWELL J AND WHELAN S. (1987). *Cancer Incidence in Five Continents*. Vol. V. IARC Scientific publication. 88. IARC: Lyon.
- PETERS RK, GARABRANT DH, YU MC AND MACK TM. (1989). A case-control study of occupational and dietary factors in colorectal cancer in young men by subsite. *Cancer Res.*, **49**, 5459-5468.
- REDDY BS, SUGIE S AND LOWENFELS A. (1988). Effect of voluntary exercise on azoxymethane-induced colon carcinogenesis in male F344 rats. *Cancer Res.*, **48**, 7079-7081.
- SAS INSTITUTE. (1992). *SAS/STAT Guide for Personal Computers*. Version 6 edition. SAS Institute: Cary, NC (USA).
- SEVERSON RK, NOMURA AMY, GROVE JS AND STEMMERMANN GN (1989). A prospective analysis of physical activity and cancer. *Am. J. Epidemiol.*, **130**, 522-529.
- SHEPHARD RJ AND SHEK PN. (1995). Exercise, aging and immune function. *Int. J. Sports Med.*, **16**, 1-6.
- SLATTERY ML, SCHUMACHER MC, SMITH KR, WEST DW AND ABD-ELGHANY N. (1988). Physical activity, diet, and risk of colon cancer in Utah. *Am. J. Epidemiol.*, **128**, 989-999.
- SLATTERY ML, ABD-ELGHANY N, KERBER R AND SCHUMACHER MC. (1990). Physical activity and colon cancer: a comparison of various indicators of physical activity to evaluate the association. *Epidemiology*, **1**, 481-485.
- SLATTERY ML, POTTER JD AND SORENSON AW. (1994). Age and risk factors for colon cancer (United States and Australia): are there implications for understanding differences in case control and cohort studies? *Cancer Causes Control*, **5**, 557-563.
- STEPHEN AM, WIGGINS HS, ENGLYST HN, COLE TJ, WAYMAN BJ AND CUMMINGS JH. (1986). The effect of age, sex, and level of intake of dietary fibre from wheat on large-bowel function in thirty healthy subjects. *Br. J. Nutr.*, **56**, 349-361.
- SUADICANI P, HEIN HO AND GYNTELBERG F. (1993). Height, weight, and risk of colorectal cancer. (1993). *Scand. J. Gastroenterol.*, **28**, 285-288.
- VENA JE, GRAHAM S, ZIELEZNY M, SWANSON MK, BARNES RE AND NOLAN J. (1985). Lifetime occupational exercise and colon cancer. *Am. J. Epidemiol.*, **122**, 357-365.
- WHITTEMORE AS, WU-WILLIAMS AH, LEE M, SHU Z, GALLAGHER RP, DENG-AO J, LUN Z, XIANGHUI W, KUN C, JUNG D, TEH C-Z, CHENGDE L, YAO XJ, PAFFENBARGER JR. RS AND HENDERSON BE. (1990). Diet, physical activity, and colorectal cancer among chinese in North America and China. *J. Natl. Cancer Inst.*, **82**, 915-926.
- WILHELMSEN L, TIBBLIN G, AURELL M, BJURE J, EKSTRØM-JODAL B AND GRIMBY G. (1976). Physical activity, physical fitness and risk of myocardial infarction. *Adv. Cardiol.*, **18**, 217-230.
- WU AH, PAGANINI-HILL A, ROSS RK AND HENDERSON BE. (1987). Alcohol, physical activity and other risk factors for colorectal cancer: a prospective study. *Br. J. Cancer.*, **55**, 687-694.

Paper III

THE INFLUENCE OF PHYSICAL ACTIVITY ON LUNG-CANCER RISK A prospective study of 81,516 men and women

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Physical activity is inversely related to mortality from respiratory diseases including lung cancer. Physical activity improves pulmonary function but its impact on lung-cancer risk has not been studied much. During 1972–1978, 53,242 men and 28,274 women, aged 20 to 49 years, participated in a population-based health survey and were followed until 31 December 1991. We observed a total of 413 men and 51 women with lung cancer. Leisure activity and work activity were assessed using a questionnaire in 4 categories. In a sub-cohort, physical activity was assessed twice at an interval of 3 to 5 years. Leisure but not work activity was inversely related to lung-cancer risk in men after adjustment for age, smoking habits, body-mass index and geographical residence (p for trend = 0.01). Men who exercised at least 4 hours a week had a lower risk than men who did not exercise [relative risk (RR) = 0.71; 95% confidence interval (CI) = 0.52–0.97]. Reduced risk of lung cancer was particularly marked for small-cell carcinoma (RR = 0.59; 95% CI = 0.38–0.94) and for adenocarcinoma (RR = 0.65; 95% CI = 0.41–1.05), with no association seen for squamous-cell carcinoma. In the sub-cohort in which physical activity was assessed twice, the risk of lung cancer was particularly reduced among men who were most active at both assessments (RR = 0.39; 95% CI = 0.18–0.85). No consistent association between physical activity and lung-cancer risk was observed among women. Our results suggest that leisure physical activity has a protective effect on lung-cancer risk in men. The small number of incident cases, combined with the narrow range of physical activity reported, may have limited our ability to detect an association between physical activity and lung cancer in women. *Int. J. Cancer*, 70:57–62, 1997.

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Pulmonary function is inversely related to mortality from respiratory diseases including lung cancer (Kuller *et al.*, 1990; Nomura *et al.*, 1991; Weiss *et al.*, 1995). A measure of pulmonary function, the forced expiratory volume in one second adjusted for height (FEV₁/height), correlates positively with strenuous physical activity and duration of exercise (Kuller *et al.*, 1990; Higgins *et al.*, 1991). In a prospective study, Nomura *et al.* (1991) observed a reduction in lung-cancer risk among subjects with high levels of FEV₁. Therefore physical activity may have an influence on subsequent lung-cancer risk. No study has specifically focused on this relationship, although a few epidemiological studies have provided indications that physical activity may reduce lung-cancer risk (Albanes *et al.*, 1989; Severson *et al.*, 1989; Sellers *et al.*, 1991; Lee and Paffenbarger, 1994).

Smoking is a well-established cause of lung cancer. The fact that most smokers never develop lung cancer has prompted interest in the role of host factors in pulmonary carcinogenesis (McDuffie *et al.*, 1993). Prospective and case-control studies have shown that histological types of lung cancer vary in their respective aetiology (Vena *et al.*, 1985; Yang *et al.*, 1989; McDuffie *et al.*, 1993). Smoking has been shown to be a strong risk factor for squamous-cell carcinoma and small-cell carcinoma, but a weak risk factor for adenocarcinoma (Brownson *et al.*, 1992; McDuffie *et al.*, 1993). Adenocarcinoma is the second most frequent histological sub-type of cancer in Norway (Cancer Registry of Norway, 1995). A recent study in the USA, however, has reported that adenocarcinoma has replaced squamous-cell carcinoma as the most common histological sub-type of lung cancer for all sexes and races combined (Travis *et al.*, 1995). This may indicate a change in the aetiology and pathogenesis of this very important cancer type. To verify this,

possible factors other than the effect of smoking on lung-cancer risk must be examined. We have reported earlier, from the same cohort, a protective effect of physical activity on prostate (Thune and Lund, 1994) and colon cancer (Thune and Lund, 1996), a finding that points to physical activity as a protective factor for certain cancer sites.

In this paper we hypothesize that physical activity may lower the risk of lung cancer. This hypothesis is based on the analysis of a large population-based prospective study of 81,516 men and women, in which careful adjustments for smoking habits were made. In a sub-cohort, 2 assessments of physical activity permitted the evaluation of sustained physical activity on the risk of lung cancer.

SUBJECTS AND METHODS

Cohort

Between 1972 and 1978, 104,485 men and women, aged 20 to 49 years, from 3 counties (Oppland, Sogn and Fjordane, and Finnmark) and 2 cities (Oslo and Tromsø) were invited to participate in a population-based health survey of risk factors for cardiovascular disease. In Tromsø, all men aged 20 to 49 years were invited to participate, whereas in Oslo men aged 40 to 49 were invited, together with a 7% random sample of men aged 20 to 39. In the 3 counties of Oppland, Sogn and Fjordane, and Finnmark, all men and women aged 35 to 49 and a 10% random sample of people aged 20 to 34 years were invited to participate. In 4 small municipalities in Finnmark, all men and women aged 20 to 34 were invited: a total of 104,485, of whom 53,622 men (73.5%) and 28,621 women (90.7%) attended the screening. In the 3 counties, 3 or 5 years later (1977–1983), people still resident there were invited to a similar health survey.

Screening procedures were almost identical in the 5 areas. Each person was initially contacted by mail, with a covering letter and a one-page questionnaire. The participants were asked to answer the questionnaire at home and to bring it to the clinical examination at which the questionnaire was checked for inconsistencies. Measurements of weight, height and blood pressure were made, and blood samples were taken at the examination. The questionnaire covered the following areas: physical activity (PhA) during recreational (R) and occupational (O) hours within the last year; history of chronic diseases, especially cardiovascular symptoms and diseases; smoking habits and stress in daily life.

Population for analysis

All 53,622 men and 28,621 women were followed up through the Norwegian Central Bureau of Statistics to identify deaths in the cohort until the end of 1991. Those who emigrated, or who had a pre-existing malignancy, or who developed a malignancy within the first year after the survey (men, $n = 380$; women, $n = 347$) were excluded from the analyses. This reduced the possibility of

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TABLE 1 - AGE-ADJUSTED¹ MEAN/DISTRIBUTION (%) OF BASELINE CHARACTERISTICS ACCORDING TO LEVEL AND CATEGORY OF PHYSICAL ACTIVITY AT FIRST SCREENING, BY GENDER

Characteristics by gender	Recreational activity			Occupational activity			
	Sedentary ²	Moderately active	Regular exercise	Sedentary	Walking	Lifting	Heavy manual
Men	(n = 10,640)	(n = 29,040)	(n = 13,522)	(n = 18,737)	(n = 13,990)	(n = 11,804)	(n = 8,414)
Age at entry (years)	42.7	43.0	41.3	42.8	42.8	41.9	42.3
BMI (kg/m ²)	25.1	24.6	24.5	24.6	24.6	24.8	25.0
Cholesterol (mmol/l)	6.92	6.83	6.61	6.72	6.77	6.87	6.86
Triglycerides (mmol/l)	2.60	2.47	2.36	2.39	2.47	2.54	2.54
Smoking habits							
Current cigarette smokers (%)	59.0	49.7	39.3	43.1	49.1	56.6	50.6
Number of cigarettes (daily)	15.0	13.4	12.2	14.2	14.2	13.3	12.8
Number of years smoked	21.0	20.3	19.8	20.1	20.3	20.7	20.7
Pipe or cigar smokers (%)	6.0	6.3	6.4	7.0	5.8	5.6	6.2
Ex-smokers (%)	18.1	22.7	24.7	25.0	22.7	19.9	18.8
Never-smokers (%)	16.7	21.0	29.2	24.5	22.2	17.4	23.9
Women	(n = 6,336)	(n = 19,100)	(n = 2,819)	(n = 3,232)	(n = 19,192)	(n = 4,462)	(n = 1,237)
Age at entry (years)	41.5	41.1	41.2	40.2	40.9	41.9	43.3
BMI (kg/m ²)	24.9	24.4	24.1	24.1	24.4	24.8	25.3
Cholesterol (mmol/liter)	6.76	6.65	6.52	6.63	6.68	6.66	6.61
Triglycerides (mmol/liter)	1.91	1.81	1.76	1.80	1.83	1.85	1.81
Smoking habits							
Current cigarette smokers (%)	42.7	38.0	33.0	41.0	39.1	39.1	22.6
Number of cigarettes (daily)	10.2	9.1	8.8	10.1	9.3	9.2	8.4
Number of years smoked	16.0	15.6	15.5	16.1	15.7	15.3	14.6
Ex-smokers (%)	11.0	13.1	14.5	14.7	13.1	11.5	8.4
Never-smokers (%)	46.1	48.7	52.2	44.1	47.6	49.2	68.7

¹Except mean age.--²Number of participants for activity categories in parentheses. For some subjects, information concerning smoking status, height and body mass index (BMI) was missing.

any undiagnosed cancer influencing the level of physical activity. Included for analysis were 53,242 men (867,822 person-years) and 28,274 women (437,785 person-years).

A sub-cohort of 25,879 men and 26,131 women participating in the first (1974-1978) and the second (1977-1983) screening were followed for an additional year, until the end of 1992 (attendance rate was 85.3% of all invited). Those who emigrated, who had a pre-existing malignancy or who developed a malignancy within the first year after attending the second survey (men, n = 270; women, n = 772) were excluded from analyses. Included for analysis in the sub-cohort were 25,609 men (306,488 person-years) and 25,629 women (309,303 person-years).

Assessment of physical activity

Self-reported physical-activity categories during recreational hours in the last year was graded from 1 to 4, according to which of the following categories provided the best description of the participant's usual level of leisure activity: R1, reading, watching TV or other sedentary activities; R2, walking, bicycling or physical activities for at least 4 hr a week; R3, exercise to keep fit, participating in recreational athletics, etc., for at least 4 hr a week; R4, regular hard training or participation in competitive sports several times a week.

Self-reported physical activity during work hours in the last year was divided into 4 categories: O1, mostly sedentary work; O2, work with a lot of walking; O3, work with a lot of lifting and walking; O4, heavy manual work. At the screening, trained nurses checked the reporting of physical activity at work and leisure time for inconsistencies, and particular attention was given to housewives. This has been described in more detail elsewhere (Thune and Lund, 1996). Heart rate and other measures of physical fitness were not assessed.

Identification of cases

The national 11-digit personal identification number enabled a linkage to the Cancer Registry of Norway. This allowed identification of every incident case of lung cancer occurring in the cohort in accordance with the seventh revision of the International Classification of Diseases. Histological classification was based on the

diagnoses in the pathology reports. Reporting of malignant and pre-malignant diagnoses is mandatory for all laboratories in the country. Cases identified only incidentally *post mortem* were not included. Histological confirmation was performed in 95.4% of the cases among men and 98.0% among women.

Statistical analysis

All analyses were gender-specific. In the main cohort, observation years at risk were calculated as the number of years from one year after entry into the study until the time of withdrawal (year of diagnosis, time of death or end of follow-up at 31 December 1991). In the sub-cohort, we calculated the observation time for each person from one year after second screening until the time of withdrawal (year of diagnosis, death or end of follow-up at 31 December 1992). This reduced the possibility of any undiagnosed cancer influencing the level of physical activity reported at both surveys. Baseline variables were age-adjusted and compared by analysis of co-variance. Cox's proportional-hazard regression technique was used to analyze the simultaneous effect of physical activity and co-variables on lung-cancer incidence in the cohort.

In the analyses, the R3 and R4 categories of leisure activity were merged because of the small numbers in category R4 in both screenings. We adjusted for age at entry into the screening (continuous variable), smoking habits, geographical regions and obesity at time of measurements. Smoking habits were adjusted according to ex-smoking, pipe and cigar smoking, current cigarette smoking, including number of cigarettes smoked, and years of smoking. In addition, stratified analyses by smoking behaviour were performed. As a measure of obesity, we used the body-mass index (BMI) [weight (kg)/(height [m])²].

To study the influence of total physical activity on lung-cancer risk, occupational (O) and recreational (R) physical activity were combined. We used sedentary leisure activity (R1) and sedentary at work (O1) as the reference group (R1/O1). To account for changes over time in physical activity and smoking habits, we used information from both screenings.

The analyses were performed with the Proc Phreg procedure in the SAS statistical package. In some analyses, only cases with

TABLE II - ADJUSTED RELATIVE RISK (RR) OF LUNG CANCER WITH 95% CONFIDENCE INTERVAL (CI) RELATED TO CATEGORIES OF OCCUPATIONAL (O) AND RECREATIONAL (R) PHYSICAL ACTIVITY AT THE FIRST SCREENING (1972-78) AMONG MEN AND WOMEN

Physical activity (PhA)	Men			Women		
	Number of cases	RR ¹	95% CI	Number of cases	RR ¹	95% CI
Occupational PhA						
Sedentary (O1)	139	1.00	(Ref)	8	1.00	(Ref)
Walking (O2)	119	1.15	(0.90-1.47)	34	0.81	(0.37-1.76)
Lifting (O3)	97	1.13	(0.87-1.47)	8	0.79	(0.30-2.12)
Heavy manual (O4)	47	0.99	(0.70-1.41)	0	—	—
Trend test		<i>p</i> = 0.71			<i>p</i> = 0.30	
Recreational PhA						
Sedentary (R1)	123	1.00	(Ref)	14	1.00	(Ref)
Moderate (R2)	217	0.75	(0.60-0.94)	32	0.91	(0.48-1.71)
Regular training (R3 + R4)	62	0.71	(0.52-0.97)	5	0.99	(0.35-2.78)
Trend test		<i>p</i> = 0.01			<i>p</i> = 0.88	
Total PhA (occupational + recreational)						
Sedentary (O1 + R1)	52	1.0	(Ref)	2	1.0	(Ref)
Active	349	0.73	(0.54-0.98)	48	0.87	(0.21-3.62)

¹Adjusted for age at entry, geographical region, smoking habits [ex-smoking, pipe/cigar smoking (males only), number of cigarettes smoked, years smoked] and BMI.

histological diagnoses of squamous-cell carcinoma, adenocarcinoma and small-cell carcinoma were taken into consideration. As a result of missing data, the number of subjects included in the individual analyses varies slightly.

RESULTS

Lung cancer was diagnosed in 413 men and 51 women with median age at diagnosis of 57.3 (39.1-67.8) years and 54.2 (42.1-62.7) years in men and women, respectively, in the main cohort. Squamous-cell carcinoma was diagnosed in 128 cases (31.0%) and 10 cases (19.6%); adenocarcinoma in 88 cases (21.3%) and 17 cases (33.3%); small-cell carcinoma in 84 cases (20.3%) and 15 cases (29.4%); other types/unspecified malignancy in 94 cases (22.8%) and 8 cases (15.7%); histology not known 19 cases (4.6%) and 1 case (2.0%), in men and women, respectively.

Table I presents baseline characteristics according to type and level of activity. Approximately 25% of the men, but only 10% of the women, reported regular leisure exercise. Gender differences were also observed during occupational hours; two thirds of the women reported frequent walking, whereas in men work activity was more equally distributed among the different activity categories. Subjects reporting more leisure activity tended to be leaner and had lower serum lipids, in contrast to those subjects who had little leisure exercise and those performing heavy manual occupational activity, who had the highest BMI among both sexes. Current cigarette smokers dominated the group, with low leisure exercise, whereas ex-smokers and never-smokers reported more frequent leisure exercise.

After adjustment for age at entry, geographical region, smoking habits (ex-smoker, pipe/cigar smoking, number of cigarettes smoked, years smoked) and BMI, the risk of lung cancer decreased with increase in leisure activity among men in a dose-response manner (*p* for trend = 0.01) (Table II). Men who exercised for at least 4 hr a week during leisure time had a reduced adjusted relative risk (RR = 0.71; 95% CI = 0.52-0.97). No such relationship was observed among women. No statistical association was observed between work activity and lung-cancer risk among men or women.

To study total physical activity, leisure and work activity were combined (Table II). Among men we observed a 27% reduction in risk among active men (non-sedentary) compared with sedentary men (O1, R1) (RR = 0.73; 95% CI = 0.54-0.98). For women, the small number of lung-cancer cases in the reference group prevented any conclusion.

For men active in their leisure time (R2 + R3 + R4) compared with inactive men (R1), the relative risk of lung cancer was particularly reduced for small-cell carcinoma (RR = 0.59, 95%

CI = 0.38-0.94) and adenocarcinoma (RR = 0.65; 95% CI = 0.41-1.05), with no significant association for squamous-cell carcinomas (Table III).

We examined the time-dependent nature of physical activity in 3 of 5 geographical areas, with initial activity assessment between 1974 and 1978 and an update after 3 to 5 years. In this sub-cohort, 142 lung-cancer cases were observed among men and 50 cases among women during a total of 615,000 person-years. Among men, median age at diagnosis was 57.0 years (41.3-66.2) whereas in women it was 55.2 years (48.7-62.7).

The impact of sustained leisure activity over time on lung-cancer risk among men is presented in Table IV. We observed the greatest reduction in lung-cancer risk among those who were active in their leisure time (R3/R4) at both screenings; the men who were inactive in their leisure time at both assessments were used as a reference group (RR = 0.39, 95% CI = 0.18-0.85; *p* for trend, 0.01). A weaker, but consistently reduced, lung-cancer risk was observed among the active men (R3/R4) relative to sedentary men (R1) at the first screening, whereas there was a reduction in risk of almost 40% in these men (R3/R4) at the second screening compared with the inactive men (R1) (RR = 0.62; 95% CI = 0.38-1.01). This association between activity at the second screening and lung-cancer risk was also observed in an inverse dose-response manner (*p* for trend, 0.05).

We also examined models stratified by smoking habits in men, to examine whether the number of cigarettes smoked influenced our results. As a consequence of the small number of lung cancer cases among never-smokers (*n* = 5) and among ex-smokers (*n* = 25), we only performed stratified analyses among current smokers by the number of cigarettes smoked (Table V). Among men smoking 15 cigarettes or more daily, we observed a reduced lung-cancer risk among active men compared with inactive men (RR = 0.59; 95% CI = 0.35-0.97). Similar results were found among men smoking less than 15 cigarettes daily, but these results did not reach significant levels (RR = 0.79; 95% CI = 0.49-1.26). The same analyses were performed for occupational activity groups, but no significant associations were observed.

DISCUSSION

In the present study we found that leisure activity reduced lung-cancer risk among men in a dose-response fashion. This protective effect was also seen for total physical activity, and was strengthened when initial and subsequent leisure activity assessments were combined.

The association between physical activity and lung-cancer risk has not been studied much. Our results however, can be compared,

TABLE III - ADJUSTED RELATIVE RISK (RR)¹ OF LUNG CANCER WITH 95% CONFIDENCE INTERVAL (CI) AMONG DIFFERENT HISTOLOGICAL SUB-TYPES ACCORDING TO RECREATIONAL (R) PHYSICAL ACTIVITY (PhA) AMONG MEN AGED 20-49 YEARS AT ENTRY IN 1972-1978 (FIRST SCREENING)

Recreational PhA	Squamous-cell carcinoma			Adenocarcinoma			Small-cell carcinoma		
	Number of cases	RR	95% CI	Number of cases	RR	95% CI	Number of cases	RR	95% CI
Sedentary (R1)	34	1.0		26	1.0		30	1.0	
Active (R2/R3/R4)	91	0.97	(0.65-1.44)	58	0.65	(0.41-1.05)	53	0.59	(0.38-0.94)

¹Adjusted for age at entry, geographical region, smoking habits (ex-smoking, pipe/cigar smoking, number of cigarettes, years of smoking) and BMI.

TABLE IV - ADJUSTED RELATIVE RISK (RR) OF LUNG CANCER WITH 95% CONFIDENCE INTERVAL (CI) ACCORDING TO RECREATIONAL (R) PHYSICAL ACTIVITY AMONG PARTICIPATING MEN AT BOTH SCREENINGS, AGED 20-49 YEARS IN 1974-78

Recreational PhA	Years of assessment of physical activity								
	1974-78			1977-83			1974-78 and 1977-83		
	Number of cases	RR ¹	95% CI	Number of cases	RR ²	95% CI	Number of cases	RR ³	95% CI
Sedentary (R1)	36	1.0		38	1.0		21	1.0	
Moderate (R2)	74	0.81	(0.54-1.21)	71	0.72	(0.48-1.07)	45	0.54	(0.32-0.91)
Regular exercise (R3, R4)	28	0.84	(0.51-1.39)	29	0.62	(0.38-1.01)	10	0.39	(0.18-0.85)
Trend test	$p = 0.46$			$p = 0.05$			$p = 0.01$		

¹Adjusted for age at entry, geographical region, smoking habits (ex-smoker, pipe/cigars, number of cigarettes, years of smoking) and BMI at first screening (1974-1978). ²Adjusted for age at entry, geographical region, smoking habits (ex-smoker, pipe/cigars, number of cigarettes, years of smoking) and BMI at second screening (1977-1983). ³Adjusted for age at entry at first screening, geographic region, smoking habits (ex-smoker, pipe/cigars, number of cigarettes, years of smoking) and BMI at second screening (1977-1983).

in part, with earlier research (Albanes *et al.*, 1989; Severson *et al.*, 1989; Sellers *et al.*, 1991; Lee and Paffenbarger, 1994). Albanes *et al.* (1989) observed, in their follow-up study of 12,500 subjects, that men in sedentary occupations had twice the risk of lung cancer of men in non-sedentary occupations. They revealed a dose-response profile, also demonstrated by others (Paffenbarger *et al.*, 1987). In contrast, increased incidence (Brownson *et al.*, 1991), as well as increased lung-cancer mortality (Garfinkel and Stellman, 1988), has been reported among physically active, as compared with inactive, men. In the first study, leisure-time activity was not assessed, whereas in the second study smoking habits were not adjusted for.

The accuracy of the self-reported recreational physical activity questions used in the present study has been validated (Wilhelmsen *et al.*, 1976; Holme *et al.*, 1981; Løchen and Rasmussen, 1992). In addition, the results for BMI and blood-lipid profiles across leisure activity groups support the validity of the assessment of physical activity. Our data also have the advantage of repeated individual estimates of total physical activity in a general population. We observed a greater protective effect of moderate and regular exercise when 2 activity assessments over a timespan of 3 to 5 years were combined. This finding may be the result of a reduction in mis-classification of physical activity, and further indicates that physical activity over a longer period is of importance. A similar effect of physical activity on risks of prostate cancer was observed in the Harvard alumni study by Lee *et al.* (1992), who observed a greater protective effect when 2 activity assessments were combined. In the ascertainment of physical activity in the present study, the questionnaire was checked for inconsistency at both screenings. It is, however, possible that reported physical exercise during

leisure time is in excess of its actual occurrence (*i.e.*, "wish" bias). Based on the same questionnaire, one study demonstrated that physical fitness increased with leisure activity (Løchen and Rasmussen, 1992). Moreover, combining 2 assessments increased the precision of physical-activity measurements, and may have reduced such "wish" bias.

Another strength of this study is the completeness of data on lung-cancer cases, thanks to the compulsory reporting of all new cancer cases by hospital departments, pathology laboratories and death certificates in Norway. In addition, there was unbiased selection of participants.

One explanation of the observed association in our study could be that a pre-clinical illness resulting in inactivity could underlie the increased risk seen among sedentary men. However, the relative-risk estimates and tests for linear trend were essentially unchanged in men and women after excluding either 1, 2 or 4 years of follow-up.

The magnitude of the impact of cigarette smoking on lung-cancer risk is well documented (Doll and Peto, 1978; Risch *et al.*, 1993). One could therefore argue that our findings are residual effects of smoking which cannot be entirely eliminated by statistical adjustments. However, adjustments were made carefully for current and former smoking behaviour, ex-smoking and number of cigarettes smoked daily, as well as for years of current smoking. In addition, among men smoking 15 cigarettes or more daily, a reduced lung-cancer risk was observed among those men who were active in their leisure time compared with sedentary men. Further, it is likely that those active men who had smoked heavily had consumed about the same number of cigarettes during their lifetime

TABLE V - ADJUSTED RELATIVE RISK (RR)¹ OF LUNG CANCER WITH 95% CONFIDENCE INTERVAL (CI) AND RECREATIONAL (R) PHYSICAL ACTIVITY STRATIFIED BY NUMBER OF CIGARETTES SMOKED IN CURRENT CIGARETTES SMOKING MEN AT THE FIRST SCREENING (1972-78)

Physical activity (PhA)	<15 Cigarettes			15 Cigarettes+		
	Number of cases	RR	95% CI	Number of cases	RR	95% CI
Recreational PhA						
Sedentary (R1)	42	1.0		71	1.0	
Moderate (R2)	94	0.77	(0.53-1.11)	96	0.71	(0.52-0.96)
Regular exercise (R3, R4)	31	0.79	(0.49-1.26)	20	0.59	(0.35-0.97)
Trend test	$p = 0.28$			$p = 0.01$		

¹Adjusted for age at entry, geographical region, pipe/cigar smoking, years of smoking and BMI.

as the sedentary men who were currently heavy smokers, because the number of reported years of smoking were the same in the 2 groups. It is plausible, therefore, that physical activity reduces lung-cancer risk, as observed in our study, and is not merely a residual effect of smoking.

In this study, we analyze the association between physical activity and lung-cancer risk by histological sub-types. We observed the inverse association between physical activity and lung cancer to be strongest for small-cell carcinoma, less marked for adenocarcinoma, with no association observed for squamous-cell carcinoma. Although cigarette smoking appears to induce lung cancer for all histological types, the magnitude of smoking-related lung-cancer risk by cell type is strongest for squamous-cell and small-cell carcinoma, and weakest for adenocarcinoma (Vena *et al.*, 1985; Brownson *et al.*, 1992; McDuffie *et al.*, 1993). However, the reliability of histological classification may be a problem, although mis-classifications will make differences in risk smaller, among different histological types, if they are non-differential. In the present study, it is probable that mis-classification is non-differential. Due to the few cases among never-smokers and ex-smokers, we could not analyze these sub-groups. In a cohort study by Lee and Paffenbarger (1994), physically active non-smokers had a reduced lung-cancer risk. When stratified by number of cigarettes smoked (more or less than 15 cigarettes), the findings of a reductive effect of leisure activity on lung-cancer risk were consistent in both groups. Overall, the data still suggest that physical activity reduces lung-cancer risk in men.

One interpretation of these observations could be that physical activity is sufficient to counteract carcinogenesis in groups with exposure to cigarette smoke or other carcinogenic agents. One plausible biological mechanism may act through the increased pulmonary function observed with increased exercise (Kuller *et al.*, 1990; Higgins *et al.*, 1991). Increased pulmonary ventilation and perfusion could reduce the interaction time and concentration of any carcinogenic agent in the airways. High or moderate levels of physical activity may thereby reduce the production of free radicals and carcinogenic metabolites produced from, for example, smoking (Tappia *et al.*, 1995; Morrow *et al.*, 1995). In our study, this supposition is supported by a positive dose-response effect with no threshold effect.

Another mechanism may be that physical activity resulting from increased pulmonary function influences particle deposition. The degree of carcinogenicity of cigarette smoke or other agents may be related to the location of particle deposition in the airways. The geometrical site of preferential particle deposition in the central airway has been demonstrated as the favoured site of cancer induction (Byers *et al.*, 1984; Yang *et al.*, 1989; Martonen, 1992). As small-cell carcinoma and, particularly, adenocarcinoma are more often located in the periphery of the lung, increased pulmonary function could be more important for these sub-types. This

may explain the lack of a protective effect of physical activity observed for squamous-cell carcinoma, the protective effect for small-cell carcinoma and adenocarcinoma of the lung in our study.

The lack of association between lung cancer and physical activity among women in our study may be explained partly by the small number of cases. Further, a narrow range of variation of both occupational and leisure physical activity in women could reduce our ability to find such a relationship. An association may therefore be present but undetectable. An indication for this could be the consistent reduction in risk among occupationally active women compared with sedentary women, supported by findings in a nested case-control study in which active women had a 60% reduction in risk of lung cancer (Sellers *et al.*, 1991). A few cases in the present study exclude any conclusion regarding the association of physical activity with lung-cancer risk in women.

Occupational physical activity did not appear, in the present study, to have the same protective effect on lung-cancer risk as leisure activity. Occupational activity could reflect a more static activity, and this is supported by the observation that subjects who carry out more leisure activities are leaner and have lower serum-lipid concentrations, a pattern not observed among occupationally active subjects. Static activity may not influence lung-cancer risk through the same biological mechanism as leisure exercise. However, total physical activity reduced lung-cancer risk among active men compared with sedentary men. This indicates a weak negative or no association between occupational physical activity and lung-cancer risk in men.

CONCLUSION

The observed negative dose-response association between recreational physical activity and lung cancer in men may be explained by exercise-induced improvement of pulmonary function and reduced carcinogenic effect of any environmental factor. The observed protective effect on small-cell carcinoma and adenocarcinoma, but not on squamous-cell carcinoma, supports explanations other than a residual smoking effect. Further studies, including information on physical fitness and pulmonary function, are needed in which both gender and histological sub-types are taken into account, together with smoking habits and physical activity over time.

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REFERENCES

- ALBANES, D., BLAIR, A.A. and TAYLOR, P.R., Physical activity and risk of cancer in the NHANES I population. *Amer. J. publ. Hlth.*, **79**, 744-750 (1989).
- BROWNSON, R.C., CHANG, J.C. and DAVIS, J.R., Gender and histologic type variations in smoking-related risk of lung cancer. *Epidemiology*, **3**, 61-64 (1992).
- BROWNSON, R.C., CHANG, J.C., DAVIS, J.R. and SMITH, C.A., Physical activity on the job and cancer in Missouri. *Amer. J. publ. Hlth.*, **81**, 639-642 (1991).
- BYERS, T.E., VENA, J.E. and RZEPKA, T.F., Predilection of lung cancer for the upper lobes: an epidemiologic inquiry. *J. nat. Cancer Inst.*, **72**, 1271-1275 (1984).
- CANCER REGISTRY OF NORWAY, *Cancer in Norway 1992*. Oslo (1995).
- DOLL, R. and PETO, R., Cigarette smoking and bronchial carcinoma: dose and time relationships among regular smokers and lifelong non-smokers. *J. Epidemiol. Community Hlth.*, **32**, 303-313 (1978).
- GARFINKEL, L. and STELLMAN, S.D., Mortality by relative weight and exercise. *Cancer*, **62**, 1844-1850 (1988).
- HIGGINS, M., KELLER, J.B., WAGENKNECHT, L.E., TOWNSEND, M.C., SPARROW, D., JACOBS, D.R. and HUGHES, G., Pulmonary function and cardiovascular risk-factor relationships in black and in white young men and women. *Chest*, **99**, 315-322 (1991).
- HOLME, I., HELGELAND, A., HJERMANN, LEREN, P. and LUND-LARSEN, P.G., Physical activity at work and at leisure in relation to coronary risk factors and social class. A 4-year mortality follow-up. The Oslo Study. *Acta. med. scand.*, **209**, 277-283 (1981).
- KULLER, L.H., OCKENE, J., MEILAHN, E. and SVENDSEN, K.H., Relation of forced expiratory volume in one second (FEV₁) to lung-cancer mortality in the multiple-risk-factor intervention trial (MRFIT). *Amer. J. Epidemiol.*, **132**, 265-274 (1990).
- LEE, I.-M. and PAFENBARGER, R.S. JR., Physical activity and its relation to cancer risk: a prospective study of college alumni. *Med. Sci. Sports Exerc.*, **26**, 831-877 (1994).

- LEE, I.-M., PAFFENBARGER, R.S. JR. and HSIEH, C.-C., Physical activity and risk of prostatic cancer among college alumni. *Amer. J. Epidemiol.*, **135**, 169-179 (1992).
- LOCHEN, M.-L. and RASMUSSEN, K., The Tromsø study: physical fitness, self-reported physical activity, and their relationship to other coronary risk factors. *J. Epidemiol. Community Hlth.*, **26**, 103-107 (1992).
- MARTONEN, T.B., Deposition patterns of cigarette smoke in human airways. *Amer. ind. Hyg. Ass. J.*, **53**, 6-18 (1992).
- MCDUFFIE, H.H., KLAASSEN, D.J. and DOSMAN, J.A., Determinants of cell type in patients with cancer of the lungs. *Chest*, **98**, 1187-1193 (1993).
- MORROW, J.D., FREI, B., LONGMIRE, A.W., GAZIANO, J.M., LYNCH, S.M., SHYR, Y., STRAUSS, W.E., OATES, J.A. and ROBERTS, L.J., Increase in circulating product of lipid peroxidation (F₂-isoprostanes) in smokers. Smoking as a cause of oxidative damage. *New Engl. J. Med.*, **332**, 1198-1203 (1995).
- NOMURA, A., STEMMERMAN, G.N., CHYOU, P.-H., MARCUS, E.B. and BUIST, A.S., Prospective study of pulmonary function and lung cancer. *Amer. Rev. respir. Dis.*, **144**, 307-311 (1991).
- PAFFENBARGER, R.S. JR., HYDE, R.T. and WING, A.L., Physical activity and incidence of cancer in diverse populations: a preliminary report. *Amer. J. clin. Nutr.*, **45**, 312-317 (1987).
- RISCH, H.A., HOWE, G.R., MEERA, J., BURCH, J.D., HOLOWATY, E.J. and MILLER, A.B., Are female smokers at higher risk for lung cancer than male smokers? A case-control analysis by histologic type. *Amer. J. Epidemiol.*, **138**, 281-293 (1993).
- SELLERS, T.A., POTTER, J.D. and FOLSOM, A.R., Association of incident lung cancer with family history of female reproductive cancers: the Iowa women's health study. *Genet. Epidemiol.*, **8**, 199-208 (1991).
- SEVERSON, R.K., NOMURA, A.M.Y., GROVE, J.S. and STEMMERMANN, G.N., A prospective analysis of physical activity and cancer. *Amer. J. Epidemiol.*, **130**, 522-529 (1989).
- TAPPIA, P.S., TROUGHTON, K.L., LANGLEY-EVANS, S.C. and GRIMBLE, R.F., Cigarette smoking influences cytokine production and anti-oxidant defences. *Clin. Sci.*, **88**, 485-489 (1995).
- THUNE, I. and LUND, E., Physical activity and the risk of prostate and testicular cancer: a cohort study of 53,000 Norwegian men. *Cancer Causes Control*, **5**, 549-556 (1994).
- THUNE, I. and LUND, E., Physical activity and the risk of colorectal cancer in men and women. *Brit. J. Cancer*, **73**, 1134-1140 (1996).
- TRAVIS, W.D., TRAVIS, L.B. and DEVESA, S.S., Lung cancer. *Cancer*, **75**, 191-202 (1995).
- VENA, J.E., BYERS, T.E., COOKFAIR, D. and SWANSON, M., Occupation and lung-cancer risk. An analysis by histologic sub-types. *Cancer*, **56**, 910-917 (1985).
- WEISS, S.T., SEGAL, M.R., SPARROW, D. and WAGER, C., Relation of FEV₁ and peripheral-blood-leucocyte count to total mortality. The normative aging study. *Amer. J. Epidemiol.*, **142**, 493-498 (1995).
- WILHELMSSEN, L., TIBBLIN, G., AURELL, M., BJURE, J., EKSTRÖM-JODAL, B. and GRIMBY, G., Physical activity, physical fitness and risk of myocardial infarction. *Advanc. Cardiol.*, **18**, 217-230 (1976).
- YANG, C.P., GALLAGHER, R.P., WEISS, N.S., BAND, P.R., THOMAS, D.B. and RUSSEL, D.A., Differences in incidence rates of cancers of the respiratory tract by anatomic sub-site and histologic type: an etiologic implication. *J. nat. Cancer Inst.*, **81**, 1828-1831 (1989).

Paper IV

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PHYSICAL ACTIVITY AND THE RISK OF BREAST CANCER

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ABSTRACT

Background Because physical activity may affect hormonal concentrations and energy balance, we decided to investigate whether everyday exercise is related to the risk of breast cancer.

Methods During 1974 to 1978 and 1977 to 1983, a total of 25,624 women, 20 to 54 years of age at entry, enrolled in health surveys and answered questionnaires about leisure-time and work activity.

Results During a median follow-up of 13.7 years, we identified 351 cases of invasive breast cancer among the 25,624 women in the cohort. Greater leisure-time activity was associated with a reduced risk of breast cancer, after adjustments for age, body-mass index (the weight in kilograms divided by the square of the height in meters), height, parity, and county of residence (relative risk, 0.63; 95 percent confidence interval, 0.42 to 0.95), among women who exercised regularly, as compared with sedentary women (P for trend=0.04). In regularly exercising women, the reduction in risk was greater in premenopausal women than in postmenopausal women, and greater in younger women (<45 years at study entry) than in older women (≥ 45 years) (relative risk, 0.38; 95 percent confidence interval, 0.19 to 0.79). In stratified analyses the risk of breast cancer was lowest in lean women (body-mass index, <22.8) who exercised at least four hours per week (relative risk, 0.28; 95 percent confidence interval, 0.11 to 0.70). The risk was also reduced with higher levels of activity at work, and again there was a more pronounced effect among premenopausal than postmenopausal women.

Conclusions Physical activity during leisure time and at work is associated with a reduced risk of breast cancer. (N Engl J Med 1997;336:1269-75.)

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VIGOROUS physical training¹⁻⁵ and even moderate exercise⁶⁻⁹ can interrupt the menstrual cycle, perhaps by suppressing the pulsatile release of gonadotropin-releasing hormone.^{10,11} This effect of physical activity may lower a woman's cumulative exposure to estro-

gen and progesterone, thereby inhibiting carcinogenesis in the breast.¹²⁻²² Energy balance might also influence the risk of breast cancer. Caloric restriction in rodents reduces the proliferative activity of the mammary glands²³ and inhibits carcinogenesis.^{24,25} However, the effect of energy balance, as indicated by energy intake, body-mass index (the weight in kilograms divided by the square of the height in meters), and energy expenditure, on the risk of breast cancer has not been examined thoroughly in humans.

In this study we evaluated the influence of physical activity, both at work and during leisure time, on the risk of breast cancer in a cohort of 25,624 premenopausal and postmenopausal women. Data on parity, dietary factors, and body-mass index allowed adjustment for potentially confounding factors, and reassessment of physical activity after three to five years gave an indication of the effect of sustained physical activity on the risk of breast cancer.

METHODS

Study Population

From 1974 to 1978, the National Health Screening Service invited people in three counties in Norway (Oppland, Sogn og Fjordane, and Finnmark) to participate in a survey of risk factors for cardiovascular disease. All women who were 35 to 49 years of age and a random sample of 10 percent of those who were 20 to 34 years of age were invited. In four municipalities in Finnmark all women who were 20 to 34 years of age were invited. A comprehensive description of these populations has been published previously.²⁶ A total of 31,556 women were invited to participate, and 28,621 (91 percent) actually did.

All women in this survey as well as a random sample of women who were 20 to 39 years of age were invited to participate in a second survey three to five years later (1977 to 1983). Of these 34,378 women, 31,209 (91 percent) participated.²⁷ This second survey was used as the base line, because no information on parity and dietary factors was collected during the first survey.

Each woman received a written invitation to participate, to-

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gether with a one-page questionnaire. The participants were asked to answer the questionnaire and bring it to the clinical examination. At screening, trained nurses checked the questionnaire for inconsistencies regarding physical activity and menopausal status, measured weight and height, and collected blood samples.

During screening in the second survey, the participants were asked to fill out a food-frequency questionnaire, to be returned by mail. After one reminder, 25,892 (83 percent) returned the questionnaire. The energy and fat intakes for each woman were derived from the sum of all food consumed. The semiquantitative food-frequency questionnaire that we used has been described in detail and validated.^{28,29}

Assessment of Physical Activity

Self-reported categories of physical activity during leisure hours in the year preceding each survey were assessed when the women entered the study and graded from 1 to 4 according to the participant's usual level of physical activity. A grade of 1 was assigned to those whose leisure time was spent reading, watching television, or engaging in other sedentary activities; a grade of 2 to those who spent at least four hours a week walking, bicycling, or engaging in other types of physical activity; a grade of 3 to those who spent at least four hours a week exercising to keep fit and participating in recreational athletics; and a grade of 4 to those who engaged in regular, vigorous training or participating in competitive sports several times a week. The self-reported level of physical activity during work hours in the preceding year was also graded on a four-point scale. A grade of 1 was assigned to those whose work was mostly sedentary; a grade of 2 to those whose job involved a lot of walking; a grade of 3 to those whose job required a lot of lifting and walking; and a grade of 4 to those engaged in heavy manual labor.

Two identical assessments of leisure-time activity were made at an interval of three to five years, and the results were combined for all groups. Women who reported moderate (grade 2) or regular (grade 3 or 4) exercise during leisure time in the first survey and regular exercise (grade 3 or 4) in the second survey were characterized as being consistently physically active. Women who were sedentary (grade 1) during leisure time in both surveys were characterized as being consistently sedentary. The women who were neither consistently sedentary nor consistently active during leisure time were characterized as being moderately active.

Follow-up and Identification of Cases of Breast Cancer

We followed a total of 25,707 women who had not been given a diagnosis of cancer before our base-line survey (1977 to 1983). We used the participants' national 11-digit personal identification numbers to identify every incident case of breast cancer reported to the Cancer Registry of Norway and Statistics Norway through the end of follow-up (December 31, 1994). A total of 98 percent of the cases were verified histologically. Women in whom cancer developed ($n=72$) or who died ($n=11$) within the first year of the study were excluded from the analyses to account for the possibility that undiagnosed cancer or severe illness might influence the level of physical activity. Through a linkage to the Central Population Register at Statistics Norway, we obtained information concerning the reproductive history of each woman, including the date of birth of each liveborn child through December 31, 1992, and deaths in the cohort through December 31, 1994.

The ultimate study cohort consisted of 25,624 women who participated in both surveys (age range, 20 to 69 years) during 359,930 person-years of follow-up.

Statistical Analysis

Base-line variables were adjusted for age and compared by analysis of covariance. Cox proportional-hazards regression analysis was carried out to investigate the simultaneous effect of physical activity and covariates on the incidence of breast cancer. To calculate the risk of breast cancer, women were observed for the de-

velopment of breast cancer from entry into the study to the date of diagnosis of any cancer, the time of death, or the end of follow-up, whichever event came first. In the analysis, grades 3 and 4 of leisure-time activity were merged because of the small numbers of women with a grade of 4 in both surveys (48 women in the first survey and 57 in the second survey). As a reference group we used women who were sedentary at work or during leisure time.

In the analyses, we adjusted for age at entry (a continuous variable), county of residence, number of children, age at birth of first child, intake of total fat and energy, and body-mass index. Women who reported that they were premenopausal at base line were treated as premenopausal until they reached the age of 50 during follow-up, at which time they were considered postmenopausal. Women who reported that they were postmenopausal at base line were treated as postmenopausal.

Because there were few women with breast cancer who were sedentary both at work and during leisure time, the effect of this combination on the risk of breast cancer could not be analyzed. All significance tests were two-tailed, and the level of significance was set at 5 percent. The analyses were performed with the SAS statistical package version 6.11.

RESULTS

There were 351 incident cases of breast cancer (100 among premenopausal women and 251 among postmenopausal women) among 25,624 women. The mean length of follow-up was 14.0 years (median, 13.7), and the median age at diagnosis was 54.7 years (range, 36.3 to 68.0).

Table 1 gives the base-line characteristics of the participants. Two thirds of the women reported moderate activity during leisure time, whereas 15 percent exercised regularly. Only 14 percent reported being sedentary at work, whereas 20 percent reported lifting and 5 percent reported doing heavy manual labor. Women who reported regularly exercising during leisure time did not differ from women who were inactive during their leisure time with respect to age at entry or number of children, but they tended to be taller and to have a lower body-mass index, a relatively low ratio of total cholesterol to high-density lipoprotein (HDL) cholesterol in serum, lower serum triglyceride levels, and higher HDL cholesterol levels. Women whose work involved lifting or heavy manual labor had a higher body-mass index and more children than those engaged in sedentary work. Energy intake was positively related to physical activity, but the association was more pronounced with work activity than with leisure-time activity.

We analyzed other possible age-adjusted risk factors for breast cancer at base line and found a 28 percent increase in risk for each additional 6 cm of height and a 13 percent reduction in risk for each child. An older maternal age at the birth of a first child was associated with a borderline increase in risk, whereas body-mass index (in the group as a whole or in the subgroups of premenopausal and postmenopausal women), energy intake, and total fat intake did not influence the overall risk of breast cancer (data not shown).

Table 2 shows the relation between the level of

PHYSICAL ACTIVITY AND THE RISK OF BREAST CANCER

TABLE 1. BASE-LINE CHARACTERISTICS OF THE WOMEN ACCORDING TO THE LEVEL OF PHYSICAL ACTIVITY IN THE 1977-1983 SURVEY.*

CHARACTERISTIC	LEVEL OF ACTIVITY DURING LEISURE TIME			LEVEL OF ACTIVITY AT WORK			
	SEDENTARY (N=4410)	MODERATE (N=17,481)	REGULAR EXERCISE (N=3719)	SEDENTARY (N=3534)	WALKING (N=15,385)	LIFTING (N=5240)	HEAVY MANUAL LABOR (N=1385)
Age at entry (yr)	45.1	45.0	45.4	44.1	45.0	45.4	46.8
Body mass index	25.5	24.8	24.5	24.3	24.8	25.2	25.6
Height (cm)	161.9	162.7	163.3	163.4	162.5	162.4	162.8
Triglycerides (mg/dl)†	139.9	127.5	124.0	125.8	130.2	130.2	122.2
HDL cholesterol (mg/dl)‡	54.5	56.1	57.3	55.7	55.7	56.5	58.0
Total cholesterol: HDL cholesterol	4.71	4.50	4.34	4.47	4.55	4.51	4.27
Parity							
No. of children	2.7	2.6	2.7	2.1	2.7	2.8	2.9
Mother's age at first birth (yr)	24.2	24.4	24.5	24.6	24.4	24.1	24.6
Daily energy intake (kJ)	5725	5761	5797	5561	5716	5854	6434
Total daily fat intake (g)	55.4	54.8	54.7	52.4	54.6	55.9	61.5
Daily smoking (%)	40.1	34.5	31.6	35.8	35.3	36.4	26.1

*All variables except age were adjusted for age. All values except those for daily smoking are means. Subjects for whom information concerning certain variables was missing are not included.

†To convert values for triglycerides to millimoles per liter, multiply by 0.01129.

‡To convert values for HDL cholesterol to millimoles per liter, multiply by 0.02586.

TABLE 2. ADJUSTED RELATIVE RISK OF BREAST CANCER ACCORDING TO THE LEVEL OF PHYSICAL ACTIVITY DURING LEISURE TIME AND AT WORK IN THE 1977-1983 SURVEY.*

LEVEL OF PHYSICAL ACTIVITY	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)†	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)‡
During leisure time				
Sedentary	66	1.00	65	1.00
Moderate	249	0.98 (0.75-1.28)	245	0.93 (0.71-1.22)
Regular exercise	36	0.67 (0.44-1.00)	36	0.63 (0.42-0.95)
P for trend		0.08		0.04
At work				
Sedentary	62	1.00	61	1.00
Walking	212	0.76 (0.57-1.01)	210	0.84 (0.63-1.12)
Lifting	64	0.66 (0.47-0.94)	63	0.74 (0.52-1.06)
Heavy manual labor	12	0.46 (0.25-0.86)	11	0.48 (0.25-0.92)
P for trend		0.004		0.02

*The sedentary group is the reference group. CI denotes confidence interval. Subjects for whom information concerning certain variables was missing are not included.

†Variables were adjusted for age at entry.

‡Variables were adjusted for age at entry, body-mass index, height, county of residence, and number of children.

leisure-time or work activity and the overall risk of breast cancer. After adjustment for age and with the sedentary group as the reference group, the relative risk of breast cancer was reduced among women whose jobs involved walking, lifting, or heavy manual labor. Adjustments for other factors (body-mass index, county of residence, number of children, and height) in addition to age changed the risk estimates only slightly. Further adjustments for age at first

birth or dietary factors (energy intake, total fat intake, and fiber intake) did not influence our estimates of relative risk and were omitted from the final model. A 52 percent reduction in risk was observed among the women who reported doing heavy manual labor (relative risk, 0.48; 95 percent confidence interval, 0.25 to 0.92). The overall adjusted risk of breast cancer decreased in a dose-response manner with increasing activity level during leisure time

TABLE 3. ADJUSTED RELATIVE RISK OF BREAST CANCER ACCORDING TO MENOPAUSAL STATUS AND THE LEVEL OF PHYSICAL ACTIVITY IN THE 1977-1983 SURVEY.*

LEVEL OF PHYSICAL ACTIVITY	PREMENOPAUSAL WOMEN		POSTMENOPAUSAL WOMEN	
	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)
During leisure time				
Sedentary	20	1.00	45	1.00
Moderate	68	0.77 (0.46-1.27)	177	1.00 (0.72-1.39)
Regular exercise	10	0.53 (0.25-1.14)	26	0.67 (0.41-1.10)
P for trend		0.10		0.15
At work				
Sedentary	22	1.00	39	1.00
Walking	62	0.82 (0.50-1.34)	148	0.87 (0.61-1.24)
Lifting or heavy manual labor	14	0.48 (0.24-0.95)	60	0.78 (0.52-1.18)
P for trend		0.03		0.24

*The sedentary group is the reference group. Variables were adjusted for age at entry, body-mass index, height, county of residence, and number of children. CI denotes confidence interval. Subjects for whom information concerning certain variables was missing are not included.

TABLE 4. ADJUSTED RELATIVE RISK OF BREAST CANCER ACCORDING TO BODY-MASS INDEX AND THE LEVEL OF PHYSICAL ACTIVITY DURING LEISURE TIME IN THE 1977-1983 SURVEY.*

LEVEL OF PHYSICAL ACTIVITY	BODY-MASS INDEX, <22.8		BODY-MASS INDEX, 22.8-25.7		BODY-MASS INDEX, >25.7	
	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)
Sedentary	21	1.00	14	1.00	30	1.00
Moderate	104	1.12 (0.70-1.79)	73	1.09 (0.61-1.93)	68	0.70 (0.46-1.08)
Regular exercise	6	0.28 (0.11-0.70)	14	0.96 (0.45-2.01)	16	0.83 (0.45-1.53)
P for trend		0.02		0.90		0.36

*The sedentary group is the reference group. Variables were adjusted for age at entry, height, county of residence, and number of children. CI denotes confidence interval. Subjects for whom information concerning certain variables was missing are not included.

(P for trend = 0.04). Women who exercised at least four hours a week during leisure time had a 37 percent reduction in the risk of breast cancer (relative risk, 0.63; 95 percent confidence interval, 0.42 to 0.95).

When the group was divided according to menopausal status (Table 3), a consistently inverse association was observed between the level of leisure-time activity and the premenopausal risk of breast cancer; the adjusted relative risk declined to 0.77 (95 percent confidence interval, 0.46 to 1.27) and further to 0.53 (95 percent confidence interval, 0.25 to 1.14) as the level of activity increased (P for trend = 0.10). A weaker association was observed between the level of leisure-time activity and the postmenopausal risk of breast cancer. The inverse association between the level of activity at work and the risk of breast cancer was also pronounced among

premenopausal women; among premenopausal women whose jobs involved lifting or heavy manual labor, the relative risk was 0.48 (95 percent confidence interval, 0.24 to 0.95).

We also divided the cohort into women who were younger than 45 years of age at entry and those who were 45 or older. Among those younger than 45 years at entry for whom data were complete (of whom breast cancer developed in 138; mean age at diagnosis, 48.3 years), the adjusted relative risk declined to 0.80 (95 percent confidence interval, 0.52 to 1.22) and further to 0.38 (95 percent confidence interval, 0.19 to 0.79) as the level of activity during leisure time increased (P for trend = 0.01). The respective adjusted relative risks were 1.03 (95 percent confidence interval, 0.72 to 1.48) and 0.84 (95 percent confidence interval, 0.51 to 1.39) (P for trend =

TABLE 5. ADJUSTED RELATIVE RISK OF BREAST CANCER ACCORDING TO BODY-MASS INDEX AND OVERALL LEVEL OF PHYSICAL ACTIVITY DURING LEISURE TIME IN THE 1974-1978 AND 1977-1983 SURVEYS.*

OVERALL LEVEL OF PHYSICAL ACTIVITY†	ALL WOMEN		BODY-MASS INDEX, <22.8		BODY-MASS INDEX, 22.8-25.7		BODY-MASS INDEX, >25.7	
	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)‡	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)§	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)§	CASES OF BREAST CANCER	RELATIVE RISK (95% CI)§
Consistently sedentary	29	1.00	13	1.00	7	1.00	9	1.00
Moderately active	283	0.90 (0.61-1.32)	112	0.76 (0.43-1.35)	81	0.87 (0.40-1.88)	90	1.14 (0.57-2.27)
Consistently active	34	0.67 (0.40-1.10)	6	0.23 (0.09-0.60)	13	0.83 (0.33-2.09)	15	1.38 (0.60-3.17)
P for trend		0.09		0.002		0.73		0.42

*The sedentary group is the reference group. CI denotes confidence interval.

†Subjects with a consistently sedentary level of activity were classified as sedentary in both surveys (grade 1). Subjects who remained active, reporting moderate (grade 2) or regular (grade 3 or 4) exercise in the first survey and regular exercise (grade 3 or 4) in the second survey, were classified as being consistently active. Subjects were classified as moderately active if they did not meet the criteria for the other two categories.

‡Variables were adjusted for age at entry, body-mass index, height, county of residence, and number of children.

§Variables were adjusted for age at entry, height, county of residence, and number of children.

0.54) among those for whom data were complete who were 45 years of age or older at entry (of whom breast cancer developed in 208; mean age at diagnosis, 58.2 years). These values indicate that physical activity had a protective effect, particularly with respect to the risk of breast cancer before and soon after menopause.

We examined models stratified according to body-mass index (Table 4). Among lean (body-mass index, <22.8), regularly exercising women, the risk of breast cancer was reduced by 72 percent (relative risk, 0.28; 95 percent confidence interval, 0.11 to 0.70). No such association was observed in the middle or upper thirds of body-mass index among regularly exercising women. In models stratified according to both body-mass index and menopausal status, this association was seen among both premenopausal and postmenopausal lean women (data not shown).

In the second survey 61.2 percent of the participants reported the same level of leisure-time activity as in the first survey, 23.5 percent reported an increased level, and 15.3 percent reported a reduced level. By combining these two assessments of leisure-time activity, we observed that the relative risk declined to 0.23 (95 percent confidence interval, 0.09 to 0.60) as the level of sustained activity increased in lean (body-mass index, <22.8) women (*P* for trend = 0.002) (Table 5). This protective effect across increasing levels of sustained leisure-time activity was observed in both lean premenopausal women (relative risk, 0.23; 95 percent confidence interval, 0.06 to 0.88; *P* for linear trend = 0.02) and lean postmenopausal women (relative risk, 0.24; 95 percent confidence interval, 0.06 to 0.96; *P* for linear trend = 0.03).

DISCUSSION

Our results support the idea that physical activity protects against breast cancer, particularly among premenopausal and younger postmenopausal women. Activity during both leisure time and work reduced the overall risk. There was a significant inverse dose-response relation between leisure-time activity and the risk of breast cancer. The protective effect was evident among lean premenopausal and postmenopausal women, and repeated assessment emphasized the preventive effect of physical activity.

The overall reduction in the risk of breast cancer among active women is consistent with findings in other cohort^{15,17} and case-control^{19,22} studies, but at variance with the findings of a few others.^{30,31} In one of these discrepant studies,³¹ most of the women were older than in the present study and breast cancer was diagnosed mainly among postmenopausal women. In the other,³⁰ physical activity at college was assessed 35 to 70 years before the diagnosis of breast cancer, and no adjustments were made for potential confounding factors. Our finding of a protective effect of work-related activity on the risk of breast cancer is also in agreement with other studies.^{18,32,33}

Precise assessment of physical activity is difficult in a population-based cohort. The accuracy of the levels of leisure-time activity reported on the questionnaire that we used has been validated previously.^{34,36} Since the level of leisure-time activity correlates with the degree of physical fitness,^{34,36} our observation that recreationally active women tended to be leaner than inactive women and had serum lipid profiles associated with regular exercise strengthens the validity of our assessments. Energy intake was also positively

related to both leisure-time and work activities, particularly work activities.

Repeated assessment of leisure-time activity is important in any analysis of the effect of sustained activity on the risk of breast cancer. The protective effect was notable among lean women who were consistently active during their leisure time. In combining the two assessments for each woman, we may also have increased the precision of our assessment of physical-activity levels, but we cannot differentiate the effect of sustained activity from any misclassification.

The population-based approach and the high participation rate in our study reduced selection bias. The almost complete reporting of incident cases of breast cancer also strengthens our results. Age at menarche was not available and could have confounded our results, but this is not likely, since an increased risk of only 4 percent was observed for each year of earlier age at menarche in a similar study population in Norway.³⁷

Information about the use of hormonal contraceptives was not available, although recent meta-analyses suggest that there is only a small increase in the risk of breast cancer among the youngest women who commonly use hormonal contraceptives.³⁸ It is probable that this information would not have confounded our results to any large extent.

How does physical activity influence the development of breast cancer? The propensity to be physically active may be inherited,³⁹ so the genotype may influence both physical activity and the predisposition to breast cancer. Social and cultural influences on exercise and energy balance seem to be more important than genetic factors,^{39,40} which points to leisure-time activity as an independent and modifiable variable with regard to its effect on the risk of breast cancer.

A reduction in the cumulative exposure to cyclic estrogens and progesterone may in part explain the preventive effect of both leisure-time and work activity. Over the long term, vigorous training and moderate leisure-time activity may decrease estradiol and progesterone secretion,^{3,6,41} reduce the length of the luteal phase,^{10,42} induce anovulation,^{7,8,41,43} delay menarche,^{4,5} and cause secondary amenorrhea.^{2,12}

Physical activity influences energy balance, and experimental studies have shown that calorie restrictions inhibit mammary carcinogenesis.^{24,25,44} Anthropometric measures such as height, body-mass index, and weight gain have been used as biomarkers of calorie intake, and increased values have been reported to be risk factors for breast cancer in humans.⁴⁵⁻⁴⁸ A diet involving a high energy intake has also been associated with early age at menarche,^{5,49} and this finding supports the hypothesis that increased net energy may increase the cumulative hormonal levels that are of importance for carcinogenesis of the breast.

Women who were active during leisure time reported only a slightly higher total energy intake than sedentary women, and they tended to be leaner, indicating that their net available energy was lower. The greater protective effect of leisure-time activity against breast cancer in lean women indicates that there may be an optimal energy balance that inhibits mammary carcinogenesis.

Triglycerides are known to displace estradiol from its tight binding to the sex hormone-binding globulin, which is found in low levels in obese women,⁵⁰ and thus triglycerides increase levels of free estradiol. Serum levels of triglycerides were higher in sedentary women than in women who were more active during their leisure time; thus, exposure to estrogen may be greater in inactive women. This underscores the importance of avoiding obesity if physical activity is to have an optimal inhibitory effect on the risk of breast cancer.

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REFERENCES

- Warren MP. The effects of exercise on pubertal progression and reproductive function in girls. *J Clin Endocrinol Metab* 1980;51:1150-7.
- Frisch RE, Gorz Welbergen AV, McArthur JW, et al. Delayed menarche and amenorrhea of college athletes in relation to age at onset of training. *JAMA* 1981;246:1559-63.
- Bullen BA, Skrinar GS, Beitins IZ, von Mering G, Turnbull BA, McArthur JW. Induction of menstrual disorders by strenuous exercise in untrained women. *N Engl J Med* 1985;312:1349-53.
- Moisan J, Meyer F, Gingras S. Leisure physical activity and age at menarche. *Med Sci Sports Exerc* 1991;23:1170-5.
- Merzenich H, Boeing H, Wahrendorf J. Dietary fat and sports activity as determinants for age at menarche. *Am J Epidemiol* 1993;138:217-24.
- Ellison PT, Lager C. Moderate recreational running is associated with lowered salivary progesterone profiles in women. *Am J Obstet Gynecol* 1986;154:1000-3.
- Bernstein L, Ross RK, Lobo RA, Hanisch R, Kraio MD, Henderson BE. The effects of moderate physical activity on menstrual cycle patterns in adolescence: implications for breast cancer prevention. *Br J Cancer* 1987;55:681-5.
- Prior JC, Vigna YM, Schechter MT, Burgess AE. Spinal bone loss and ovulatory disturbances. *N Engl J Med* 1990;323:1221-7. [Erratum, *N Engl J Med* 1993;328:1724.]
- Harlow SD, Matanoski GM. The association between weight, physical activity, and stress and variation in the length of the menstrual cycle. *Am J Epidemiol* 1991;133:38-49.
- Loucks AB, Mortola JF, Girton L, Yen SSC. Alterations in the hypothalamic-pituitary-ovarian and the hypothalamic-pituitary-adrenal axes in athletic women. *J Clin Endocrinol Metab* 1989;68:402-11.
- Walberg Rankin J, Franke WD, Gwazdauskas FC. Response of beta-endorphin and estradiol to resistance exercise in females during energy balance and energy restriction. *Int J Sports Med* 1992;13:542-7.
- Henderson BE, Ross RK, Judd HL, Kraio MD, Pike MC. Do regular ovulatory cycles increase breast cancer risk? *Cancer* 1985;56:1206-8.
- Pike MC, Spicer DV, Dahmouh L, Press MF. Estrogens, progesterone, normal breast cell proliferation, and breast cancer risk. *Epidemiol Rev* 1993;15:17-35.
- Whelan EA, Sandler DP, Root JL, Smith KR, Weinberg CR. Menstrual cycle patterns and risk of breast cancer. *Am J Epidemiol* 1994;140:1081-90.
- Frisch RE, Wyshak G, Albright NL, et al. Lower prevalence of breast cancer and cancers of the reproductive system among former college athletes compared to non athletes. *Br J Cancer* 1985;52:885-91.
- Vena JE, Graham S, Zielezny M, Brasure J, Swanson MK. Occupational exercise and risk of cancer. *Am J Clin Nutr* 1987;45:Suppl:318-27.

17. Albanes D, Blair AA, Taylor PR. Physical activity and risk of cancer in the NHANES I population. *Am J Public Health* 1989;79:744-50.
18. Zheng W, Shu XO, McLaughlin JK, Chow W-H, Gao YT, Blot WJ. Occupational physical activity and the incidence of cancer of the breast, corpus uteri, and ovary in Shanghai. *Cancer* 1993;71:3620-4.
19. Bernstein L, Henderson BE, Hanisch R, Sullivan-Halley J, Ross RK. Physical exercise and reduced risk of breast cancer in young women. *J Natl Cancer Inst* 1994;86:1403-8.
20. Friedenreich CM, Rohan TE. Physical activity and risk of breast cancer. *Eur J Cancer Prev* 1995;4:145-51.
21. Mittendorf R, Longnecker MP, Newcomb PA, et al. Strenuous physical activity in young adulthood and risk of breast cancer (United States). *Cancer Causes Control* 1995;6:347-53.
22. D'Avanzo B, Nanni O, La Vecchia C, et al. Physical activity and breast cancer risk. *Cancer Epidemiol Biomarkers Prev* 1996;5:155-60.
23. Welsh CW, DeHoog JV, O'Connor DH, Scheffield LG. Influence of dietary fat levels on development and hormone responsiveness of the mouse mammary gland. *Cancer Res* 1985;45:6147-54.
24. Boissonneault GA, Elson CE, Pariza MW. Net energy effects of dietary fat on chemically induced mammary carcinogenesis in F344 rats. *J Natl Cancer Inst* 1986;76:335-8.
25. Cohen LA, Choi KW, Wang C-X. Influence of dietary fat, caloric restriction, and voluntary exercise on *N*-nitrosomethylurea-induced mammary tumorigenesis in rats. *Cancer Res* 1988;48:4276-83.
26. Bjartveit K, Foss OP, Gjervig T, Lund-Larsen PG. The Cardiovascular Disease Study in Norwegian counties: background and organization. *Acta Med Scand Suppl* 1979;634:1-70.
27. The Cardiovascular Disease Study in Norwegian counties: results from second screening. Oslo, Norway: National Health Screening Services, 1988.
28. Løken EB, Solvoll K. The reproducibility of a self-administered diet questionnaire. *Var Foda* 1987;1:Suppl 1:33-7.
29. Gaard M, Sandvin O, Løken EB. Calculation of nutrient intake based on dietary data from 2nd cardio-vascular disease survey in Finnmark, Sogn og Fjordane, and Oppland 1977-83. Technical report 1. Oslo: Cancer Registry of Norway, 1996.
30. Paffenbarger RS Jr, Hyde RT, Wing AL. Physical activity and incidence of cancer in diverse populations: a preliminary report. *Am J Clin Nutr* 1987;45:Suppl:312-7.
31. Dorgan JF, Brown C, Barrett M, et al. Physical activity and risk of breast cancer in the Framingham Heart Study. *Am J Epidemiol* 1994;139:662-9.
32. Vihko VJ, Apter DL, Pukkala EL, Oinonen MT, Hakulinen TR, Vihko RK. Risk of breast cancer among female teachers of physical education and languages. *Acta Oncol* 1992;31:201-4.
33. Dosemeci M, Hayes RB, Vetter R, et al. Occupational physical activity, socioeconomic status, and risk of 15 cancer sites in Turkey. *Cancer Causes Control* 1993;4:313-21.
34. Wilhelmson L, Tibblin G, Aurell M, Bjure J, Ekstrom-Jodal B, Grimby G. Physical activity, physical fitness and risk of myocardial infarction. *Adv Cardiol* 1976;18:217-30.
35. Holme I, Helgeland A, Hjertmann I, Leren P, Lund-Larsen PG. Physical activity at work and at leisure in relation to coronary risk factors and social class: a 4-year mortality follow up: the Oslo Study. *Acta Med Scand* 1981;209:277-83.
36. Løchen M-L, Rasmussen K. The Tromsø study: physical fitness, self reported physical activity, and their relationship to other coronary risk factors. *J Epidemiol Community Health* 1992;46:103-7.
37. Kvåle G, Heuch I. Menstrual factors and breast cancer risk. *Cancer* 1988;62:1625-31.
38. Collaborative Group on Hormonal Factors in Breast Cancer. Breast cancer and hormonal contraceptives: collaborative reanalysis of individual data on 53 297 women with breast cancer and 100 239 women without breast cancer from 54 epidemiological studies. *Lancet* 1996;347:1713-27.
39. Pérusse L, Tremblay A, Leblanc C, Bouchard C. Genetic and environmental influences on level of habitual physical activity and exercise participation. *Am J Epidemiol* 1989;129:1012-22.
40. Slattery ML, McDonald A, Bild DE, et al. Associations of body fat and its distribution with dietary intake, physical activity, alcohol, and smoking in blacks and whites. *Am J Clin Nutr* 1992;55:943-9.
41. Broocks A, Pirke KM, Schweiger U, et al. Cyclic ovarian function in recreational athletes. *J Appl Physiol* 1990;68:2083-6.
42. Beitins IZ, McArthur JW, Turnbull BA, Skrinar GS, Bullen BA. Exercise induces two types of human luteal dysfunction: confirmation by urinary free progesterone. *J Endocrinol Metab* 1991;72:1350-8.
43. Russell JB, Mitchell D, Musey PJ, Collins DC. The relationship of exercise to anovulatory cycles in female athletes: hormonal and physical characteristics. *Obstet Gynecol* 1984;63:452-6.
44. Kritchevsky D. Caloric restriction and experimental carcinogenesis. In: Jacobs MM, ed. *Exercise, calories, fat, and cancer*. Vol. 322 of *Advances in experimental medicine and biology*. New York: Plenum Press, 1992:131-41.
45. Tretli S. Height and weight in relation to breast cancer morbidity and mortality: a prospective study of 570,000 women in Norway. *Int J Cancer* 1989;44:23-30.
46. Taioli E, Barone J, Wynder EL. A case-control study on breast cancer and body mass. *Eur J Cancer* 1995;31A:723-8.
47. Freni SC, Eberhardt MS, Turturro A, Hine RJ. Anthropometric measures and metabolic rate in association with risk of breast cancer (United States). *Cancer Causes Control* 1996;7:358-65.
48. Ziegler RG, Hoover RN, Nomura AMY, et al. Relative weight, weight change, height, and breast cancer risk in Asian-American women. *J Natl Cancer Inst* 1996;88:650-60.
49. Meyer F, Moisan J, Marcoux D, Bouchard C. Dietary and physical determinants of menarche. *Epidemiology* 1990;1:377-81.
50. Haffner SM, Katz MS, Dunn JF. Increased upper body and overall adiposity is associated with decreased sex hormone binding globulin in postmenopausal women. *Int J Obes* 1991;15:471-8.

Paper V

**PHYSICAL ACTIVITY IMPROVES THE METABOLIC RISK PROFILES
IN MEN AND WOMEN**

A seven-year follow-up study with repeated assessments of leisure time activity: The Tromsø Study

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Objective.—To examine effects of leisure time physical activity on the metabolic profiles.

Design.— Population-based cohort study, following subjects attending two surveys (1979–80 and 1986–87) with repeated assessments of self-reported leisure time physical activity

Participants.— 5220 men and 5869 women, aged 20–49 years at entry.

Main Outcome Measures.— Measurements of body mass index (BMI in kg/m^2), triglycerides, total cholesterol (Total-C), high-density-lipoprotein cholesterol (HDL-C) related to four levels of physical activity.

Results.— There was a dose-response relationship between serum lipids and BMI, and levels of physical activity in both sexes after adjustments for potential confounders. These differences of BMI and serum lipids between sedentary and sustained exercising groups were consistently more pronounced after 7 years than at baseline, especially in the oldest age group. Men reporting sustained hard training compared with sedentary men, had a lower concentration of Total-C (5.65 mmol/l vs 6.21 mmol/l), triglycerides (1.34 mmol/l vs 1.85 mmol/l), Total-C:HDL-C ratio by 19.0% and BMI (23.9 kg/m^2 vs 25.7 kg/m^2), and a higher HDL-C concentration (1.52 mmol/l vs 1.36 mmol/l). Women reporting sustained regular or hard training compared with sedentary women, had a lower concentration of Total-C (5.70 mmol/l vs 5.90 mmol/l), triglycerides (1.03 mmol/l vs 1.18 mmol/l), Total-C:HDL-C ratio by 7.5% and BMI (23.1 kg/m^2 vs 23.6 kg/m^2), and a higher HDL-C concentration (1.73 mmol/l vs 1.66 mmol/l). An increase in leisure time activity over the 7 years improved metabolic profiles, whereas a decrease worsened them in both sexes.

Conclusions.— Sustained high levels and change from sedentary to higher levels of physical activity improved the metabolic risk profile in both sexes. The differences observed are sufficiently large to have a beneficial effect on the risk of certain chronic diseases.

INTRODUCTION

Physical activity strengthens the musculoskeletal system, improves cardiovascular capacity and pulmonary function,¹ influences the cumulative exposure to certain hormones,² enhances the immune system³ and increases insulin sensitivity.⁴ It also represents the largest source of variability in energy requirements,⁵⁻⁶ influencing body weight⁶⁻⁷. The benefits of physical activity and fitness have been associated with decreased overall mortality⁸⁻¹⁰ and reduced incidence of cardiovascular diseases,¹¹⁻¹² non-insulin-dependent diabetes mellitus^{7,12-13} and certain types of cancer such as colon cancer¹⁴⁻¹⁵ and breast cancer.¹⁶

Serum lipids are important biological risk markers for chronic diseases, particularly cardiovascular diseases,^{11,17-18} that can be modified by physical activity.¹⁹⁻²⁴ In addition, obesity increases the risk of cardiovascular diseases,²⁵⁻²⁶ diabetes^{4,27} and certain types of cancer.^{14,28}

Changes in the level of physical activity can result in weight gain or weight loss if there is no calorie compensation. Weight gain in adult life may be a marker of a new metabolic steady state, which may have serious health consequences.^{4,26,28-29} This emphasizes the importance of age and body mass index (BMI) in the evaluation of the influence of physical activity on metabolic risk profiles.

Few population-based prospective studies have evaluated the importance of both sustained and change in level

of physical activity over time on weight gain and lipids, taking age into account. Most studies are of men, less is known about women.^{20,30-32}

The aim of the present study was to elucidate the impact of physical activity on the metabolic risk profile over a certain timespan. This was done in a population-based study of 5220 men and 5869 women with two self-reported assessments of leisure time physical activity with a 7-year interval. Biological markers of a metabolic risk profile were defined by BMI, triglycerides, total cholesterol, high-density-lipoprotein cholesterol (HDL-C) and the ratio of total cholesterol to HDL cholesterol (Total-C:HDL-C). Data on smoking habits, menopausal status, dietary factors, consumption of alcohol and coffee, and use of hormonal contraceptives allowed for adjustment of use and change in potentially confounding factors during follow-up. Heart rate and physical fitness (in women only), made a validation of self-reported physical activity possible.

MATERIAL AND METHODS

Study subjects

The study subjects are men and women who participated in two population surveys carried out with a 7-year interval in the municipality of Tromsø, northern Norway. In the 1979-80 survey, all men aged 20-54 years and all women aged 20-49 years, registered in the municipality, were invited and 16 621 subjects attended, i.e. 78% of the invited

population. The total number of individuals examined at the 1986-87 survey was 21 826 subjects, 81.3% of the eligible population. Eligible for the present study were those 11 508 subjects (5423 men and 6085 women) who were 20-49 years at baseline (1979-80) and attended both surveys (88.3% of those invited to both surveys). We excluded subjects who reported previous myocardial infarction, stroke, angina pectoris, diabetes mellitus and taking antihypertensive medications, (192 men, 207 women) at baseline (1979-80), and those with missing information about leisure time activity at either surveys (11 men, 9 women). Hence, the present cohort consisted of 5220 men and 5869 women aged 20-49 years in 1979-80.

Screening procedures

The methods and questionnaires used in the two surveys were almost identical and are described in detail elsewhere.³³ The screening comprised administration of a main questionnaire about disease, symptoms and smoking. At the two surveys, trained nurses checked the questionnaire for inconsistencies and asked about time since last meal, menopausal status, use of oral contraceptives and hormonal replacement therapy (1986-87). Height was measured to the nearest centimetre and weight to the nearest half-kilogram on regularly calibrated scales. As a measure of obesity, we used the body mass index (BMI) (weight (kg)/[height (m)]²).

In the 1979/80 survey, blood pressure was measured by personnel

trained according to tape recordings produced by the London School of Hygiene and Tropical Medicine. In the 1986-87 survey, an automatic device (Dinamap, Criticon, Tampa) was used. Heart rate is derived from the median pulse-to-pulse interval during the time of the blood pressure measurement. Three recordings of heart rate were made at 2-minute intervals, and the lowest measurement recorded was used in the present study.

A non-fasting blood sample for lipid analyses was taken. Total serum cholesterol was measured directly by the enzymatic oxidase method. HDL-cholesterol was assayed by the same procedure after precipitation of low-density lipoproteins with heparin and manganese chloride. Triglycerides were enzymatically determined as glycerol.

A second questionnaire, given to the participants at both surveys, was a combined food frequency questionnaire and a questionnaire about chronic diseases and drug use. The nutritional part covered type and quantity of table fat, milk, coffee drinking, vegetables, fruit and alcohol habits. It was filled in at home and returned by 88% and 92% of the participants at the 1979-80 and 1986-87 surveys, respectively. Energy or fat intake for each individual could not be calculated because of insufficient information about nutritional habits.

Assessment of physical activity and physical fitness

The main questionnaire covering physical activity was filled in at home

and checked at the screening for inconsistencies and incomplete data. Physical activity in leisure time was graded from I to IV: level I: reading, watching TV or other sedentary activities; level II: walking, bicycling or physical activities for at least 4 hours a week; level III: exercises to keep fit for at least 4 hours a week; and level IV: regular hard training or exercise for competition several times a week.

Physical fitness was assessed by a graded submaximal or maximal bicycle exercise test with a pedalling frequency of 60/min in a random subgroup in the 1986–87 survey.³⁴ The initial workload was set at 25 watts, with a 25-W increment every minute, up to a maximum of 250 W after 10 min. Physical fitness was defined as the maximum workload performed. As a result of incomplete testing in men – about 75% reached the maximum workload – only measurements of 220 women could be used in analyses.

Statistical analysis

All analyses were sex specific. The primary aim was to analyse differences and changes in metabolic risk factors (serum lipids and BMI) in leisure time sedentary and active attendees over the 7 years of follow-up. Baseline characteristics were age adjusted and compared, based on the four reported levels of leisure time activity using analysis of co-variance. In the analysis of changes in BMI, cholesterol, HDL-cholesterol, triglycerides and Total-C:HDL-C ratio, comparison groups were

defined as:

- ≤ -2 changes – those who reduced at least two levels of leisure time activity
- -1 change – those who reported one level reduction in leisure activity between surveys
- Unchanged – those who reported the same activity level after 7 years of follow-up
- $+1$ change – those who reported one level increase in activity
- $\geq +2$ change – those who increased at least two levels of leisure time activity.

We adjusted, for age at entry into the 1979–80 survey, use and change in smoking habits, coffee drinking, type of table fat used and menopausal status, when analysing the effect of sustained and change in physical activity in metabolic profiles, respectively. Time since last meal was included as a co-variate only in the analyses of triglyceride levels. Analysis of covariance were used for adjustments. Because very few women reported regular hard exercise (level IV), levels III and IV were merged in some analyses.

We examined models stratified by age at entry (20–29, 30–39 and 40–49 years) and BMI (tertiles) to analyse whether there was any effect modification by age and BMI of leisure time activity. As a result of missing data, the number of subjects included in the separate analyses varied slightly. Tests for linear trends were performed by linear regression. All significance tests were two-tailed

and the significance level was chosen at 5%. SAS statistical package version 6.11 was used.

RESULTS

At baseline the mean age was 34.4 years and 33.7 years in men and women, respectively. Sedentary leisure time activity was reported by 19.4% of men and 21.9% of women; regular exercise was reported in 29.3% of men and 12.2% of women;

hard exercise was performed by 6.3% of men and 1.0% of women (Table 1).

Men reporting higher levels of daily leisure time activity at baseline tended to be slightly leaner, to have an increased daily intake of fruit and vegetables and low fat milk, to have lower diastolic blood pressure, to smoke less, to consume fewer cups of coffee and to have a lower daily alcohol and saturated table fat intake compared with sedentary men (Table 1). Men who performed regular hard

Table 1. Baseline characteristics* (mean/distribution (%)) according to level of leisure time physical activity in men and women, aged 20-49 years at entry at survey 1979-80.

CHARACTERISTICS	LEISURE TIME PHYSICAL ACTIVITY								
	Sedentary (n=1015)†	Men				Women			
		Moderate (n=2344)	Regular (n=1532)	Hard (n=329)	Sedentary (n=1284)	Moderate (n=3811)	Regular (n=715)	Hard (n=59)	
Age at entry (years)	34.4	35.2	34.3	29.6	32.9	33.9	34.4	27.7	
BMI (kg/m ²)	24.5	24.2	24.1	24.0	22.6	22.7	22.4	22.3	
Height (cm)	176.6	177.1	177.2	177.2	163.5	163.6	164.5	165.1	
Systolic blood pressure (mmHg)	129.4	129.6	129.3	130.2	121.3	121.2	119.3	119.3	
Diastolic blood pressure (mmHg)	81.7	81.8	80.9	80.5	77.9	78.0	77.7	77.1	
Dietary intake (%)									
unsaturated table fat	71.3	72.2	74.4	75.8	71.8	75.3	75.4	67.1	
low fat milk	12.1	13.7	17.2	21.0	17.7	26.7	33.2	33.2	
fruit/vegetables (daily)	31.8	40.6	47.5	48.5	52.7	65.4	74.4	67.2	
Coffee consumption (% ≥5 cups/daily)	68.1	61.0	59.1	46.2	55.8	50.5	47.4	35.5	
Alcohol (% ≥ 2-3 times weekly)	17.8	13.7	13.6	11.7	4.4	3.9	4.8	6.6	
Daily smoking (%)	59.8	48.6	41.5	25.7	54.2	44.9	41.9	37.4	
Use of oral contraceptives (%)					8.4	7.3	7.0	11.1	
Premenopausal (%)					95.2	95.8	96.5	90.1	

* Age-adjusted

† Number of participants for activity categories in parentheses. For some subjects, information concerning certain variables was missing.

Table 2. Changes in serum lipids and body mass index (BMI) according to changes in the level of leisure time physical activity after 7 years of follow-up (1986-87)

Characteristics*	Baseline†	1986-87†	CHANGE IN LEVEL OF LEISURE TIME ACTIVITY					p for trend
			Mean ±SD	Mean ±SD	≤-2	-1	0	
			(n=261)‡	(n=1337)	(n=2681)	(n=823)	(n=118)	
MEN								
Cholesterol								
(mmol/l)	5.93±1.26	6.05 ± 1.23	0.32	0.14	0.12	0.05	0.09	0.0002
[mg/dl]	[229±49]	[234±48]	[12]	[5]	[5]	[2]	[3]	
Triglycerides								
(mmol/l)	1.63 ±0.96	1.67 ± 1.07	0.14	0.08	0.03	0.04	-0.16	0.14
[mg/dl]	[144±85]	[148±95]	[12]	[7]	[3]	[4]	[-14]	
HDL chol.								
(mmol/l)	1.45±0.45	1.37±0.35	-0.11	-0.09	-0.09	-0.07	0.01	0.16
[mg/dl]	[56±17]	[53±14]	[-3]	[-3]	[-3]	[-3]	[0.4]	
Tot.-C:HDL-C	4.36 ±1.46	4.71± 1.63	0.59	0.40	0.35	0.22	0.02	0.0001
BMI (kg/m ²)	24.2 ±2.8	24.9 ± 2.9	1.0	0.8	0.7	0.6	0.2	0.0001
			(n=134)	(n=1147)	(n=3604)	(n=917)	(n= 67)	
WOMEN								
Cholesterol								
(mmol/l)	5.73 ±1.19	5.83 ± 1.26	0.30	0.15	0.09	0.04	-0.08	0.004
[mg/dl]	[222±46]	[225±49]	[12]	[6]	[3]	[2]	[-3]	
Triglycerides								
(mmol/l)	1.09 ±0.61	1.15 ± 0.66	0.13	0.06	0.06	0.01	0.03	0.14
[mg/dl]	[97±54]	[102±58]	[12]	[5]	[5]	[1]	[3]	
HDL chol.								
(mmol/l)	1.75 ±0.42	1.66 ± 0.39	-0.10	-0.10	-0.09	-0.07	-0.05	0.05
[mg/dl]	[68±16]	[64±15]	[-4]	[-4]	[-3]	[-3]	[-2]	
Tot.-C:HDL-C	3.43 ±1.03	3.69 ± 1.19	0.35	0.31	0.26	0.19	0.04	0.0013
BMI (kg/m ²)	22.6 ±3.2	23.5 ± 3.5	1.2	1.0	0.9	0.7	0.4	0.001

* Adjusted for age at baseline and change in: smoking habits, table fat, coffee drinking, menopausal status (women)

† Values are presented as means ± SD

‡ Number of participants for activity categories in parentheses. For some subjects, information concerning certain variables was missing.

exercise tended to be younger. Leisure time active women tended to be taller, to have a lower systolic blood pressure and a higher daily intake of low fat milk, to consume fewer cups of coffee and to smoke less than sedentary women. Only 7.5% of the women were current users of oral contraceptives

After 7 years, more individuals reported decreased (men: 30.6%, women: 21.8%) rather than increased physical activity (men: 18.0%, women: 16.8%). There was a decline

in level of physical activity with increasing age. By comparing the regular exercise groups at follow-up survey, with the groups who are 10 years older at baseline, there was, however, a decline by 3-5% in subjects who trained regularly; this was not related to the average age effect in the cohort (results not presented).

There was an overall increase in BMI, triglycerides, cholesterol and Total C:HDL-C ratio, and a decrease in HDL-C in both sexes after 7 years

in the cohort relative to baseline values (Table 2). After adjustment for age at entry there was a highly significant inverse dose-response pattern from most reduced to most increased changes in activity level in both sexes for serum cholesterol concentration, Total-C: HDL-C ratio and BMI. Adjustment for change in smoking habits, dietary fat intake, coffee drinking and menopausal status did not change these results. Comparing the two extremes in the change of physical activity level during follow-up (≤ 2 levels decrease vs ≥ 2 levels increase), there was a significant difference also in triglycerides and HDL-C for both sexes. Further adjustments for present use of or change in fruit/vegetable, use of oral contraceptives or alcohol consumption did not influence this association and were omitted from the final model.

To analyse the influence of sustained leisure activity on metabolic risk factors over time, we focused on men and women who maintained their activity after 7 years of follow-up. After multivariate adjustments the differences in the level of serum lipids and BMI, between the sedentary and exercising groups (regular and hard), were consistently more pronounced after 7 years than at baseline (Table 3). This was especially marked among those who sustained activity compared with all participants. This was true for both men and women. Men reporting sustained hard training had reduced levels, compared with sustained sedentary men, of: Total-C by 9.0%, triglycerides by 27.6% and

Total-C:HDL-C ratio by 19.0%; there was also an increase in HDL-C of 13.2% plus a 7.0% smaller BMI. Women who were regular and hard trainers in their leisure time had a reduced concentration, compared with sedentary women, of: Total-C by 3.4%, triglycerides by 12.7% and Total-C:HDL-C ratio by 7.5%; they also had an increase in HDL-C of 4.0% plus a 2.1% smaller BMI. All tests for linear trend with increasing physical activity showed a highly significant dose-response effect ($p < 0.001$) for men and for women (except HDL-C).

The effect of sustained physical activity on metabolic profiles in different age groups of men is demonstrated in Figure 1. First, the differences in BMI, and the levels of cholesterol, triglycerides, HDL-C and Total-C: HDL-C, when compared for the sedentary and hard exercise group, were more pronounced in the oldest (40-49 years) than in the youngest (20-29 years) age groups; this effect increased after 7 years of follow-up. Second, the average levels of the metabolic profiles were consistently higher after 7 years in the sedentary and moderately active individuals of all ages. It is possible to reduce the levels of BMI, cholesterol and triglycerides below the baseline level by regular and hard training after 7 years, but this occurs mainly in the oldest segment of the cohort. Similar effects were seen for women, although to a lesser extent (Figure 2). This indicates an important effect of sustained physical activity on metabolic profiles.

A possible effect modification of weight was elucidated by stratified analyses of BMI. Leisure time activity had the same effect on the serum lipids in all three tertiles of BMI in both sexes (Figure 3). Within each tertile of BMI, the serum concentration of Total-C and

Table 3. Serum lipids* and BMI* at baseline among all participants and after 7 years of follow-up among men and women who sustained† activity across levels of leisure time physical activity

Characteristics	LEISURE TIME PHYSICAL ACTIVITY									
	All participants survey 1979/80					Sustained activity survey 1986/87				
	Sedentary	Moderate	Regular	Hard	p for trend	Sedentary	Moderate	Regular	Hard	p for trend
MEN	(n=1015)§	(n=2344)	(n=1532)	(n=329)		(n=499)	(n=1485)	(n=616)	(n=81)	
Cholesterol										
(mmol/l)	6.09	5.93	5.90	5.69	0.0001	6.21	6.13	5.96	5.65	0.001
[mg/dl]	[235]	[229]	[228]	[220]		[240]	[237]	[230]	[218]	
Triglycerides										
(mmol/l)	1.73	1.62	1.59	1.48	0.0001	1.85	1.69	1.54	1.34	0.0001
[mg/dl]	[153]	[143]	[141]	[131]		[164]	[150]	[119]	[136]	
HDL chol.										
(mmol/l)	1.43	1.45	1.46	1.52	0.007	1.36	1.36	1.39	1.52	0.0014
[mg/dl]	[55]	[56]	[56]	[59]		[53]	[53]	[54]	[59]	
Tot.C:HDL-C	4.58	4.38	4.28	3.95	0.0001	4.84	4.80	4.56	3.92	0.0001
BMI (kg/m²)	24.5	24.2	24.1	23.9	0.0001	25.7	25.0	24.5	23.9	0.0001
WOMEN‡	Sedentary	Moderate	Regular/Hard			Sedentary	Moderate	Regular/Hard		
	(n=1284)§	(n=3811)	(n=774)			(n=581)	(n=2832)	(n=213)		
Cholesterol										
(mmol/l)	5.74	5.74	5.62	0.07	5.90	5.85	5.70	0.05		
[mg/dl]	[222]	[222]	[217]		[228]	[226]	[220]			
Triglycerides										
(mmol/l)	1.15	1.09	1.01	0.03	1.18	1.14	1.03	0.007		
[mg/dl]	[102]	[97]	[89]		[105]	[101]	[91]			
HDL chol.										
(mmol/l)	1.73	1.75	1.79	0.22	1.66	1.67	1.73	0.09		
[mg/dl]	[67]	[68]	[69]		[64]	[65]	[67]			
Tot.C:HDL-C	3.47	3.44	3.30	0.07	3.75	3.69	3.47	0.009		
BMI (kg/m²)	22.6	22.7	22.3	0.14	23.6	23.5	23.1	0.03		

* Adjustments were done for age at baseline and current; smoking habits, coffee drinking, table fat, menopausal status (women) and time since last meal (triglycerides) at 1979-80 or 1986-87 survey, respectively.

† Sustained activity level; men and women who reported the same level of leisure time activity in 1979-80 and in 1986-87.

‡ Regular and hard exercise are combined in women

§ For some subjects, information concerning certain variables was missing.

Table 4. Mean values of heart rate (beats/min)* at survey 1986/87 in both sexes by level and changes of leisure time physical activity at survey 1979/80 and 1986/87 combined.

Change in leisure time physical activity 1979/80 to 1986/87	Leisure time physical activity 1986/87				<i>p</i> for trend
	Sedentary	Moderate	Regular exercise	Hard exercise	
MEN					
Increased (<i>n</i> = 941)	-	71.5	68.4	63.3	<i>p</i> < 0.0001
Sustained (<i>n</i> = 2 676)	72.8	71.6	66.2	56.8	<i>p</i> < 0.0001
Decreased (<i>n</i> = 1 598)	72.8	69.1	62.6	-	<i>p</i> < 0.0001
WOMEN					
Increased (<i>n</i> = 984)	-	75.5	72.5	69.4	<i>p</i> < 0.0001
Sustained (<i>n</i> = 3 604)	77.6	75.0	71.1	69.4	<i>p</i> < 0.0001
Decreased (<i>n</i> = 1 281)	75.8	73.3	70.0	-	<i>p</i> = 0.0002

* Adjusted for age at baseline (1979-80)

Table 5. Physical fitness and heart rate in relation to leisure time physical activity among 220 women at survey 1986/87.

Leisure time physical activity	Number	Physical fitness (Watts) \pm SEM	Heart rate (beats/min) \pm SEM
Sedentary	55	144.1 \pm 4.59	78.2 \pm 1.74
Moderate	148	158.7 \pm 2.80	75.4 \pm 1.06
Regular/Hard exercise	17	186.1 \pm 8.25	69.0 \pm 3.13
<i>p</i> for trend		<i>p</i> < 0.0001	<i>p</i> = 0.02

Values are presented as age-adjusted means \pm SEM

triglycerides, and the Total-C:HDL-C ratio were reduced and HDL-C was increased with increasing leisure time activity.

The impact of physical activity on metabolic profiles in non-smokers did not differ from the results presented

for smokers and non-smokers combined (results not presented).

A highly significant lower heart rate with increasing physical activity level was demonstrated (Table 4) for both sexes in all groups (*p* for linear trend < 0.0001). When comparing

those who increased or decreased their activity level with those who sustained their activity level between the two surveys, a lower heart rate among those who increased and a higher heart rate among those who decreased their activity level was found. The lowest heart rate was observed among those who reported the highest level of leisure time physical activity in both surveys in both sexes, indicating high correlation between physical activity and heart rate.

Physical fitness, assessed in a small random sample of 220 women, increased with increasing level of leisure time activity ($p < 0.0001$) (Table 5), and heart rate was also reduced with increasing level of leisure activity ($p = 0.02$).

DISCUSSION

With the steady decline in occupational physical activity, exercise at leisure time has become more important for determining metabolic and health-related effects of physical activity. About 36% of the men and 13% of the women in our study performed regular exercise at baseline, which dropped to 26% and 9%. This implies both an age-dependent decline in level of physical activity in the cohort, and a general decrease in activity in the population, independent of age.

The large sample size, and repeated assessment of physical activity, made

it possible to test for the influence of both sustained and change in physical activity on weight and serum lipids in an adult population. Sustained physical activity at leisure time reduced the age-related weight gain and improved lipid profiles across all BMI strata. Maintenance of a high level of physical activity after 7 years strengthened the associations observed at baseline. Men and women reporting sustained regular or hard training had significant improvement of serum lipids and BMI, compared with sedentary men and women. We were also able to demonstrate that an increase in leisure time activity over the 7 years improved the metabolic profiles, whereas a decrease in activity worsened the metabolic profile in both sexes.

Self-reports of physical activity provide a possibility for under- or over-estimating activity levels.³⁵ A lower heart rate and higher physical fitness (women), with increasing physical activity, support real differences in the level of physical activity among groups. Additionally, the physical activity assessment used has previously been validated,^{34, 36-37} and exclusion of subjects with chronic diseases improves the quality of the physical activity data in the study.

An individual's propensity to be physically active may be inherited.³⁸⁻³⁹ Lower Total-C and higher HDL-C concentrations at baseline may be markers for men and women who are genetically endowed with muscle

Fig 1. Age-adjusted mean values of body mass index (BMI) and serum lipids* across levels of leisure time physical activity at 1979-80 ●—● and 1986-87 ○---○ in different age-groups among men who sustained the same activity level at both surveys.
 *Triglycerides were adjusted for time since last meal

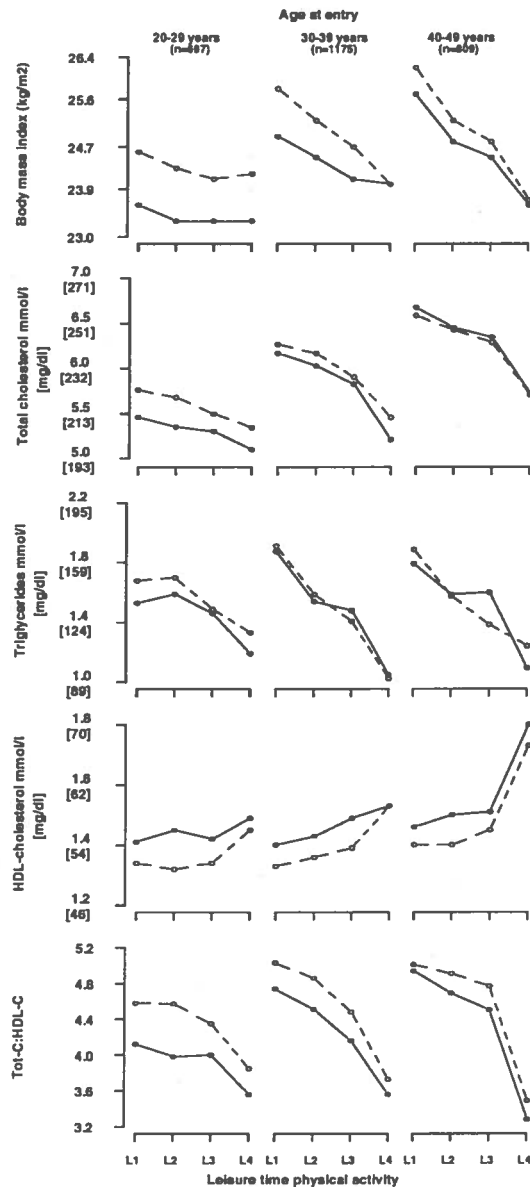


Fig 2. Age-adjusted mean values of body mass index (BMI) and serum lipids* across levels of leisure time physical activity at 1979-80 ●—● and 1986-87 ○---○ in different age-groups among women who sustained the same activity level at both surveys.
 *Triglycerides were adjusted for time since last meal

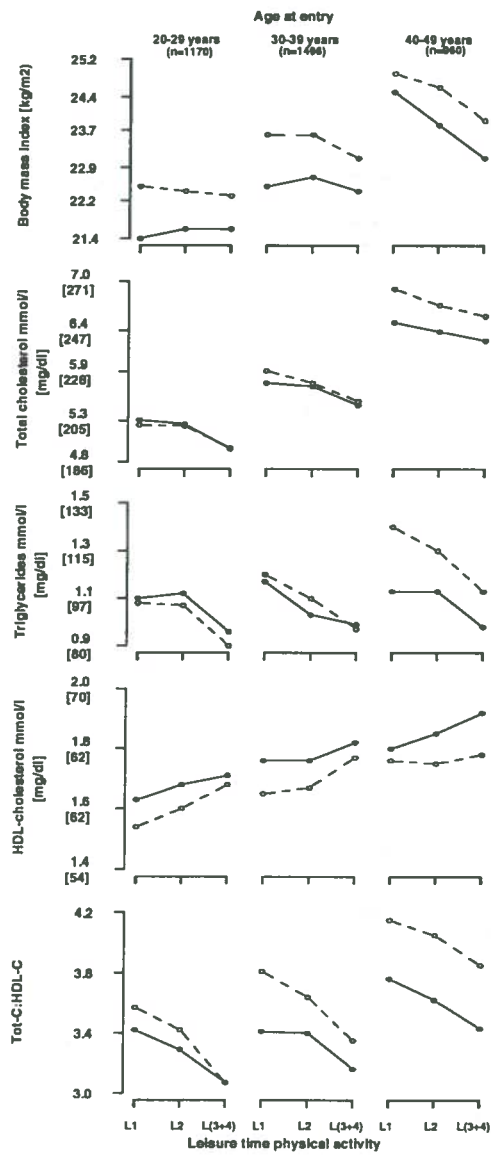
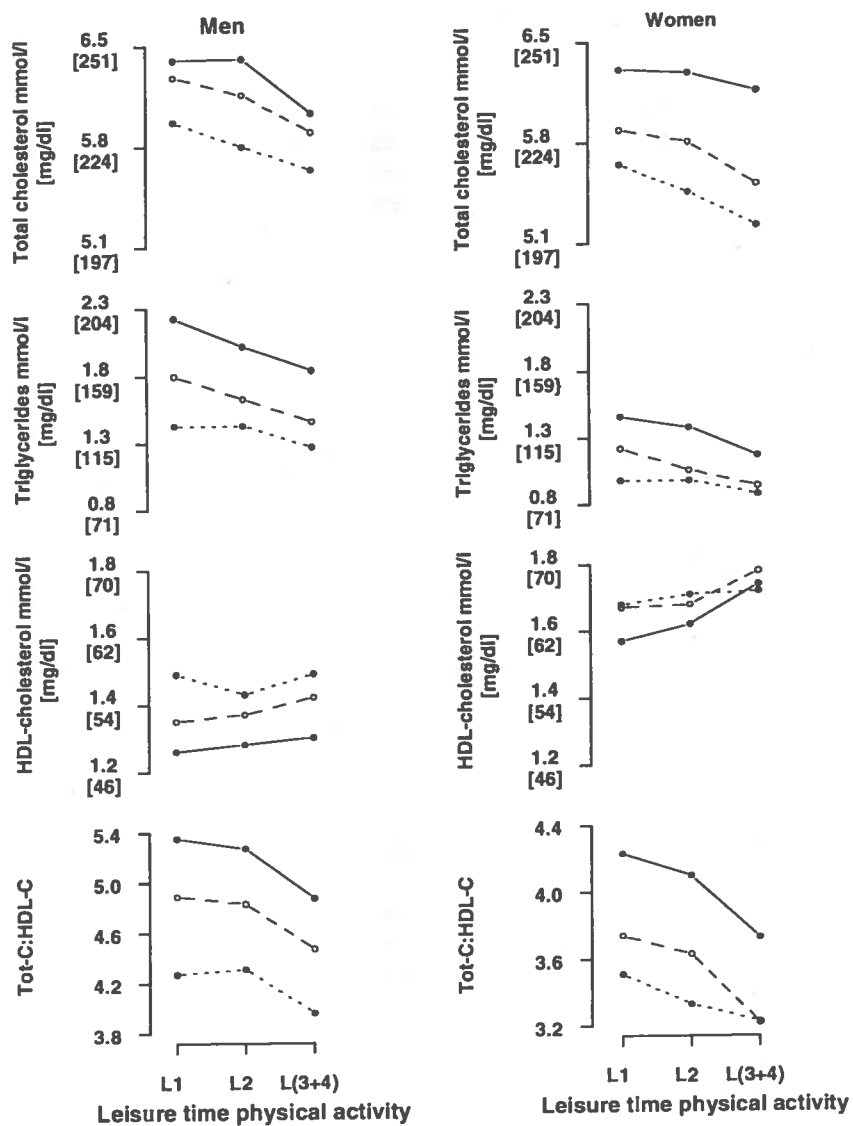


Fig. 3. Age-adjusted mean values of serum lipids* across levels of leisure time physical activity stratified by tertiles of body mass index (1986-87) among men; BMI < 23.5 kg/m² ●---●, BMI 23.5-25.9 kg/m² ○---○, BMI > 25.9 kg/m² ●---● and women; BMI < 21.7 kg/m² ●---●, BMI 21.7-24.2 kg/m² ○---○, BMI > 24.2 kg/m² ●---●, who sustained the same activity level at both surveys
*Triglycerides were adjusted for time since last meal



fibre types that make physical activity easier. Changes in the level of physical activity, however, influenced both body weight and lipid profiles in this study, and has been observed by others.⁴⁰ These changes indicate true metabolic effects of physical activity, not merely a genetic predisposition.

The population-based approach and high attendance rate reduce any selection bias in this study. It could be argued that the improvement of metabolic profiles could result from a residual effect among smokers. However, in separate analyses of non-smokers, the impact of physical activity on the estimates and tests for linear trends of the metabolic factors studied was no different from that in the total cohort.

It is possible that physically active subjects in this study may have under- or over-estimated the amount of certain foods consumed to a greater extent than inactive subjects. Due to incomplete information about nutritional habits, total energy or fat intake for each individual could not be calculated. Only certain items could be used as markers for foods generally consumed. Thus, a residual effect of diet composition, a cluster of healthy habits can not be excluded. However, regular exercise in overweight men and women improves the plasma lipoprotein levels in the present study and is additional to the effect of nutrition.⁴¹

Weight

Physical activity is an important component of long-term weight control. The inverse association

between physical activity and body weight at baseline corroborates previous cross-sectionally observations.¹⁴⁻¹⁵ The increase in BMI between sedentary and exercising groups, in those who maintained their activity level, is particularly interesting. This finding is consistent with a recent prospective study,⁶ in which American male adults in the lowest category of leisure activity weighed, on average, 1.1 kg/m² more than men in the highest category. In our study we observed that sedentary men weighed, on average, 1.8 kg/m² more than exercising men at the follow-up survey. This difference was most marked in the oldest age group, with 2.5 kg/m² low vs high physical activity; it supports the finding that inactivity results in more weight gain in older than in younger individuals.⁷ Even in those who sustained hard training, some weight gain was observed after 7 years of follow-up. This "age effect" could be explained by an age-dependent metabolic change in calorie utilization or a reduction in physical activity within, but not between, each level of activity. It could also be related to changes in nutrition.

It has been suggested that leisure time activity may also be a consequence of weight change.⁶ People can put on weight because their weight makes it more difficult to exercise. Women who decreased their physical activity over the follow-up period weighed, on average, 1.2 kg/m² more after 7 years, whereas those women who

increased their activity level had only a 0.4 kg/m² weight gain. Comparable results were observed for men; this points to the possibility of changing physical activity behaviour during adult life, independent of weight. Our study demonstrates a strong relationship between physical activity and weight gain in adult life, and supports the hypothesis that physical activity is a main determinant of body weight throughout adulthood. The effects of physical activity on body weight are mediated through the mechanisms of direct energy expenditure during exercise, preserving fat-free mass, increasing resting metabolic rate⁴²⁻⁴³ and the thermic effect of food,⁴² and inducing a decreased dietary intake.⁴⁴ Even if a person has not gained much weight, the mass of adipose tissue may have increased with a concomitant reduction in muscle mass as a result of physical inactivity.

Serum lipids

Cross-sectional and interventional studies have found lower concentrations of Total-C and triglycerides^{19, 21, 40} and higher concentrations of HDL-C^{24, 34, 45} in physically active compared with inactive individuals. We found more pronounced differences in the concentrations of cholesterol, triglycerides and HDL-C, and Total-C:HDL-C between sedentary and exercising men and women at follow-up; we are not aware of any other population-based study that demonstrates this effect on lipid profiles using repeated assessment of

physical activity. These findings indicate an important effect of sustained physical activity on lipid profiles in the general population. This effect was most pronounced among the oldest members of the cohort.^{21, 40, 46}

A certain intensity and duration of physical activity are necessary to achieve the desired effects on lipid metabolism.^{23, 47} Regular exercise for at least 4 hours a week, as in the present study, may include both the intensity and timespan needed to improve certain metabolic profiles. We found an inverse dose-response relationship between both sustained and increased physical activity levels and total cholesterol during follow-up, indicating that even moderate physical activity influences the level of total cholesterol. This was not the case with HDL-C and triglycerides because no significant dose-response relationship was observed between increase in the level of physical activity over 7 years and these lipids in men. By comparing the extremes in changes of activity (reduction versus increased), the HDL-C concentration was improved significantly in both men and women. This indicates that improvement in levels of triglycerides and HDL-C requires a high level of physical activity. This may also explain the weaker association of serum lipids and physical activity in population-based studies,²¹ because these studies may have had a smaller distribution of activity than intervention studies.^{24, 31, 40}

High BMI values have often been

associated with higher levels of serum cholesterol and triglycerides,^{21, 48} as was observed in this study. Differences in body weight are therefore frequently cited as the reason for differences in serum cholesterol, triglyceride and HDL levels between physically active and inactive people.^{41, 49} However, in the present study we observed an effect of physical activity on lipid profiles across all BMI values. This supports an influence of physical activity on lipid profiles independent of the metabolic effect of weight gain or loss.⁵⁰⁻⁵²

The increase in aerobic metabolism, oxygen uptake and increase in fatty acid use for muscular energy provides significant control parameters for lipid metabolic processes that occur during training.^{19, 22, 45, 47} These mechanisms may explain why physical activity is able to bring about a reduction in concentration of serum triglycerides and serum cholesterol, and an increase in serum HDL-C.^{22, 45, 47} It also supports the assumption that exercise over a long period (common in leisure time activity) has more impact than exercise over a short period – static exercise (common in occupational activities) – on lipid profiles;⁵³ this underlines the importance of leisure time activity over work activity.

Sex

It has been suggested that women have different physiological responses to physical activity than men in terms of BMI⁵⁴ and lipid

profiles.^{20, 31} We observed almost identical effects on BMI across all physical activity levels and in those who sustained their activity level during the follow-up. This supports common physiological responses to weight of physical activity in both sexes.

Our data also contradict the assumption that the generally higher HDL-C concentrations in women compared with men limit the potential for any further increase with exercise.²⁰ It has been observed that there is a smaller increase in HDL-C concentration in exercising women relative to men in some studies,²⁰ but not in others.²⁴ This may result from a smaller distribution in the level of physical activity in women and may not be related to true differences in biological effect between the sexes. The larger difference observed in heart rate according to level of leisure time activity in men than in women further emphasizes the larger range of physical activity carried out by men. Similarly, in this study, women, particularly in the oldest age group, reported hard exercise less frequently than men. This may also explain the smaller effect of physical activity on HDL-C concentration in women. However, women have to perform comparable levels of physical activity at a comparable intensity to achieve the same improvements as men in the concentrations of lipids.

Implications for chronic diseases

Not only are large weight gains during adulthood associated with

increased morbidity or mortality of chronic diseases;^{55, 56} even a modest weight gain in a normal weighted population is associated with increased risk of coronary heart disease,²⁹ non-insulin-dependent diabetes mellitus^{4, 7} and breast cancer.²⁸ The importance of the dose-response association between recent weight gain and breast cancer risk²⁸ underlines the importance of adult weight control in women. The improvement of lipid profiles by 9.0–27.6%, as achieved in men who carried out sustained hard training, may represent, from previous studies,^{57, 58} a reduction in morbidity and mortality from coronary heart disease of 20% and 25%, respectively. The reduction in triglyceride levels may be important as a risk factor for cardiovascular diseases¹⁷ and breast cancer.⁵⁹

The benefits of physical activity to risk and mortality from chronic diseases^{7-8, 10-12} may be hypothesized as acting through a common link. Weight gain during adulthood and physical inactivity^{4, 7, 54} may give a diminished sensitivity to insulin, which is associated with increased risk for non-insulin-dependent diabetes mellitus and cardiovascular diseases;⁶⁰ they have recently also been suggested to be important in carcinogenesis of the colon¹⁴ and breast.⁶¹

Conclusion

This population-based prospective study has demonstrated that sustained physical activity over 7 years reduces the age-related weight

gain, in a dose-response pattern, and improves lipid profiles across all BMI values in both sexes. A change from sedentary to higher levels of physical activity during adulthood improves metabolic profiles whereas a reduction in activity worsens the profiles. However, only sustained regular or hard exercise gives metabolic effects that are sufficient to compensate for the age-dependent worsening of metabolic profiles. Women have similar metabolic effects to men, but, as a result of a narrower distribution in the level of physical activity, there are more limited effects on lipids and BMI. These observations strengthen the importance of leisure time physical activity as a preventive factor against certain chronic diseases.

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REFERENCES

1. Higgins M, Keller JB, Wagenknecht LE, et al. Pulmonary function and cardiovascular risk factor relationships in black and in white young men and women. *Chest*. 1991;99:315-322.
2. Pike MC, Spicer DV, Dahmouch L, Press MF. Estrogens,

- progestogens, normal breast cell proliferation, and breast cancer risk. *Epidemiol Rev.* 1993;15:17-35.
3. Shinkai S, Kohno H, Kimura K, et al. Physical activity and immune senescence in men. *Med Sci Sports Exerc.* 1995;27:1516-1526.
 4. Folsom AR, Jacobs DR, Wagenknecht LE, et al. Increase in fasting insulin and glucose over seven years with increasing weight and inactivity of young adults. *Am J Epidemiol.* 1996;144:235-246.
 5. Danforth E. Diet and obesity. *Am J Clin Nutr.* 1985;41:1132-1145.
 6. Williamson DF, Madans J, Anda RF, Kleinman JC, Kahn HS, Byers T. Recreational physical activity and ten-year weight change in a US national cohort. *Int J Obes.* 1993;17:279-286.
 7. Helmrich SP, Ragland DR, Leung RW, Paffenbarger RS Jr. Physical activity and reduced occurrence of non-insulin-dependent diabetes mellitus. *N Engl J Med.* 1991;325:147-152.
 8. Paffenbarger RS Jr, Hyde RT, Wing AL, Lee I-M, Jung DL, Kampert JB. The association of changes in physical activity level and other lifestyle characteristics with mortality among men. *N Engl J Med.* 1993;328:538-545.
 9. Lee I-M, Hsieh C-C, Paffenbarger RS Jr. Exercise intensity and longevity in men. The Harvard alumni health study. *JAMA.* 1995;273:1179-1184.
 10. Lissner L, Bengtsson C, Björkelund C, Wedel H. Physical activity levels and changes in relation to longevity. *Am J Epidemiol.* 1996;143:54-62.
 11. Berlin JA, Colditz GA. A meta-analysis of physical activity in the prevention of coronary heart disease. *Am J Epidemiol.* 1990;132:612-628.
 12. Kujala UM, Kaprio J, Taimela S, Sarna S. Prevalence of diabetes, hypertension, and ischemic heart disease in former elite athletes. *Metabolism.* 1994;43:1255-1260.
 13. Burchfiel CM, Sharp DS, Curb JD, et al. Physical activity and incidence of diabetes: The Honolulu heart program. *Am J Epidemiol.* 1995;141:360-368.
 14. Giovannucci E, Ascherio A, Rimm EB, Colditz GA, Stampfer MJ, Willett WC. Physical activity, obesity, and risk for colon cancer and adenoma in men. *Ann Intern Med.* 1995;122:327-334.
 15. Thune I, Lund E. Physical activity and risk of colorectal cancer in men and women. *Br J Cancer.* 1996;73:1134-1140.
 16. Bernstein L, Henderson BE, Hanisch R, Sullivan-Halley J, Ross RK. Physical exercise and reduced risk of breast cancer in young

- women. *J Natl Cancer Inst.* 1994;86:1403-1408.
17. Stampfer MJ, Krauss RM, Ma J, et al. A prospective study of triglycerides level, low-density lipoprotein particle diameter, and risk of myocardial infarction. *JAMA.* 1996;276:882-888.
 18. Njølstad I, Arnesen E, Lund-Larsen PG. Smoking, serum lipids, blood pressure, and sex differences in myocardial infarction. A 12-year follow-up of the Finnmark Study. *Circulation.* 1996;93:450-456.
 19. Kiens B, Lithell H. Lipoprotein metabolism influenced by training-induced changes in human skeletal muscle. *J Clin Invest.* 1989; 83:558-564.
 20. Lokey EA, Tran ZV. Effects of exercise training on serum lipid and lipoprotein concentrations in women: A meta-analysis. *Int J Sports Med.* 1989;10:424-429.
 21. Lakka TA, Salonen JT. Physical activity and serum lipids: A cross-sectional population study in eastern Finnish men. *Am J Epidemiol.* 1992;136:806-818.
 22. Podl TR, Zmuda JM, Yurgalevitch SM, et al. Lipoprotein lipase activity and plasma triglyceride clearance are elevated in endurance-trained women. *Metabolism.* 1994;43:808-813.
 23. Marrugat J, Elosua R, Covas M-I, Molina L, Rubies-Prat J and the MARATHOM investigators. Amount and intensity of physical activity, physical fitness, and serum lipids in men. *Am J Epidemiol.* 1996;143:562-569.
 24. Williams PT. High-density lipoprotein cholesterol and other risk factors for coronary heart disease in female runners. *N Engl J Med.* 1996;334:1298-1303.
 25. Manson JE, Colditz GA, Stampfer MJ, et al. A prospective study of obesity and risk of coronary heart disease in women. *N Engl J Med.* 1990;322:882-889.
 26. Rimm EB, Stampfer MJ, Giovannucci E, et al. Body size and fat distribution as predictors of coronary heart disease among middle-aged and older US men. *Am J Epidemiol.* 1995;141:1117-1127.
 27. Colditz GA, Willett WC, Stampfer MJ, et al. Weight as a risk factor for clinical diabetes in women. *Am J Epidemiol.* 1990;132:501-513.
 28. Ziegler RG, Hoover RN, Nomura AM, et al. Relative weight, weight change, height, and breast cancer risk in Asian-American women. *J Natl Cancer Inst.* 1996;88:650-660.
 29. Willett WC, Manson JE, Stampfer MJ, et al. Weight, weight change and coronary heart disease in women: risk within the "normal"

- weight range. *JAMA*. 1995;273:461-465.
30. Eaton CB, Lapane KL, Garber CA, Assaf AR, Lasater TM, Carleton RA. Sedentary lifestyle and risk of coronary heart disease in women. *Med Sci Sports Exerc*. 1995;27:1535-1539.
31. Brownell KD, Bachorik PS, Ayerle RS. Changes in plasma lipid and lipoprotein levels in men and women after a program of moderate exercise. *Circulation*. 1982;65:477-484.
32. Suter E, Marti B. Little effect of long-term, self-monitored exercise on serum lipid levels in middle aged women. *J Sports Med Phys Fitness*. 1992;32:400-411.
33. Bønaa KH, Arnesen E. Association between heart rate and atherogenic lipid fraction in a population. The Tromsø Study. *Circulation*. 1992;86:394-405.
34. Løchen M-L, Rasmussen K. The Tromsø study: physical fitness, self reported physical activity, and their relationship to other coronary risk factors. *J Epidemiol Community Health*. 1992;26:103-107.
35. Klesges RC, Eck LH, Mellon MW, Fulliton W, Somes GW, Hanson CL. The accuracy of self-reports of physical activity. *Med Sci Sports Exerc*. 1990;22:690-697.
36. Saltin B, Grimby G. Physiological analysis of middle-aged and old former athletes. *Circulation*. 1968;38:1104-1115.
37. Wilhelmsen L, Tibblin G, Aurell M, Bjure J, Ekström-Jodal B, Grimby G. Physical activity, physical fitness and risk of myocardial infarction. *Adv Cardiol*. 1976;18:217-230.
38. Pérusse L, Tremblay A, Leblanc C, Bouchard C. Genetic and environmental influences on level of habitual physical activity and exercise participation. *Am J Epidemiol*. 1989;129:1012-1022.
39. Bouchard C, Lesage R, Lortie G, et al. Aerobic performance in brothers, dizygotic and monozygotic twins. *Med Sci Sports Exerc*. 1986;18:639-646.
40. Hellénus M-L, de Faire U, Berglund B, Hamsten A, Krakau I. Diet and exercise are equally effective in reducing risk for cardiovascular disease. Results of a randomized controlled study in men with slightly to moderately raised cardiovascular risk factors. *Atherosclerosis*. 1993;103:81-91.
41. Wood PD, Stefanick ML, Williams PT, Haskell WL. The effects on plasma lipoproteins of a prudent weight-reducing diet, with or without exercise in overweight men and women. *N Engl J Med*. 1991;325:461-466.

42. Poehlman ET, Melby CL, Badylak SF. Resting metabolic rate and postprandial thermogenesis in highly trained and untrained males. *Am J Clin Nutr.* 1988;47:793-798.
43. Bullough RC, Gillette CA, Harris MA, Melby CL. Interaction of acute changes in exercise energy expenditure and energy intake on resting metabolic rate. *Am J Clin Nutr.* 1995;61:473-481.
44. Lichtman SW, Pisarska K, Berman ER, et al. Discrepancy between self-reported and actual caloric intake and exercise in obese subjects. *N Engl J Med.* 1992;327:1893-1898.
45. Hartung GH, Lally DA, Prins J, Goebert DA. Relation of high-density lipoprotein cholesterol to physical activity levels in men and women. *Med Exerc Nutr Health.* 1992;1:293-300.
46. Stevenson ET, Davy KP, Seals DR. Hemostatic, metabolic and androgenic risk factors for coronary heart disease in physically active and less active postmenopausal women. *Arterioscler-Thromb-Vasc-Biol.* 1995;15:669-677.
47. Berg A, Frey I, Baumstark MW, Halle M, Keul J. Physical activity and lipoprotein lipid disorders. *Sports Med.* 1994;17:6-21.
48. Anderson AJ, Sobocinski KA, Freedman DS, Barboriak JJ, Rimm AA, Gruchow HW. Body fat distribution, plasma lipids, and lipoproteins. *Arteriosclerosis.* 1988;8:88-94.
49. Thompson PD, Culliname EM, Sady SP, et al. High density lipoprotein metabolism in endurance athletes and sedentary men. *Circulation.* 1991;84:140-152.
50. Higuchi M, Hashimoto I, Yamakawa K, Tsuji E, Nishimuta M, Suzuki S. Effect of exercise training on plasma high-density lipoprotein cholesterol level at constant weight. *Clin Physiol.* 1984;4:125-133.
51. Sopko G, Leon AS, Jacobs DR, et al. The effects of exercise and weight loss on plasma lipids in young obese men. *Metabolism.* 1985;34:227-236.
52. Tremblay A, Despres J-P, Maheux J, et al. Normalization of the metabolic profile in obese women by exercise and a low fat diet. *Med Sci Sports Exerc.* 1991;23:1326-1331.
53. Clarkson PM, Hintermister R, Fillyaw M, Stylos L. High density lipoprotein cholesterol in young adult weight lifters, runners, and untrained subjects. *Human biology.* 1981;53:251-257.
54. Krotkiewski M, Björntorp P. Muscle tissue in obesity with different distribution of adipose tissue. Effects of physical training. *Int J Obes.* 1986;10:331-341.

55. Hamm P, Shekelle RB, Stamler J. Large fluctuations in body weight during young adulthood and twenty-five-year risk of coronary death in men. *Am J Epidemiol.* 1989;129:312-318.
56. Ballard-Barbash R, Schatzkin A, Taylor PR, Kahle LL. Association of change in body mass with breast cancer. *Cancer Res.* 1990;50:2152-2155.
57. Lipid Research Clinics Program. The Lipid Research Clinics Coronary Primary Prevention Trial Results. II The relationship of reduction in incidence of coronary heart disease to cholesterol lowering. *JAMA.* 1984;251:365-374.
58. Gordon DJ, Probstfield JL, Garrison RJ, et al. High-density lipoprotein cholesterol and cardiovascular disease. Four prospective American studies. *Circulation.* 1989;79:8-15.
59. Bruning PF, Bonfrère JMG. Free fatty acid concentrations correlated with the available fraction of estradiol in human plasma. *Cancer Res.* 1986;46:2606-2609.
60. Després J-P, Lamarche B, Mauriège P, et al. Hyperinsulinemia as an independent risk factor for ischemic heart disease. *N Engl J Med.* 1996;334:952-957.
61. Bruning PF, Bonfrère JMG, Van Noord PAH, Hart AAM, De Jong-Bakker M, Nuijten WJ. Insulin resistance and breast-cancer risk. *Int J Cancer.* 1992;52:511-516.

Appendix I

Questionnaire; Oslo 1972-73

Finnmark 1974-75

Tromsø 1974

Sogn og Fjordane 1975-76

Oppland 1976-78

English translation of the questionnaire used in the cardiovascular disease study in Norwegian counties 1977-83 (Finnmark, Sogn og Fjordane, Oppland) and Tromsø 1979-80

English translation; Mrs. Anne Clancy and Mr. Kevin McCafferty

Tick "yes/no" or "yes", as appropriate.

Part A

Have you, or have you had:
a heart attack?
angina pectoris (heart cramp)?
any other heart disease?
arteriosclerosis of the legs?
a cerebral stroke?
diabetes?

Are you being treated for:
high blood pressure?
Do you use:
nitroglycerine?

Part B

Do you have pain or discomfort in the chest when:
- walking up hills or stairs, or walking fast on level ground?
- walking at normal pace on level ground?

If you get pain or discomfort in the chest when walking, do you usually:
(1) stop?
(2) slow down?
(3) carry on at the same pace?

If you stop or slow down, does the pain disappear:
(1) within 10 minutes?
(2) after more than 10 minutes?

Do you have pain in the calf while:
- walking?
- resting?

If you get pain in the calf, then:
- does the pain increase when you walk faster or uphill?
- does the pain disappear if you stop?

Do you usually have:
- cough in the morning?
- phlegm chest in the morning?

Part C

Exercise and physical exertion in *leisure time*.
If your activity varies much, for example between summer and winter, then give an average. The questions refer only to the last twelve months.

Tick "YES" beside the description that fits best:

- (1) Reading, watching TV, or other sedentary activity?
- (2) Walking, cycling, or other forms of exercise at least 4 hours a week? (including walking or cycling to place of work, Sunday-walking, etc.)
- (3) Participation in recreational sports, heavy gardening, etc.? (note: duration of activity at least 4 hours a week).
- (4) Participation in hard training or sports competitions, regularly several times a week?

Part D

Do you smoke daily at present?

If "Yes":
Do you smoke cigarettes daily?
(handrolled or factory made)

If you do not smoke cigarettes at present:
Have you previously smoked cigarettes daily?

If "Yes", how long is it since you stopped?
(1) Less than 3 months?
(2) 3 months to 1 year?
(3) 1 to 5 years?
(4) More than 5 years?

For those who smoke or have smoked previously:
How many years altogether have you smoked daily? *Number of years*

How many cigarettes do you, or did you, smoke daily? Give number of cigarettes per day (handrolled + factory made)

Number of cigarettes

Do you smoke tobacco products other than cigarettes daily?

- cigars or cigarillos?
- a pipe?

If you smoke a pipe, how many packs of tobacco (50 grams) do you smoke per week?

Give average number of packs per week.

Number of tobacco packs

Part E

Do you usually work shifts or at night?

Can you usually come home from work:

- every day?
- every weekend?

Are there periods during which your working days are longer than usual? (e.g.: fishing season, harvest)

During the last year, have you had: (Tick "YES" beside description that fits best):

- (1) mostly sedentary work? (e.g., office work, watchmaker, light manual work)
- (2) work that requires a lot of walking? (e.g., shop assistant, light industrial work, teaching)
- (3) work that requires a lot of walking and lifting? (e.g., postman, heavy industrial work, construction)
- (4) heavy manual labour? (e.g., forestry, heavy farmwork, heavy construction)

During the last 12 months, have you had to move house for work reasons?

Is housekeeping your main occupation?

Have you within the last 12 months received unemployment benefit?

Are you at present on sick leave, or receiving rehabilitation allowance?

Do you receive a complete or partial disability pension?

Part F (alternatives: yes, no, don't know)

Have one or more of your parents or sisters or brothers had a heart attack (heart wound) or angina pectoris (heart cramp)?

In Finnmark and Tromsø only:

Are two or more of your grandparents of Finnish origin?

Are two or more of your grandparents of Lapp origin?

Part G

Has anyone in your household (other than yourself), been called in to a doctor for further medical examination after the previous cardiovascular disease survey?

Original questionnaire, Tromsø 1974

Identical in; Finnmark 1974-75
Sogn og Fjordane 1975-76
Oppland 1976-78

Innkallings Brev

UNDERSØKELSE AV RISIKO FOR HJERTE- OG KARSYKDOMMER

Hjerte- og karsykdommer er mer utbredt i Nord-Norge enn i andre landsdeler. Vi vil nå be om Deres samarbeid i kampen mot disse sykdommer.

Med sikte på å sette i verk forebyggende tiltak, vil det i løpet av 1974 bli gjennomført en undersøkelse av samtlige menn i Tromsø i alderen 20—49 år.

Undersøkelsen utføres av Universitetet i Tromsø og Tromsø helseråd. Undersøkelsen er frivillig, men forutsetningen for at den skal få verdi, er at alle møter opp.

UNDERSØKELSENS GJENNOMFØRING

1. Vennligst fyll ut vedlagte spørreskjema.
2. Ta med dette skjema og møt fram til undersøkelse i Poliklinikkbygget, Sentralsykehuset ca. 1 uke etter at dette brev er mottatt. Poliklinikken er åpen mandag, tirsdag, onsdag, torsdag og fredag, kl. 08.30—18.00.

Undersøkelsen vil omfatte:

- a. Måling av blodtrykk.
- b. Gjennomgåelse av det utfylte skjema, samt utfylling av tilleggs skjema.
- c. Måling av høyde og vekt.
- d. Blodprøve.

Undersøkelsen vil ta ca. 30 minutter. Opplysningene behandles konfidensielt.

En del av dem som blir undersøkt, vil etter bestemte regler bli innkalt til undersøkelse på et senere tidspunkt.

Hvis undersøkelsen gir holdepunkter for at De kan ha nytte av behandling, vil De bli henvist til lege for videre kontroll.

Dersom det er vanskelig for Dem å møte fram ca. 2 uker etter at dette brev er mottatt, vennligst ring telefon 81 100.

TROMSØUNDERSØKELSEN 1974

Vi håper på et godt samarbeid og 100 % frammøte. På forhånd takk.

Hilsen

Arne Nordøy
prof.dr.med.

Hans Ånstad
stadslege

A		JA	NEI	D		JA	NEI
Har De, eller har De hatt:				Røyker De daglig for tiden? 52			
Hjerteinfarkt? 53				Hvis svaret var „JA“ på forrige spørsmål, besvar da:			
Angina pectoris (hjertekrampe)? 54				Røyker De sigaretter daglig? 53			
Annen hjertesykdom? 55				(håndrullede eller fabrikkframstilte)			
Åreforkalkning i beina? 56				Hvis De ikke røyker sigaretter nå, besvar da:			
Hjerneslag? 57				Har De røykt sigaretter daglig tidligere? . . . 54			
Sukkersyke? 58				Hvis De svarte „JA“, hvor lenge er det siden De sluttet?			
Er De under behandling for:				1 Mindre enn 3 måneder? 55			
Høyt blodtrykk? 59				2 3 måneder - 1 år? 56			
Bruker De:				3 1 - 5 år? 57			
Nitroglycerin? 60				4 Mer enn 5 år? 58			
B		JA	NEI	Bevares av dem som røyker nå eller har røykt tidligere:			
Får De smerter eller ubehag i brystet når De:				Hvor mange år tilsammen har De røykt daglig? 54-57			
Går i bakker, trapper eller fort på flat mark? 61				Hvor mange sigaretter røyker eller røykte De daglig? Oppgi antall pr. dag (håndrullede + fabrikkframstilte) 59-61			
Går i vanlig takt på flat mark? 62				Røyker De noe annet enn sigaretter daglig?			
Hvis De får smerter eller ubehag i brystet ved gange, pleier De da å:				Sigarer eller serutter/cigarillos? . . . 62			
1 Stanse? 63				Pipe? 63			
2 Sakte farten? 64				Hvis De røyker pipe, hvor mange pakker tobakk (50 gram) bruker De i pipa pr. uke? 64-66			
3 Fortsette i samme takt? 65				Oppgi gjennomsnittlig antall pakker pr. uke. 64-66			
Hvis De stanser eller saktner farten, forsvinner smertene da:				E			
1 Etter mindre enn 10 minutter? 66				Har De vanligvis skiftarbeid eller nattarbeid? 67			
2 Etter mer enn 10 minutter? 67				Kan De vanligvis komme hjem fra arbeidet:			
Får De smerter i tykkleggen når De:				Hver dag? 68			
Går? 68				Hver helg? 69			
Er i ro? 69				Har De i perioder lengre arbeidsdager enn vanlig? 70			
Hvis De får leggsmerter, besvar da:				(f.eks. under sesongfiske, onnearbeid)			
Forverres smertene ved raskere tempo eller i bakker? 67				Har De i løpet av siste året hatt:			
Gir smertene seg når De stopper? . . . 68				Sett kryss i den ruten hvor „JA“ passer best			
Har De vanligvis:				1 Overveiende stillesittende arbeid? . . 71			
Hoste om morgenen? 69				(f.eks. skrivebordsarb., urmakerarb., montering)			
Oppspytt fra brystet om morgenen? . . 70				2 Arbeid som krever at De går mye? . . 72			
C				(f.eks. ekspeditørarb., lett industriarb., undervien.)			
Bevegelse og kroppslig anstrengelse i Ders fritid				3 Arbeid hvor De går og løfter mye? . . 73			
Hvis aktiviteten varierer meget (f.eks. mellom sommer og vinter så ta et gjennomsnitt)				(f.eks. postbud, tyngre industriarb., bygningsarb.)			
Spørsmålet gjelder bare det siste året.				4 Tungt kroppsarbeid? 74			
Sett kryss i den ruten hvor „JA“ passer best.				(f.eks. skogsarbeid, tungt jordbruksarb., tungt bygningsarb.)			
1 Leser, ser på fjernsyn eller annen stillesittende beskjeftigelse? 71				Har De i løpet av de siste 12 mnd måttet flytte fra hjemstedet på grunn av forandring i arbeidssituasjonen? 72			
2 Spaserer, sykler eller beveger Dem på annen måte minst 4 timer i uken? . . (Heri medregnes også gang eller sykling til arbeidstedet, søndagsturer m.m.)				Er husmorarbeid Dares hovedyrke? 73			
3 Driver mosjonsidrett, tyngre høgearbeid e.l.? 72				Har De i løpet av de siste 12 mnd fått arbeidsledighetstrygd? 74			
(Merk at yrkesheten skal være minst 4 timer i uken.)				Er De for tiden sykmeldt, eller får De attføringspenger? 75			
4 Trener hardt eller driver konkurranseidrett, regelmessig og flere ganger i uken? 73				Har De full eller delvis uførepensjon? . . 76			
D				F			
Har en eller flere av foreldre eller søsken hatt hjerteinfarkt (sår på hjertet) eller angina pectoris (hjertekrampe)? . . 77				Har en eller flere av foreldre eller søsken hatt hjerteinfarkt (sår på hjertet) eller angina pectoris (hjertekrampe)? . . 77			
Er to eller flere av Dares besteforeldre av finak ætt? 78				Er to eller flere av Dares besteforeldre av finak ætt? 78			
Er to eller flere av Dares besteforeldre av samisk ætt? 79				Er to eller flere av Dares besteforeldre av samisk ætt? 79			

Original questionnaire, Oslo 1972-73

FRAMMØTESTED: St. Olavs pl. 5 II.

FRAMMØTETID: Mandag, tirsdag: Menn A-L. Kl. 8.15 - 11.30 og kl. 13.00 - 14.30
Torsdag, fredag: Menn M-Å. Kl. 8.15 - 11.30 og kl. 13.00 - 14.30
De kan også møte mandag, tirsdag og torsdag kl. 15.30 - 18.00, men da må De vennligst ringe 20 10 70 linje 660 og 661 kl. 8.30 - 14.30 for å avtale tid.

De bes møte innen en uke etter at De har mottatt denne innkalling.
Vennligst ta med tuberkulinkort om De har.

NB: Møt helst om formiddagen.

UNDERSØKELSE AV RISIKO FOR HJERTE - KARSYKDOMMER

Samtidig med skjermbildefotograferingen ber vi Dem om - på helt frivillig grunnlag - å medvirke i en stor befolkningsundersøkelse blant menn i Oslo røttet mot hjerte - infarkt og lignende tilstander.

Dersom De er villig, må De på forhånd, så godt som mulig, krysse av svarene på baksiden. Ta skjemaet med ved frammøtet.

Hjerteundersøkelsen vil omfatte:

Måling av høyde og vekt.

Kontroll av skjemaet, med anledning til å drøfte tvilsspørsmål ved utfyllingen.

Måling av blodtrykk.

Blodprøve (ca. 10 ml blod fra vene).

Blodprøven vil bli analysert på Ullevål sykehus.

Som hovedregel gjelder at opplysninger om funnene hos enkeltpersoner ikke gis - heller ikke til den undersøkte lege.

Men en del av dem som blir undersøkt i Oslo Helseråd, vil etter bestemte regler bli innkalt til en etterundersøkelse ved Ullevål sykehus, vanligvis i løpet av 1-3 uker.

Hvis etterundersøkelsen på Ullevål gir som resultat at en mann vurderes som sannsynlig behandlingstrengende, vil han bli henvist til lege for videre kontroll og eventuelt behandling.

Hvis etterundersøkelsen gir som resultat at mennens risiko for hjerte - karsykdom ligger i et nærmere definert grenseområde, vil han bli spurt om han er villig til å delta i et vitenskapelig forsøk på å senke risikoen. Slike forsøk må gå over flere år, og fordelingen av mennene på behandlings- og kontrollgruppe vil skje ved loddtrekning.

Vennlig hilsen
OSLO HELSERÅD

HJERTE - KARSYKDOMMER

SPØRRESKJEMA

		JA	NEI			JA	NEI
Har De, eller har De hatt:							
Hjerteinfarkt?	13	<input type="checkbox"/>	<input type="checkbox"/>				
Angina pectoris (hjertekrampe)?	14	<input type="checkbox"/>	<input type="checkbox"/>				
Armen hjertesykdom?	15	<input type="checkbox"/>	<input type="checkbox"/>				
Åreforkalkning i bena?	16	<input type="checkbox"/>	<input type="checkbox"/>				
Hjerneslag?	17	<input type="checkbox"/>	<input type="checkbox"/>				
Sukkersyke?	18	<input type="checkbox"/>	<input type="checkbox"/>				
Er De under behandling for:							
Høyt blodtrykk?	19	<input type="checkbox"/>	<input type="checkbox"/>				
Bruker De:							
Nitroglyserin?	20	<input type="checkbox"/>	<input type="checkbox"/>				
				JA			
Bevegelse og kroppslig anstrengelse i Deres arbeid. Spørsmålet gjelder bare det siste året. Sett kryss i den ruten hvor "JA" passer best.							
Har De:							
Overveiende stillesittende arbeid? (f.eks. skrivebondsarbeid, umakerarbeid, montering)	21	<input type="checkbox"/>	<input type="checkbox"/>				
Arbeid som krever at De går mye? (f.eks. ekspeditørarb., lett industriarb., undervisning)	22	<input type="checkbox"/>	<input type="checkbox"/>				
Arbeid hvor De går og løfter mye? (f.eks. postbud, tynge industriarb., bygningsarbeid)	23	<input type="checkbox"/>	<input type="checkbox"/>				
Tungt kroppsarbeid? (f.eks. skogerarbeid, tungt jordbruksarb., tungt bygearb.)	24	<input type="checkbox"/>	<input type="checkbox"/>				
				JA			
Bevegelse og kroppslig anstrengelse i Deres fritid. Hvis aktiviteten varierer meget f.eks. mellom sommer og vinter så ta et gjennomsnitt. Spørsmålet gjelder bare det siste året. Sett kryss i den ruten hvor "JA" passer best.							
Leser, ser på fjernsyn eller annen stillesittende baskjefingelse?				25	<input type="checkbox"/>	<input type="checkbox"/>	
Spiser, sykler eller beveger Dem på annen måte minst 7 timer i uken? (Her medregnes også gang eller sykling til arbeidstødet, søndagstur m.m.)				26	<input type="checkbox"/>	<input type="checkbox"/>	
Driver mosjonsidrett, tynge hagearbeid e.l.? (Merk at virksomheten skal være minst 4 timer i uken.)				27	<input type="checkbox"/>	<input type="checkbox"/>	
Trener hardt eller driver konkurransetidrett, regelmessig og flere ganger i uken?				28	<input type="checkbox"/>	<input type="checkbox"/>	
				JA NEI			
Har De i det siste året følt Dem merkbart mere anspent eller irritabel enn før?				29	<input type="checkbox"/>	<input type="checkbox"/>	
Regner De Dem selv for å være en person som presser Dem selv og ofte velger et høyt tempo?				30	<input type="checkbox"/>	<input type="checkbox"/>	
Eller regner De Dem selv for en person som gir avkall på slik innsats, for heller å ha ro og fredelige dager?				31	<input type="checkbox"/>	<input type="checkbox"/>	
Mener De at det er mere enn vanlig press på tidsfrister i Deres arbeidssituasjon?				32	<input type="checkbox"/>	<input type="checkbox"/>	
Roker De daglig for tiden?				33	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis svaret var "JA" på forrige spørsmål, besvar de:							
Inhalerer De?				34	<input type="checkbox"/>	<input type="checkbox"/>	
Roker De sigaretter daglig? (håndrullede eller fabrikkframslåtte)				35	<input type="checkbox"/>	<input type="checkbox"/>	
Bruker De mest filtersigaretter?				36	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis De ikke roker daglig nå, besvar de:							
Har De røkt sigaretter daglig tidligere?				37	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis De svarte "JA", hvor lenge er det siden De sluttet?							
Mindre enn 3 måneder?				38	<input type="checkbox"/>	<input type="checkbox"/>	
3 måneder - 1 år?				39	<input type="checkbox"/>	<input type="checkbox"/>	
1 - 5 år?				40	<input type="checkbox"/>	<input type="checkbox"/>	
Mere enn 5 år?				41	<input type="checkbox"/>	<input type="checkbox"/>	
Hvor mange sigaretter røker eller røkter De daglig? (håndrullede + fabrikkframslåtte)							
1 - 4?				42	<input type="checkbox"/>	<input type="checkbox"/>	
5 - 9?				43	<input type="checkbox"/>	<input type="checkbox"/>	
10 - 14?				44	<input type="checkbox"/>	<input type="checkbox"/>	
15 - 19?				45	<input type="checkbox"/>	<input type="checkbox"/>	
20 - 24?				46	<input type="checkbox"/>	<input type="checkbox"/>	
25 eller flere?				47	<input type="checkbox"/>	<input type="checkbox"/>	
Roker De noe annet enn sigaretter daglig?							
Sigarer eller senutter/cigarillos?				48	<input type="checkbox"/>	<input type="checkbox"/>	
Pipe?				49	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis De røker pipe, hvor mange pakker pipe- tobakk (50 gram) bruker De pr. uke?							
1/2 pakke eller mindre pr. uke?				50	<input type="checkbox"/>	<input type="checkbox"/>	
1/2 - 2 pakker pr. uke?				51	<input type="checkbox"/>	<input type="checkbox"/>	
Mere enn 2 pakker pr. uke?				52	<input type="checkbox"/>	<input type="checkbox"/>	
				JA NEI			
Får De smerter eller ubehag i brystet når De:							
Går i bakker, trapper eller fort på flat mark?				53	<input type="checkbox"/>	<input type="checkbox"/>	
Går i vanlig takt på flat mark?				54	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis De får smerter eller ubehag i brystet ved gange, pleier De da å:							
Stanse?				55	<input type="checkbox"/>	<input type="checkbox"/>	
Sakte farten?				56	<input type="checkbox"/>	<input type="checkbox"/>	
Fortsette i samme takt?				57	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis De stanser eller saktner farten, forsvinner smertene da:							
Etter mindre enn 10 minutter?				58	<input type="checkbox"/>	<input type="checkbox"/>	
Etter mere enn 10 minutter?				59	<input type="checkbox"/>	<input type="checkbox"/>	
Får De smerter i tykkleggen når De:							
Går?				60	<input type="checkbox"/>	<input type="checkbox"/>	
Er i ro?				61	<input type="checkbox"/>	<input type="checkbox"/>	
Hvis De får leggesmerter, besvar de:							
Forverres smertene ved raskere tempo eller i bakker?				62	<input type="checkbox"/>	<input type="checkbox"/>	
Gir smertene seg når De stopper?				63	<input type="checkbox"/>	<input type="checkbox"/>	

Appendix II

Questionnaire; Finnmark 1977-78

Sogn og Fjordane 1980-81

Oppland 1981-83

Tromsø 1979-80 - Questionnaire I

English translation of the questionnaire used in the cardiovascular disease study in Oslo* 1972-73, Norwegian counties 1974-78 (Finnmark, Oppland and Sogn og Fjordane) and Tromsø 1974.

English translation; Mr. Kevin McCafferty

Tick "yes/no" or "yes", as appropriate.

Part A

Have you, or have you had:

- a heart attack?
- angina pectoris (heart cramp)?
- any other heart disease?
- hardened arteries in the legs?
- a cerebral stroke?
- diabetes?

Are you being treated for:

- high blood pressure?

Do you use:

- nitroglycerine?

Part B

Do you have pain or discomfort in the chest when:

- walking up hills or stairs, or walking fast on level ground?
- walking at normal pace on level ground?

If you get pain or discomfort in the chest when walking, do you usually:

- (1) stop?
- (2) slow down?
- (3) carry on at the same pace?

If you stop or slow down, does the pain disappear:

- (1) within 10 minutes?
- (2) after more than 10 minutes?

Do you have pain in the calf while:

- walking?
- resting?

If you get pain in the calf, then:

- does the pain increase when you walk faster or uphill?
- does the pain disappear if you stop?

Do you usually have:

- cough in the morning?
- phlegm chest in the morning?

Part C

Exercise and physical exertion in *leisure time*.

If your activity varies much, for example between summer and winter, then give an average. The questions refer only to the last twelve months.

Tick "YES" beside the description that fits best:

- (1) Reading, watching TV, or other sedentary activity?
- (2) Walking, cycling, or other forms of exercise at least 4 hours a week? (including walking or cycling to place of work, Sunday-walking, etc.)
- (3) Participation in recreational sports, heavy gardening, etc.? (note: duration of activity at least 4 hours a week).
- (4) Participation in hard training or sports competitions, regularly several times a week?

Part D*

Do you smoke daily at present?

If "Yes":

- Do you smoke cigarettes daily? (handrolled or factory made)

If you do not smoke cigarettes at present:

- Have you previously smoked cigarettes daily?

If "Yes", how long is it since you stopped?

- (1) Less than 3 months?
- (2) 3 months to 1 year?
- (3) 1 to 5 years?
- (4) More than 5 years?

For those who smoke or have smoked previously:

How many years altogether have you smoked daily? Number of years

How many cigarettes do you, or did you, smoke daily? Give number of cigarettes per day (handrolled + factory made)
Number of cigarettes

Do you smoke tobacco products other than cigarettes daily?
- cigars or cigarillos?
- a pipe?

If you smoke a pipe, how many packs of tobacco (50 grams) do you smoke per week?
Give average number of packs per week.
Number of tobacco packs

Part E

Do you usually work shifts or at night?
Can you usually come home from work:
- every day?
- every weekend?

Are there periods during which your working days are longer than usual? (e.g.: fishing season, harvest)

*In Oslo preset groups of cigarettes smoked per day and packs of pipe tobacco smoked per day (see original questionnaire)

During the last year, have you had: (Tick "YES" beside description that fits best):

- (1) mostly sedentary work? (e.g., office work, watchmaker, light manual work)
- (2) work that requires a lot of walking? (e.g., shop assistant, light industrial work, teaching)
- (3) work that requires a lot of walking and lifting? (e.g., postman, heavy industrial work, construction)
- (4) heavy manual labour? (e.g., forestry, heavy farmwork, heavy construction)

During the last 12 months, have you had to move house for work reasons?

Is housekeeping your main occupation?

Have you within the last 12 months received unemployment benefit?

Are you at present on sick leave, or receiving rehabilitation allowance?

Do you receive a complete or partial disability pension?

Part F (alternatives: yes, no, don't know)

Have one or more of your parents or sisters or brothers had a heart attack (heart wound) or angina pectoris (heart cramp)?

In Finnmark and Tromsø only:

Are two or more of your grandparents of Finnish origin?

Are two or more of your grandparents of Lapp origin?

**MELDING OM SKJERMBILDEFOTOGRAFERING
OG HJERTE-KARUNDERSØKELSE**

(Gjelder bare den person brevet er adressert til)

Skjermbildefotograferingen kommer nå til
Deres distrikt.

Tid og sted for Deres frammøte vil De finne
nedenfor.

Også denne gangen vil en del av befolkningen
få tilbud om hjerte-karundersøkelse. De tilhører
denne gruppe. En orientering om undersøkelsen
er gitt i vedlagte brosjyre.

Vennligst fyll ut spørreskjemaet på baksiden
og ta det med til undersøkelsen. Ta også med
tuberkulinkort eller helsebok, om De har.

Fravær bes eventuelt meldt på vedlagte seddel.

Med hilsen

HELSE RÅDET FYLKESLEGEN
STATENS SKJERMBILDEFOTOGRAFERING

Født dato	Personnr.	Kommune	Kretsnr.	
Møtested		Kjønn	Første bokstav etternavn Dag og dato	Klokkeslett

SKRIV IKKE HER!

T S M 17 M 18 19 / 24 25 / 30 31 32

A		JA	NEI	D		JA	NEI
Har De, eller har De hatt:				Røyker De daglig for tiden? 52			
Hjerteinfarkt? 53				Hvis svaret var „JA“ på forrige spørsmål, besvar da:			
Angina pectoris (hjertekrampe)? 54				Røyker De sigaretter daglig? 53			
Annen hjertesykdom? 55				(håndrullede eller fabrikkframstilte)			
Åreforkalkning i bena? 56				Hvis De ikke røyker sigaretter nå, besvar da:			
Hjerneslag? 57				Har De røykt sigaretter daglig tidligere? . . 54			
Sukkersyke? 58				Hvis De svarte „JA“, hvor lenge er det siden De sluttet?			
Er De under behandling for:				1 Mindre enn 3 måneder? 55			
Høyt blodtrykk? 59				2 3 måneder - 1 år?			
Bruker De:				3 1 - 5 år?			
Nitroglycerin? 40				4 Mer enn 5 år?			
B		JA	NEI	Besvares av dem som røyker nå eller har røykt tidligere:			
Får De smerter eller ubehag i brystet når De:				Hvor mange år tilsammen har De røykt daglig? 54-57			
Går i bakker, trapper eller fort på flat mark? . . 41				Hvor mange sigaretter røyker eller røykte De daglig? Oppgi antall pr. dag (håndrullede + fabrikkframstilte) 58			
Går i vanlig takt på flat mark? 42				Røyker De noe annet enn sigaretter daglig?			
Hvis De får smerter eller ubehag i brystet ved gange, pleier De da å:				Sigaretter eller serutter/cigarillos? 58			
1 Stanse? 43				Pipe? 59			
2 Sakne farten?				Hvis De røyker pipe, hvor mange pakker tobakk (50 gram) bruker De i pipa pr. uke? 54-56			
3 Fortsette i samme takt?				Oppgi gjennomsnittlig antall pakker pr. uke. 54-56			
Hvis De stanser eller saktnar farten, forsvinner smertene da:				E			
1 Etter mindre enn 10 minutter? 44				Har De vanligvis skiftarbeid eller nattarbeid? . . 67			
2 Etter mer enn 10 minutter?				Kan De vanligvis komme hjem fra arbeidet:			
Får De smerter i tykkleggen når De:				Hver dag? 68			
Går? 45				Hver helg? 69			
Er i ro? 46				Har De i perioder lengre arbeidsdager enn vanlig? 70			
Hvis De får leggsmerter, besvar da:				(f.eks. under sesongfiske, onnearbeid)			
Forverres smertene ved raskere tempo eller i bakker? 47				Har De i løpet av siste året hatt:			
Gir smertene seg når De stopper? 48				Sett kryss i den ruten hvor „JA“ passer best.			
Har De vanligvis:				1 Overveiende stillesittende arbeid? 71			
Hoste om morgenen? 49				(f.eks. skrivebordsarb., urmakerarb., monterer)			
Oppspytt fra brystet om morgenen? . . . 50				2 Arbeid som krever at De gjør mye?			
C				(f.eks. ekspeditørarb., lett industriarb., underviser)			
Bevegelse og kroppslig anstrengelse i Deres fritid.				3 Arbeid hvor De går og løfter mye?			
Hvis aktiviteten varierer meget f.eks. mellom sommer og vinter så ta et gjennomsnitt.				(f.eks. postbud, tyngre industriarb., bygningsarb.)			
Spørsmålet gjelder bare det siste året.				4 Tungt kroppsarbeid?			
Sett kryss i den ruten hvor „JA“ passer best.				(f.eks. skogsarbeid, tungt jordbruksarb., tungt byggningsarb.)			
1 Leser, ser på fjernsyn eller annen stillesittende beskjeftigelse? 51				Har De i løpet av de siste 12 mnd måttet flytte fra hjemstedet på grunn av forandring i arbeidssituasjonen? 72			
2 Spaserer, sykler eller beveger Dem på annen måte minst 4 timer i uken?				Er husmorarbeid Deres hovedyrke? 73			
(Heri medregnes også gang eller sykling til arbeidstedet, søndagsturer m.m.)				Har De i løpet av de siste 12 mnd fått arbeidsledighetstrygd? 74			
3 Driver mosjonsidrett, tyngre hagearbeid e.l.?				Er De for tiden sykmeldt, eller får De attføringspenger? 75			
(Merk at virksomheten skal være minst 4 timer i uken.)				Har De full eller delvis uførepensjon? . . . 76			
4 Trener hardt eller driver konkurranseidrett, regelmessig og flere ganger i uken?				F			
G				Har en eller flere av foreldre eller søsken hatt hjerteinfarkt (sår på hjertet) eller angina pectoris (hjertekrampe)? . . 77			
Har noen i Deres husstand (utenom Dem selv) vært innlagt til nærmere undersøkelse hos distriktslegen eller forrige hjerte-kar undersøkelse? 80				Er to eller flere av Deres besteforeldre av finsk ætt? 78			
				Er to eller flere av Deres besteforeldre av samisk ætt? 79			

Appendix III

Questionnaire; Food frequency questionnaire

Food frequency questionnaire

Prepared by the Section for Dietary Research

University of Oslo

English version in: Gaard M. *Nutritional aspects of cancer of the breast and colon*
An epidemiological study. Thesis Oslo, 1997

Questionnaire

In connection with the present examination, we would like to ask some questions about your dietary habits.

Please, fill in the questionnaire and return it in the envelope provided. The postage is paid by the recipient.

If several in your household have received a questionnaire, each one is asked to fill it in.

All information that you give will be regarded as strictly confidential.

With regards

The Board of Health

County Medical Officer of Health

*Section for Dietary Research
University of Oslo*

National Health Screening Service

Guidance

Answer each question by checking the most appropriate box.

If it is difficult to give an accurate answer, then answer in accordance with your best judgement. Perhaps there will be questions which you cannot answer at all. Leave these questions, and answer as many as possible of the other questions.

14 Do you live on a diet?

1 Yes 2 No

If you are on a regimen, try to fill in the questionnaire, nevertheless.

15 How many slices of bread do you usually eat daily?

1 Less than 2 slices a day

2 2-4 slices a day

3 5-6 slices a day

4 7-8 slices a day

5 9-12 slices a day

6 13 or more slices a day

16 What type of bread do you eat most frequently?

1 Factory made

2 Home made

17 If factory made bread, what type do you eat most often?

1 White bread

2 Medium brown bread

3 Brown bread

18 If home made bread, how much whole meal flour is used?

- 1 Do not use whole meal flour
- 2 Less than 1/4 whole meal flour
- 3 1/4-1/2 whole meal flour
- 4 More than 1/2 whole meal flour

19 What type of fat do you usually spread on bread?

- 1 Nothing
- 2 Butter
- 3 Margarine

20 If you spread margarine on your bread, what brand do you usually use?
.....

21 Check the appropriate package

- 1 Packet
- 2 Beaker

22 Which sandwich spreads do you usually use? Check all the appropriate boxes.

- 22 White cheese
- 23 Whey cheese
- 24 Honey, syrup, sugar
- 25 Jam, marmalade
- 26 Other sweet spreads
- 27 Mayonnaise, salads
- 28 Liver paste
- 29 Cold cuts, bologna
- 30 Sardines, pickled herring

31 How many glasses/cups of milk do you usually drink daily?

- 1 Do not drink milk, or drink less than 1 glass/cup a day
- 2 1 glass/cup a day
- 3 2 glasses/cups a day
- 4 3 glasses/cups a day
- 5 4 glasses/cups a day
- 6 5 or more glasses/cups a day

32 What type of milk do you usually drink?

- 1 Do not drink milk
- 2 Whole milk, sweet or sour
- 3 Skim milk, sweet or sour
- 4 Hand-skimmed milk
- 5 Both whole and skimmed milk

33 How many cups of coffee do you usually drink daily?

- 1 Do not drink coffee or less than 1 cup a day
- 2 1-2 cups a day
- 3 3-4 cups a day
- 4 5-6 cups a day
- 5 7-8 cups a day
- 6 9 or more cups a day

34 How much sugar do you use with/in your coffee?

- 1 Do not drink coffee
- 2 Do not use sugar
- 3 1-2 lumps per cup
- 4 3-4 lumps per cup
- 5 5-6 lumps per cup
- 6 7 lumps per cup

35 How many eggs (boiled, fried) do you usually eat during a week?

- 1 Do not eat, or less than 1 egg a week
- 2 1 egg a week
- 3 2 eggs a week
- 4 3-4 eggs a week
- 5 5-6 eggs a week
- 6 7 or more eggs a week

36 How many oranges do you usually eat during a week?

- 1 Do not eat, or less than 1 orange a week
- 2 1 orange a week
- 3 2 oranges a week
- 4 3-4 oranges a week
- 5 5-6 oranges a week
- 6 7 or more oranges a week

37 How often do your main meal contain fish?

- 1 Less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5-6 times a week
- 5 7 times a week

38 How often do your main meal contain meat (dishes with blood and/or offal included)?

- 1 Less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5-6 times a week
- 5 7 times a week

39 How often do your main meal contain other dishes like porridge, pancakes etc.?

- 1 Less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5 or more times a week

40 How often do you use melted fat (butter, margarine, bacon fat etc.) on or with meat dishes?

- 1 Never, or less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5 or more times a week

41 How often do you use melted fat (butter, margarine, bacon fat etc.) on or with fish dishes?

- 1 Never, or less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5 or more times a week

42 How often do you eat fish liver (when fish liver is available)?

- 1 Never, or less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5 or more times a week

43 How often do you eat potatoes with your main meal in the course of an ordinary week?

- 1 Less than 3 times a week
- 2 3-5 times a week
- 3 6-7 times a week

44 How many potatoes do you usually eat per dinner?

- 1 Less than one per meal
- 2 1 potato per meal
- 3 2 potatoes per meal
- 4 3-4 potatoes per meal
- 5 5 or more per meal

45 How often do you drink soft drinks during an ordinary week?

- 1 Never, or less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5-6 times a week
- 5 7 or more times a week

46 How often do you eat cakes, cookies etc. during an ordinary week?

- 1 Never, or less than once a week
- 2 1-2 times a week
- 3 3-4 times a week
- 4 5-6 times a week
- 5 7 or more times a week

Do you use some of these products during an ordinary week? Check the appropriate boxes.

- 47 Potato chips
- 48 Chocolate, candy
- 49 Wine, liquor
- 50 Beer
- 51 Cod liver oil
- 52 Vitamin supplements

HOW MANY TIMES PER MONTH DO YOU USE ANY OF THE FOLLOWING TYPES OF DISHES WITH YOUR MAIN MEAL?

53 Poached or fried sausages etc.?

- 1 Never, or less than once a month
- 2 1-2 times a month
- 3 3-4 times a month
- 4 5-8 times a month
- 5 More than 8 times a month

54 Meat balls, hamburgers, rissoles etc.

- 1 Never, or less than once a month
- 2 1-2 times a month
- 3 3-4 times a month
- 4 5-8 times a month
- 5 More than 8 times a month

55 Meat stews?

- 1 Never, or less than once a month
- 2 1-2 times a month
- 3 3-4 times a month
- 4 5-8 times a month
- 5 More than 8 times a month

56 Fried or roast meat?

- 1 Never, or less than once a month
- 2 1-2 times a month
- 3 3-4 times a month
- 4 5-8 times a month
- 5 9-16 times a month
- 6 More than 16 times a month

- 57 Poached fish?
1 Never, or
less than once a month
2 1-2 times a month
3 3-4 times a month
4 5-8 times a month
5 9-12 times a month
6 13-16 times a month
7 More than 16 times a
month
- 58 Fish cakes, fish balls, processed
fish?
1 Never, or
less than once a month
2 1-2 times a month
3 3-4 times a month
4 5-8 times a month
5 More than 8 times a month
- 59 Fried fish?
1 Never, or
less than once a month
2 1-2 times a month
3 3-4 times a month
4 5-8 times a month
5 9-12 times a month
6 13-16 times a month
7 More than 16 times a
month
- 60 Fruit soups, stewed fruit?
1 Never, or
less than once a month
2 1-2 times a month
3 3-4 times a month
4 5-8 times a month
5 9-12 times a month
6 13-16 times a month
7 More than 16 times a
month
- 61 How many times do you usually
eat per day? (Include coffee breaks)
1 2 times per day
2 3 times per day
3 4 times per day
4 5 times per day
5 6 or more times per day
- 62 At what time do you eat or drink
for the first time in the morning?
1 Before 6 a.m.
2 Between 6 a.m. and 8 a.m.
3 Between 8 a.m. and 10
a.m.
4 At 10 a.m. or later
- 63 How many bread meals do you
usually have per day?
1 Do not eat bread
2 Once a day
3 2-3 times a day
4 4 or more times a day
- 64 Do you have a household on your
own or with others?
1 Private household alone
2 Private household with
other adults
3 Private household with
adults and children
4 Usually eat in a canteen

65 The drawings below show cubes of butter or margarine in a true scale. *

Mark the cube with which best resembles the amount you spread on a slice of bread. If in doubt, try buttering a slice.

* (see original questionnaire)

- 1 Do not use
- 2 3 grammes
- 3 5 grammes
- 4 8 grammes
- 5 12 grammes

66 Do you make any attempts to change your body weight?

- 1 Yes 2 No

I forbindelse med den undersøkelsen De er med på, vil vi stille Dem noen spørsmål om Deres kosthold og endringer av dette de siste 3 årene.

Vi vil også spørre om endringer av den fysiske aktivitet i fritiden og av røykevaner.

Vennligst fyll ut dette spørreskjemaet og returner det i den vedlagte svarkonvolutt. Portoen vil bli betalt av mottakeren.

Om det skulle være flere i Deres husstand som har fått spørreskjema, ber vi om at hver enkelt fyller det ut.

Opplysningene De gir vil bli behandlet strengt fortrolig.

Med hilsen

Helserådet Fylkeslegen
Avdeling for kostholdsforskning
Universitetet i Oslo
Statens skjermbildefotografering

VEILEDNING FOR UTFYLING AV SPØRRESKJEMAET.

Besvar de enkelte spørsmål ved å sette kryss i den som passer.
Hvis De ikke kan gi et helt nøyaktig svar, vennligst svar da etter beste skjønn.
Det kan forekomme spørsmål som De finner at De i det hele tatt ikke er i stand til å besvare. La disse spørsmål stå åpne, og besvar så mange som mulig av de øvrige.

14 Er De på diett (spesiell kost) nå? 1 <input type="checkbox"/> Ja 2 <input type="checkbox"/> Nei Om De er på diett, så prøv likevel å fylle ut skjemaet.	20 Hvis De bruker margarin på brødet, hvilket merke bruker De vanligvis?
15 Hvor mange brødskeer spiser De vanligvis pr. dag? 1 <input type="checkbox"/> Mindre enn 2 skiver pr. dag 2 <input type="checkbox"/> 2 - 4 skiver pr. dag 3 <input type="checkbox"/> 5 - 6 skiver pr. dag 4 <input type="checkbox"/> 7 - 8 skiver pr. dag 5 <input type="checkbox"/> 9 - 12 skiver pr. dag 6 <input type="checkbox"/> 13 eller flere skiver pr. dag	21 Kryss av for den aktuelle pakning. 1 <input type="checkbox"/> Pakke 2 <input type="checkbox"/> Bordpakning (beget)
16 Hva slags brød spiser De oftest? 1 <input type="checkbox"/> Kjøpt 2 <input type="checkbox"/> Hjemmebakt	22 Hvilke påleggslag bruker De vanligvis? 30 Kryss av i alle ruter som er aktuelle. 22 <input type="checkbox"/> Hvit (gul) ost 23 <input type="checkbox"/> Brun ost 24 <input type="checkbox"/> Honning, sirup, sukker (på brød) 25 <input type="checkbox"/> Syltetøy, marmelade 26 <input type="checkbox"/> Andre søte påleggslag (sunda, sjokolade, banan, nøtte m.v.) 27 <input type="checkbox"/> Majones, salater 28 <input type="checkbox"/> Leverpostei 29 <input type="checkbox"/> Spekepølse (salt pølse) og annet kjøttpølegg 30 <input type="checkbox"/> Sardin, sursild, speket fisk og annet fiskepølegg
17 Hvis kjøpt brød, hva slags oftest? 1 <input type="checkbox"/> Løff 2 <input type="checkbox"/> Fint (lyst) brød 3 <input type="checkbox"/> Grovt (mørkt) brød	31 Hvor mange glass eller kopper melk drikker De vanligvis pr. dag? 1 <input type="checkbox"/> Drikker ikke, eller mindre enn 1 glass eller kopp pr. dag 2 <input type="checkbox"/> 1 glass eller kopp pr. dag 3 <input type="checkbox"/> 2 glass eller kopper pr. dag 4 <input type="checkbox"/> 3 glass eller kopper pr. dag 5 <input type="checkbox"/> 4 glass eller kopper pr. dag 6 <input type="checkbox"/> 5 eller flere glass eller kopper pr. dag
18 Hvis hjemmebakt brød, hvor stor andel av melet er grovt (mørkt)? 1 <input type="checkbox"/> Bruker ikke grovt mel 2 <input type="checkbox"/> Mindre enn 1/4 grovt mel 3 <input type="checkbox"/> 1/4 - 1/2 grovt mel 4 <input type="checkbox"/> Mer enn 1/2 grovt mel	
19 Hva pleier De vanligvis å smøre på brødet? 1 <input type="checkbox"/> Bruker ikke noe 2 <input type="checkbox"/> Smør (meierismør) 3 <input type="checkbox"/> Margarin	

estrykk

<p>32 Hva slags melk drikker De vanligvis?</p> <p>1 <input type="checkbox"/> Drikker ikke melk</p> <p>2 <input type="checkbox"/> Melk (helmelk), søt, sur</p> <p>3 <input type="checkbox"/> Skummet melk, søt, sur</p> <p>4 <input type="checkbox"/> Håndskummet melk</p> <p>5 <input type="checkbox"/> Både helmelk og skummet melk</p>	<p>39 Hvor ofte består middagsmåltidet av andre typer retter, som grøt, pannekaker m.v.?</p> <p>1 <input type="checkbox"/> Sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 eller flere ganger i uken</p>
<p>33 Hvor mange kopper kaffe drikker De vanligvis pr. dag?</p> <p>1 <input type="checkbox"/> Drikker ikke, eller mindre enn 1 kopp pr. dag</p> <p>2 <input type="checkbox"/> 1 - 2 kopper pr. dag</p> <p>3 <input type="checkbox"/> 3 - 4 kopper pr. dag</p> <p>4 <input type="checkbox"/> 5 - 6 kopper pr. dag</p> <p>5 <input type="checkbox"/> 7 - 8 kopper pr. dag</p> <p>6 <input type="checkbox"/> 9 eller flere kopper pr. dag</p>	<p>40 Hvor ofte bruker De fett (smør, margarin, kjøttfett eller fleskefett) til eller på kjøtt?</p> <p>1 <input type="checkbox"/> Aldri eller sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 eller flere ganger i uken</p>
<p>34 Hvor mye sukker bruker De vanligvis til eller i kaffen?</p> <p>1 <input type="checkbox"/> Drikker ikke kaffe</p> <p>2 <input type="checkbox"/> Bruker ikke sukker til/i kaffen</p> <p>3 <input type="checkbox"/> 1 - 2 biter/teskjeer pr. kopp</p> <p>4 <input type="checkbox"/> 3 - 4 biter/teskjeer pr. kopp</p> <p>5 <input type="checkbox"/> 5 - 6 biter/teskjeer pr. kopp</p> <p>6 <input type="checkbox"/> 7 eller flere biter eller teskjeer pr. kopp</p>	<p>41 Hvor ofte bruker De fett (smør, margarin, kjøttfett eller fleskefett) til eller på fisk?</p> <p>1 <input type="checkbox"/> Aldri eller sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 eller flere ganger i uken</p>
<p>35 Hvor mange egg (kokte eller stekte) spiser De vanligvis i uken?</p> <p>1 <input type="checkbox"/> Spiser ikke, eller mindre enn 1 egg i uken</p> <p>2 <input type="checkbox"/> 1 egg i uken</p> <p>3 <input type="checkbox"/> 2 egg i uken</p> <p>4 <input type="checkbox"/> 3 - 4 egg i uken</p> <p>5 <input type="checkbox"/> 5 - 6 egg i uken</p> <p>6 <input type="checkbox"/> 7 eller flere egg i uken</p>	<p>42 Hvor ofte spiser De fiskelever (i perioder fiskelever er å fisk)?</p> <p>1 <input type="checkbox"/> Aldri eller sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 eller flere ganger i uken</p>
<p>36 Hvor mange appelsiner spiser De vanligvis i uken?</p> <p>1 <input type="checkbox"/> Spiser ikke, eller mindre enn 1 appelsin i uken</p> <p>2 <input type="checkbox"/> 1 appelsin i uken</p> <p>3 <input type="checkbox"/> 2 appelsiner i uken</p> <p>4 <input type="checkbox"/> 3 - 4 appelsiner i uken</p> <p>5 <input type="checkbox"/> 5 - 6 appelsiner i uken</p> <p>6 <input type="checkbox"/> 7 eller flere appelsiner i uken</p>	<p>43 Hvor ofte spiser De poteter til middag i løpet av en vanlig uke?</p> <p>1 <input type="checkbox"/> Sjeldnere enn 3 ganger i uken</p> <p>2 <input type="checkbox"/> 3 - 5 ganger i uken</p> <p>3 <input type="checkbox"/> 6 - 7 ganger i uken</p>
<p>37 Hvor ofte består middagsmåltidet av fisk eller retter med fisk?</p> <p>1 <input type="checkbox"/> Sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 - 6 ganger i uken</p> <p>5 <input type="checkbox"/> 7 ganger i uken</p>	<p>44 Hvor mange poteter spiser De vanligvis til middag?</p> <p>1 <input type="checkbox"/> Mindre enn én potet pr. måltid</p> <p>2 <input type="checkbox"/> 1 potet pr. måltid</p> <p>3 <input type="checkbox"/> 2 poteter pr. måltid</p> <p>4 <input type="checkbox"/> 3 - 4 poteter pr. måltid</p> <p>5 <input type="checkbox"/> 5 eller flere poteter pr. måltid</p>
<p>38 Hvor ofte består middagsmåltidet av kjøtt eller retter med kjøtt (også retter med blod og innmat)?</p> <p>1 <input type="checkbox"/> Sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 - 6 ganger i uken</p> <p>5 <input type="checkbox"/> 7 ganger i uken</p>	<p>45 Hvor ofte drikker De saft, brus eller andre søte drikker i løpet av en vanlig uke?</p> <p>1 <input type="checkbox"/> Aldri eller sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 - 6 ganger i uken</p> <p>5 <input type="checkbox"/> 7 eller flere ganger i uken</p>
	<p>46 Hvor ofte spiser De kaker, kjeks, vafler eller lefser i løpet av en vanlig uke?</p> <p>1 <input type="checkbox"/> Aldri eller sjeldnere enn én gang i uken</p> <p>2 <input type="checkbox"/> 1 - 2 ganger i uken</p> <p>3 <input type="checkbox"/> 3 - 4 ganger i uken</p> <p>4 <input type="checkbox"/> 5 - 6 ganger i uken</p> <p>5 <input type="checkbox"/> 7 eller flere ganger i uken</p>

47 Bruker De noe av de følgende varer i løpet av en vanlig uke?
Kryss av i alle ruter som er aktuelle.

- 47 Potetgull (potetchips)
48 Sjokolade, konfekt, drops eller pastiller
49 Vin, brennevin
50 Øl (uansett type)
51 Tran
52 Vitaminpiller eller vitaminpreparat

HVOR MANGE GANGER I MÅNEDEN SPISER DE NOEN AV DE FØLGENDE RETTER TIL MIDDAG?
Gjelder spørsmålene 53-60.

53 Kokte eller stekte pølser

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden (inntil én gang i uken)
4 5 - 8 ganger i måneden (inntil 2 ganger i uken)
5 Mer enn 8 ganger i måneden (mer enn 2 ganger i uken)

54 Kjøttkaker, karbonader og liknende

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 Mer enn 8 ganger i måneden

55 Kokt kjøtt, fårikål, kjøttsuppe, lapskaus

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 Mer enn 8 ganger i måneden

56 Stekte kjøttretter (koteletter, småstek m.v.)

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 9 - 16 ganger i måneden
6 Mer enn 16 ganger i måneden

57 Kokt fisk

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 9 - 12 ganger i måneden
6 13 - 16 ganger i måneden
7 Mer enn 16 ganger i måneden

58 Fiskekaker, fiskepudding, fiskeboller

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 Mer enn 8 ganger i måneden

59 Stekt fisk

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 9 - 12 ganger i måneden
6 13 - 16 ganger i måneden
7 Mer enn 16 ganger i måneden

60 Søtsuppe, fruktsuppe, fruktgrøt, kompot

- 1 Aldri eller sjeldnere enn én gang i måneden
2 1 - 2 ganger i måneden
3 3 - 4 ganger i måneden
4 5 - 8 ganger i måneden
5 9 - 12 ganger i måneden
6 13 - 16 ganger i måneden
7 Mer enn 16 ganger i måneden

61 Hvor mange ganger spiser De vanligvis pr. dag (tell også med kaffemåltider)?

- 1 2 ganger pr. dag
2 3 ganger pr. dag
3 4 ganger pr. dag
4 5 ganger pr. dag
5 6 eller flere ganger pr. dag

62 Når spiser eller drikker De første gang om morgenen?

- 1 Før kl. 0600
2 Mellom kl. 0600 og kl. 0800
3 Mellom kl. 0800 og kl. 1000
4 Kl. 1000 eller senere

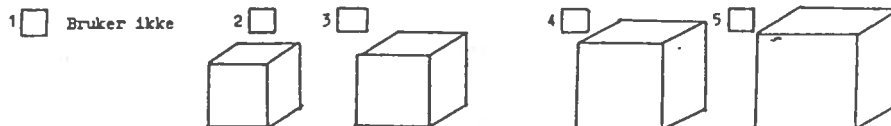
63 Hvor mange ganger om dagen spiser De brødmat?

- 1 Spiser ikke brød
2 1 gang pr. dag
3 2 - 3 ganger pr. dag
4 4 eller flere ganger pr. dag

64 Har De husholdning alene eller sammen med andre?

- 1 Har privat husholdning alene
2 Har privat husholdning sammen med voksne
3 Har privat husholdning sammen med voksne og barn
4 Spiser hovedsaklig i messe, kantine (storbusholdning)

65 Nedestående tegninger forestiller terninger av smør eller margarin i naturlig størrelse. Kryss av for den terning som likner mest på den mengde De bruker til en skive brød. Er De i tvil, forsøk å prøvesmøre en skive.



66 Gjør De noe forsøk på å forandre kroppsvekten Deres?

1 Ja 2 Nei

Dette spørreskjemaet er utarbeidet av Avdeling for kostholdsforskning, Universitetet i Oslo for bruk i Statens Skjermbildefotograferings hjerte- karundersøkelser. For bruk i Finnmark II ble skjemaet også oversatt til samisk.

Siden skjemaet belyser spesielle sider av kostholdet er det ikke uten videre egnet til å gi en generell beskrivelse. Vi ber om at andre grupper som måtte være interessert i å bruke skjemaet eller deler av det, først kontakter oss, og at det blir gitt kildehenvisning.

*Avdeling for kostholdsforskning
Universitetet i Oslo*

Appendix IV

Questionnaire; Tromsø 1979-80 - Questionnaire II

ADDITIONAL QUESTIONS FOR PERSONS ATTENDING THE MASS X-RAY EXAMINATION IN TROMSØ.

English translation; Mrs. Anne Clancy and Mr. Kevin McCafferty

Together with the invitation to attend you received a questionnaire from the National Mass Radiography Service. You delivered this questionnaire at the examination.

Cardiovascular diseases are, however, a complex group of diseases. The causes are still partly unknown. In Tromsø we are therefore trying to obtain a more complete description of factors which may be of importance for the course of these diseases, such as diet, psychological pressure ("stress"), social conditions, and occurrence of disease in relatives. We hope you will take the trouble to complete this questionnaire as well, and return it to the Tromsø Board of Health in the enclosed envelope.

All information in connection with the mass x-ray examination will be treated as strictly confidential.

I YOUR OWN DIET

1. What type of bread do you usually eat?
Tick the most appropriate box; Yes
- | | |
|---------------------------------|--------------------------|
| White bread (e.g. French bread) | <input type="checkbox"/> |
| Ordinary bread (light texture) | <input type="checkbox"/> |
| Whole meal (brown) bread | <input type="checkbox"/> |
| Home-made (brown) bread | <input type="checkbox"/> |

2. What type of butter or margarine do you usually eat?
Tick the most appropriate box; Yes
- | | |
|-----------------------|--------------------------|
| Butter | <input type="checkbox"/> |
| Ordinary margarine | <input type="checkbox"/> |
| Plant margarine | <input type="checkbox"/> |
| Soft margarine spread | <input type="checkbox"/> |

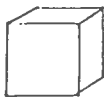
3. How many slices of bread do you usually eat daily?
Tick the most appropriate box; Yes
- | | |
|----------------------|--------------------------|
| Less than two slices | <input type="checkbox"/> |
| 2-6 slices | <input type="checkbox"/> |
| 7-12 slices | <input type="checkbox"/> |
| 13 or more slices | <input type="checkbox"/> |

4. What type of milk do you usually drink?
Tick the most appropriate box; Yes
- | | |
|---|--------------------------|
| Do not drink milk | <input type="checkbox"/> |
| Full cream milk: ordinary type or curdled | <input type="checkbox"/> |
| Skimmed milk: ordinary type or curdled | <input type="checkbox"/> |
| Mixture of full cream and skimmed milk | <input type="checkbox"/> |

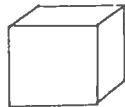
5. The drawings below show cubes of butter or margarine (actual size).
Tick the box above the cube which best resembles the amount you spread on a slice of bread. If in doubt, try buttering a slice.

Do not use butter or margarine

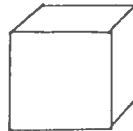
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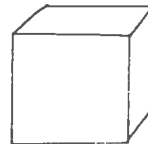
2.



3.



4.



6. How many glasses/cups of milk do you usually drink daily?

Tick the most appropriate box Yes

- Do not drink milk, or drink less than
- 1 glass /cup
- 1-2 glasses
- 3-4 glasses/cups
- 5 or more glasses/cups

7. How many cups of coffee do you usually drink daily?

Tick the most appropriate box Yes

- Do not drink coffee or drink less than
- 1 cup
- 1 - 4 cups
- 5 - 8 cups
- 9 or more cups

8. Are you a teetotaler? Yes No

If "No" :

How often do you usually drink beer?

Tick the most appropriate box Yes

- Never or just a few times a year
- Once or twice a month
- About once a week
- 2-3 times a week
- More or less daily

How often do you usually drink wine?

Tick the most appropriate box Yes

- Never or just a few times a year
- Once or twice a month
- About once a week
- 2-3 times a week
- More or less daily

How often do you usually drink spirits?

Tick the most appropriate box Yes

- Never or a just few times a year
- Once or twice a month
- About once a week
- 2-3 times a week
- More or less daily

9. Approximately how often during the past 12 months have you drunk so much wine, beer or spirits that you got drunk?

Tick the most appropriate box

- Have never been drunk, or have not been drunk during the past year Yes
- A few times during the last year
- Once or twice a month
- Once or twice a week
- 3 or more times a week

10. How often does your main meal consist of fish or fish dishes?

Tick the most appropriate box Yes

- Less than once a week
- Once or twice a week
- 3 - 4 times a week
- 5 - 6 times a week
- 7 days a week

11. How often do you eat fruit or vegetables?

Tick the most appropriate box Yes

- Never eat fruit or vegetables
- A few times a year
- Once or twice a month
- About once a week
- 2 to 3 times a week
- More or less daily

12. How many times a month do you eat boiled sausages or fried meat balls, processed meat, etc.?

Tick the most appropriate box Yes

- Never or less than once a month
- Once or twice a month
- 3 - 4 times a month (up to once a week)
- 5 - 8 times a month (up to twice a week)
- More than 8 times a month, (more than twice a week)

13. Have you made any changes in your diet during the last 5 years as regards the following food items?

Tick each item in the appropriate box

- | | As before | More now | Less now |
|-------------------------------|--------------------------|--------------------------|--------------------------|
| Ordinary margarine or butter: | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Skimmed milk : | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Lean meat: | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Full cream milk: | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Soya margarine (soft): | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Fatty meat: | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |

II. OWN ILLNESSES PAST OR PRESENT

Tick the appropriate box "Yes" or "No"

14. Have you ever had ? *Yes No*
 - Sudden paralysis or numbness on one side of your face or body, in your hand or foot
 -Sudden loss of ability to speak
 -Sudden loss of eyesight, complete or partial, or sudden onset of double vision

15. Have you had a peptic ulcer? *Yes No*

 Do you often have a gnawing pain in the upper part of your stomach?
 Do you suffer much from heartburn or regurgitation of gastric juices?
 Do you suffer much from wind and rumbling in your stomach?
 Do you often get cramps in your stomach ?
 Have you ever had your large intestine x-rayed?
 Have you ever had gall stones?

16. Have you had kidney stones or stones in the urinary tract? *Yes No*

 If yes, how many times ?
 and
 When did you have your last attack? Year:.....

17. Have you ever had cancer? *Yes No*
 If "yes", in what year was the disease discovered?
 Year:

18. Do you have, or have had you the skin disease psoriasis? *Yes No*

19. Have you had allergy-induced eczema on your hands during the last 12 months? *Yes No*

20. Have you been on sick leave, or been unable to work due to allergic eczema on your hands at any time during the past 3 years? *Yes No*

21. Have you ever had arthritis? *Yes No*
 (chronic rheumatoid arthritis)

22. Have you suffered from back pain during the past 12 months lasting for more than 4 weeks? *Yes No*

If "Yes" did the back pain improve if you exercised? *Yes No*

23. Have you suffered from morning stiffness in your back lasting more than 30 minutes? *Yes No*

24. Have you suffered from pains lasting more than 3 months, in the joints listed below during the last 3 years? *Yes No*
 Knees:
 Elbows:
 Innermost finger joints:
 Other joints:

If "Yes", did you suffer from stiff joints in the mornings lasting more than 30 minutes? *Yes No*

25. Have you had any infectious disease during the past 14 days? *Yes No*

 (influenza , common cold , vomiting, diarrhoea, etc.)

26. Have you taken iron tablets during the past 14 days? *Yes No*

27. How often do you take painkillers such as Globoid, Novid, Dispril, Albyl, etc.? *Yes*
 Tick the appropriate box
 1 - 3 times a week
 1 - 3 times a month
 Seldom or never

Have you used such painkillers during the past 14 days? *Yes No*

28. Have you changed the amount of physical exercise you take in leisure during time the last five years? *Yes*
 Tick the most appropriate box.
 As before
 More than before
 Less than before

III ILLNESS IN PARENTS AND SIBLINGS

29. Have any of these relatives had:	mother	father	sister	brother
Cerebral stroke or brain haemorrhage	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Diabetes	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Arthritis (chronic rheumatoid arthritis)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Cancer	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Kidney stones or stone in urinary tract	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Psoriasis	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Peptic ulcer	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
None of the above-mentioned illnesses	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

IV SOCIAL CONDITIONS AND PSYCHOLOGICAL PRESSURE ("STRESS")

30. How many years schooling have you had? (including secondary and folk high schools) number of years

31. What was your family's financial situation when you were growing up?

Tick the appropriate box *Yes*

Very good

Good

Poor

Very poor

32. Do you suffer from sleeplessness? *Yes No*

If "yes", at what time of the year do you suffer from sleeplessness?

Tick the appropriate box *Yes*

No particular time

Especially during the 'dark time'

Especially during the arctic summer (midnight sun)

Especially in spring and autumn

What form your sleeplessness take?

Tick the most appropriate box *Yes*

Difficult to fall asleep at night?

Wake up a lot during the night?

Wake up very early in the morning?

33. Have you had difficulty sleeping in the past couple of weeks?

Tick the most appropriate box *Yes*

Not at all

No more than usual

Rather more than usual

Much more than usual

34. Have you felt unhappy and depressed during the past couple of weeks?

Tick the appropriate box *Yes*

Not at all

No more than usual

Rather more than usual

Much more than usual

35. Have you felt unable to cope with your difficulties during the past couple of weeks?

Tick the appropriate box *Yes*

Not at all

No more than usual

Rather more than usual

Much more than usual

TILLEGGSPØRSMÅL FOR DEM SOM HAR VÆRT
TIL SKJERMBILDEUNDERØKELSE I TROMSØ

ETIKETT

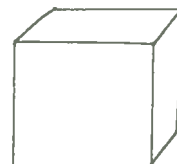
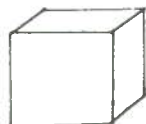
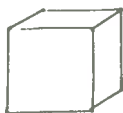
Sammen med innkallingen fikk De et spørreskjema fra Statens Skjermbilde-
fotografering. Dette leverte De ved undersøkelsen.

Hjertekarsykdommene er imidlertid en mangeartet sykdomsgruppe med tildels dårlig
kjente årsaksforhold. I Tromsø vil vi derfor forsøke å få en mer fullstendig kartlegging
av forhold som kan være av betydning for sykdommens forløp, feks. kosthold, psykisk
press ("stress"), sosiale forhold og sykdomsforekomst blant slektninger. Vi håper De vil
være brydd med å fylle ut også dette skjema, og sende det tilbake til Tromsø Helseråd
i den utleverte konvolutt.

Alle opplysninger i forbindelse med skjermbildeundersøkelsen vil bli behandlet
strengt konfidensielt.

<p>I EGET KOSTHOLD</p> <p>1. Hva slags brød spiser De oftest? Sett kryss i den ruten der „JA“ passer best.</p> <p>Løff 1</p> <p>Fint (lyst) brød, alminnelig brød 2</p> <p>Grovt (mørkt) brød, kneipp o.l. 3</p> <p>Hjemmebakt (grovt) brød 4</p>	JA	<p>3. Hvor mange brødskeer spiser De vanligvis daglig? Sett kryss i den ruten der „JA“ passer best.</p> <p>Mindre enn 2 skiver 1</p> <p>2-6 skiver 2</p> <p>7-12 skiver 3</p> <p>13 skiver eller flere 4</p>	JA
<p>2. Hva slags smør eller margarin braker De oftest? Sett kryss i den ruten der „JA“ passer best.</p> <p>Meieri eller fjellsmør 1</p> <p>Vanlig margarin 2</p> <p>Plantemargarin 3</p> <p>Myk (soft) margarin 4</p>	JA	<p>4. Hva slags melk drikker De vanligvis? Sett kryss i den ruten der „JA“ passer best.</p> <p>Drikker ikke melk 1</p> <p>Melk (helmelk), søt, sur 2</p> <p>Skummet melk, søt, sur 3</p> <p>Blanding av skummet og helmelk 4</p>	JA

5. Tegningen nedenfor forestiller terninger av smør eller margarin i naturlig størrelse.
Kryss av for den terning som likner mest på den mengde De bruker til 1 skive brød.
Er De i tvil, forsøk å prøvesmøre en skive.
Bruker ikke smør eller margarin.....



6. Hvor mange glass /kopper melk drikker De vanligvis daglig? JA

Sett kryss i den ruten der „JA“ passer best.

Drikker ikke, eller mindre enn et glass/en kopp 1

1-2 glass /kopper 2

3-4 glass /kopper 3

5 eller flere glass /kopper 4

7. Hvor mange kopper kaffe drikker De vanligvis daglig? JA

Sett kryss i den ruten der „JA“ passer best.

Drikker ikke, eller mindre enn en kopp 1

1-4 kopper 2

5-8 kopper 3

9 eller flere kopper 4

8. Er De totalavholdsmann/kvinne? JA NEI

Hvis nei,

— Hvor ofte pleier De å drikke øl?

Sett kryss i den ruten der „JA“ passer best.

Aldri, eller noen få ganger i året 1

1-2 ganger i måneden 2

Omtrent 1 gang i uken 3

2-3 ganger i uken 4

Omtrent hver dag 5

— Hvor ofte pleier De å drikke vin?

Sett kryss i den ruten der „JA“ passer best.

Aldri, eller noen få ganger i året 1

1-2 ganger i måneden 2

Omtrent 1 gang i uken 3

2-3 ganger i uken 4

Omtrent hver dag 5

— Hvor ofte pleier De å drikke brennevin?

Sett kryss i den ruten der „JA“ passer best.

Aldri, eller noen få ganger i året 1

1-2 ganger i måneden 2

Omtrent 1 gang i uken 3

2-3 ganger i uken 4

Omtrent hver dag 5

9. Omtrent hvor ofte har De i løpet av de siste 12 måneder drukket så mye øl, vin eller brennevin at De har vært beruset? JA

Sett kryss i den ruten der „JA“ passer best.

Har aldri vært beruset eller ikke vært beruset i løpet av siste år 1

Noen få ganger i året 2

1-2 ganger i måneden 3

1-2 ganger i uken 4

3 eller flere ganger i uken 5

10. Hvor ofte består middagsmåltidet av fisk eller retter med fisk? JA

Sett kryss i den ruten der „JA“ passer best.

Sjeldnere enn 1 gang i uken 1

1-2 ganger i uken 2

3-4 ganger i uken 3

5-6 ganger i uken 4

7 ganger i uken 5

11. Hvor ofte bruker De frukt eller grønnsaker? JA

Sett kryss i den ruten der „JA“ passer best.

Bruker aldri frukt eller grønnsaker 1

Noen få ganger i året 2

1-2 ganger i måneden 3

Omtrent 1 gang i uken 4

2-3 ganger i uken 5

Omtrent hver dag 6

12. Hvor mange ganger i måneden spiser De kokte eller stekte pølser, kjøttkaker eller annen opplaget kjøttmat? JA

Sett kryss i den ruten der „JA“ passer best.

Aldri eller sjeldnere enn 1 gang i måneden 1

1-2 ganger i måneden 2

3-4 ganger i måneden (inntil 1 gang i uken) 3

5-8 ganger i måneden (inntil 2 ganger i uken) 4

Mer enn 8 ganger i måneden, (mer enn 2 ganger i uken) 5

13. Har De i løpet av de siste 5 årene forandret
Deres kosthold når det gjelder disse varene?
Sett ett kryss for hver enkelt vare.

Vanlig margarin eller smør

Skummet melk

Magert kjøtt

Helmelk

Soya (soft) margarin

Kjøtt med mye fett

Som før	Mer nå	Mindre nå
<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

18. Har De, eller har De hatt
hudsykdommen psoriasis?

19. Har De i løpet av de siste 12 måneder
hatt allergisk eksem på hendene?

20. Har De i løpet av de siste 3 år vært
sykemeldt eller arbeidsufør p.g.a.
allergisk eksem på hendene?

21. Har De, eller har De hatt leddgikt?
(Kronisk reumatisk artritt).

II TIDLIGERE/NÅVÆRENDE EGNE SYKDOMMER

14. Har De noen gang hatt?

Plutselig lammelse eller nummenhet
i en side av kropp eller ansikt,
i en hånd eller fot

Plutselig tap av taleevnen

Plutselig tap av synet helt eller
delvis, eller plutselig dobbeltsyn

15. Har De hatt magesår?

Har De ofte sugende smerter
øverst i magen?

Har De mye plager med sure
oppstøt eller halsbrann?

Er De mye plaget av oppblåsthet
og rumling i magen?

Har De ofte knipsmerter i magen?

Har De noen gang tatt røntgenbilde
av tykktarmen?

Har De hatt gallestein?

16. Har De hatt nyresteinsanfall
(nyregrus) eller stein i urinveier?

Hvis ja, hvor mange ganger?

og når hadde De siste anfall?

17. Har De noen gang hatt kreftsykdom?

Hvis ja, hvilket år ble sykdommen
oppdaget?

22. Har De i løpet av de siste 12 måneder
vært plaget av smerter i ryggen
som har vært lenger enn 4 uker?
Hvis ja, bedrer ryggsmertene seg
dersom De beveger Dem?

23. Har De vært plaget av stivhet i
ryggen om morgenen som varte
lenger enn 30 minutter?

24. Har De i løpet av de siste 3 år vært
plaget av smerter i noen av de følgende
ledd i mer enn 3 måneder?

Kneleddene

Albuleddene

De innerste fingerleddene

Andre ledd

Hvis ja, merket De stivhet i
leddene om morgenen av mer
enn 30 minutters varighet?

25. Har De hatt noen infeksjonssykdom
de siste 14 dagene?
(Influensa, forkjølelse, "ræksjuka," el.l.)

26. Har De brukt jerntabletter de
siste 14 dagene?

JA NEI

JA NEI

JA NEI

JA NEI

JA NEI

JA NEI

JA NEI

ÅRSTALL:

ANTALL GÅRDER:

ÅRSTALL:

27. Hvor ofte bruker De smertestillende midler som Globoid, Novid, Dispril, Albyl et.l.? Selt kryss iden ruten der „JA“ passer best.	JA	NEI	28. Har De endret mengden av fysisk aktivitet i fritiden de siste 5 årene? Selt kryss i den ruten der „JA“ passer best.	JA
1-3 ganger i uken			Som før	
1-3 ganger i måneden			Mer enn før	
Sjelden eller aldri			Mindre enn før	
Har De brukt slike smertestillende midler de siste 14 dagene?				

III SYKDOMMER HOS FORELDRE OG SØSKEN

29. Har noen av disse slektninger hatt:

	Mor	Far	Søstre	Barn
Hjerneslag eller hjerneblødning				
Sukker sykke				
Leddgiikt (kronisk reumatisk artritt)				
Kreft				
Nyrestein eller stein i urinveier				
Psoriasis				
Magesår				
Ingen av nevnte sykdommer				

IV SOSIALE FORHOLD OG PSYKISK PRESS ("STRESS")		33. Har De i de siste par ukene hatt vansker med å sove? Selt kryss i den ruten der „JA“ passer best.	JA
30. Hvor mange års skolegang har De? (Medregnet folkeskole og ungdomsskole.)	ANTALL ÅR:	Ikke i det hele tatt	
31. Hvordan var de økonomiske forhold i familien under Deres oppvekst? Selt kryss i den ruten der „JA“ passer best.	JA	Ikke mer enn vanlig	
Meget gode		Heller mer enn vanlig	
Gode		Mye mer enn vanlig	
Vanskelige			
Meget vanskelige			
32. Hender det at De er plaget av søvnløshet? Hvis ja, når på året pleier De å være plaget? Selt kryss i den ruten der „JA“ passer best.	JA	34. Har De i de siste par ukene følt Dem ulykkelig og nedtrykt (deprimert)? Selt kryss i den ruten der „JA“ passer best.	JA
Ingen spesiell tid		Ikke i det hele tatt	
Særlig i mørketiden		Ikke mer enn vanlig	
Særlig i midnattstiltiden		Heller mer enn vanlig	
Særlig høst og vår		Mye mer enn vanlig	
Hvordan arter søvnløsheten seg?	JA	35. Har De i de siste par ukene følt Dem ute av stand til å mestre Deres vanskeligheter? Selt kryss i den ruten der „JA“ passer best.	JA
Vanskelig å sove om kvelden		Ikke i det hele tatt	
Våkner ofte i løpet av natten		Ikke mer enn vanlig	
Våkner tidlig om morgenen		Heller mer enn vanlig	
		Mye mer enn vanlig	

Appendix V

Questionnaire; Tromsø 1986-87 - Questionnaire I
Tromsø 1986-87 - Questionnaire II

**QUESTIONNAIRE I, TROMSØ
SURVEY 1986-87**

English translation; Mrs. Anne Clancy and
Mr. Kevin McCafferty

A FAMILY

Have one or both of your parents, or any of
your siblings (brothers and sisters) had a
heart attack or angina pectoris
(heart cramp)?

Yes No Don't know

B OWN ILLNESSES

Have you, or have you had: Yes No
A heart attack?
Angina pectoris (heart cramp)?
A cerebral stroke?
Diabetes?

Are you receiving treatment for: Yes No
High blood pressure?

Do you use nitroglycerine?

C SYMPTOMS

Do you get pain or discomfort
in the chest, when: Yes No
Walking up hills, stairs or walking
fast on level ground?
Walking at ordinary pace
on level ground?

If you get pain or discomfort in your
chest when walking, do you usually :
Yes
Stop
Slow down
Carry on at the same pace

If you stop or slow down, does the pain
disappear: Yes
After less than 10 minutes?
After more than 10 minutes?

D EXERCISE

Exercise and physical exertion in leisure
time. If your activity varies much, for
example between summer and winter, then
give an average. The questions refer only to
the last twelve months.

Tick "yes" in the most appropriate box:

- Reading, watching TV or other sedentary activity? Yes
- Walking, cycling or other forms of exercise at least 4 hours a week?
- (including walking or cycling to place of work, Sunday walking, etc.)
- Participation in recreational sports, heavy gardening, etc.? (Note: duration of activity at least 4 hours a week)
- Participation in hard training or sports competitions regularly several times a week?

E SALT/ FAT

How often do you use salted meat or salted fish for dinner?

- Tick the appropriate box Yes
- Never or less than once a month
 - Once a week or less
 - Twice a week or less
 - More than twice a week

How often do you add extra salt to your dinner?

- Tick the appropriate box Yes
- Rarely or never
 - Sometimes or often
 - Always or nearly always

What type of margarine or butter do you usually use on your bread?

- Tick the most appropriate box Yes
- Do not use margarine or butter on bread
 - Butter
 - Margarine
 - Soft (soya) margarine spread
 - Butter/ margarine mixtures

What type of cooking fat do you normally use in your household?

- Tick the appropriate box. Yes
- Butter or hard margarine
 - Soft (soya) margarine or oil
 - Butter/ margarine mixtures

F SMOKING

Do you smoke daily at present? *Yes No*

If "Yes":

Do you smoke cigarettes daily?
(hand-rolled or factory made)

If you do not smoke cigarettes at present:

Have you previously smoked *Yes No*
cigarettes on a daily basis?

If "Yes", how long is it since you gave up smoking? *Yes*
More than 3 months?
3 months to 1 year?
1 - 5 years?
More than 5 years?

The following questions are to be answered by those who smoke at present or who have smoked previously.

How many years altogether have you smoked on a daily basis:

How many cigarettes do you smoke or did you smoke daily:
(hand-rolled + factory made)

Do you smoke anything else other than cigarettes daily? *Yes*
Cigars, cigarillos, cheroots?
Pipe?

If you smoke a pipe, how many packets of tobacco (50 gr.) do you smoke in a week? Give the average number of packets a week:

G COFFEE

How many cups of coffee do you usually drink daily?

Tick the most appropriate box Yes

Do not drink coffee, or less than one cup
1 - 4 cups
5 - 8 cups
9 or more cups

What type of coffee do you usually drink daily?

Coarse ground coffee for brewing (boiled)
Finely ground filter coffee
Instant coffee
Caffeine free coffee
Do not drink coffee

H EMPLOYMENT

Have you received unemployment benefit within the past 12 months? *Yes No*

Are you at present on sick leave, or receiving rehabilitation allowance?

Are you on a full time or partial disability pension? *Yes No*

Do you usually work shifts or do night work?

During the past year have you had :
Tick the most appropriate box. Yes

- Mostly sedentary work? (office work, watchmaker, light manual work)
- Work requiring a lot of walking? (shop assistant, light industrial work, teaching)
- Work requiring a lot of walking and lifting? (postman, heavy industrial work, construction)
- Heavy manual labour? (forestry, heavy farmwork, heavy construction)

Is house-keeping your main occupation? *Yes No*

I FOLLOW - UP EXAMINATION

Has any one in your household (other than yourself) been called in to a doctor for further medical examination after the previous cardiovascular disease survey? *Yes No*

If as a result of this survey you need further medical examination, which general practitioner do you wish to be referred to ?
Write the doctor's name here:

.....

No particular doctor

HELSEUNDERSØKELSEN I TROMSØ

(Gjelder bare den person som brevet er adressert til.)

Helseundersøkelsen kommer nå til Deres distrikt.

Tid og sted for frammøte vil De finne nedenfor.

De finner en orientering om undersøkelsen i den vedlagte brosjyren.

Vi ber Dem vennligst fylle ut spørreskjemaet på baksiden og ta med dette til undersøkelsen.

Vi ber Dem eventuelt melde fra om fravær på den vedlagte fraværsmeldingen.

Med hilsen

KOMMUNEHELSETJENESTEN I TROMSØ
FYLKESLEGEN I TROMS UNIVERSITETET I TROMSØ
STATENS HELSEUNDERSØKELSER

Født dato Personnr. Kommune Kretsnr.
Motested Kjønn Første bokstav i etternavn Dag og dato Klokkeslett

<input type="text"/> <input type="text"/> <input type="text"/> HOYDE	<input type="text"/> <input type="text"/> <input type="text"/> <input type="text"/> <input type="text"/> <input type="text"/> VEKT	<input type="text"/> <input type="text"/> ANM 70	<input type="text"/> M	<input type="text"/> P	<input type="text"/> O	<input type="text"/> <input type="text"/> <input type="text"/> <input type="text"/> KODE 75	<input type="text"/> AVVIK	<input type="text"/> <input type="text"/> ARM	<input type="text"/> MAN	<input type="text"/> <input type="text"/> <input type="text"/> <input type="text"/> APP. NR	<input type="text"/> <input type="text"/> <input type="text"/> <input type="text"/> TS. 1 82
MÅLING 1			MÅLING 2				MÅLING 3				
MAR <input type="text"/>	S <input type="text"/>	65	68	MAR <input type="text"/>	S <input type="text"/>	91	94	MAR <input type="text"/>	S <input type="text"/>	97	100
HR <input type="text"/>	D <input type="text"/>	103	106	HR <input type="text"/>	D <input type="text"/>	109	112	HR <input type="text"/>	D <input type="text"/>	115	118

A FAMILIE		F RØYKING	
Har en eller flere av foreldre eller søsken hatt hjerteinfarkt (sår på hjertet) eller angina pectoris (hjertekrampe)? 12		Røyker De daglig for tida? 30	
		Dersom svaret er «JA», svar da på dette:	
B EGEN SYKDOM		Røyker De sigaretter daglig? 31	
Har De, eller har De hatt:		(håndrullede eller fabrikkframstilte)	
Hjerteinfarkt? 13		Dersom De ikke røyker sigaretter nå, svar da på dette:	
Angina pectoris (hjertekrampe)? 14		Har De røykt sigaretter daglig tidligere? 32	
Hjerneslag? 15		Dersom De svarte «JA», hvor lenge er det da siden De sluttet?	
Sukkersyke? 16		Mindre enn 3 måneder? 33	
Er De under behandling for:		3 måneder - 1 år? 34	
Høyt blodtrykk? 17		1-5 år? 35	
Bruker De:		Mer enn 5 år? 36	
Nitroglycerin? 18		Skal besvares av de som røyker nå eller som har røykt tidligere:	
C SYMPTOMER		Hvor mange år til sammen har De røykt daglig? 34	
Får De smerter eller ubehag i brystet når De:		Hvor mange sigaretter røyker eller røykte De daglig? 36	
Går i bakker, trapper eller fort på flat mark? 19		Gi opp antallet sigaretter daglig (håndrullede + fabrikkframstilte) 36	
Går i vanlig takt på flat mark? 20		Røyker De noe annet enn sigaretter daglig?	
Dersom De får smerter eller vondt i brystet ved gange, pleier De da:		Sigarer eller serutter/sigarillos? 40	
Stoppe? 21		Pipe? 41	
Sakne farten? 22		Dersom De røyker pipe, hvor mange pakker tobakk (50 gram) bruker De i pipen på en uke?	
Fortsette i samme takt? 23		Gi opp gjennomsnittlig tall på pakker i uken 42	
Dersom De stopper eller saktner farten, går da smertene bort:		G KAFFE	
Etter mindre enn 10 minutter? 22		Hvor mange kopper kaffe drikker De vanligvis hver dag?	
Etter mer enn 10 minutter? 23		Sett kryss i den ruten som passer best.	
Har De vanligvis:		Drikk ikke kaffe, eller mindre enn en kopp 45	
Hoste om morgenen? 23		1 - 4 kopper 46	
Oppspytt fra brystet om morgenen? 24		5 - 8 kopper 47	
D MOSJON		9 eller flere kopper 48	
Bevegelse og kroppslig aktivitet i Deres fritid. Dersom aktiviteten varierer mye, f.eks. mellom sommer og vinter, så ta ett gjennomsnitt. Spørsmålet gjelder bare det siste året.		Hva slags kaffe drikker De vanligvis hver dag?	
Sett kryss i den ruten som passer best.		Kokekaffe 46	
Leser, ser på fjernsyn eller annen stillesittende beskjeftigelse? 25		Filterkaffe 47	
Spaserer, sykler eller beveger Dem på annen måte minst 4 timer i uken? 26		Pulverkaffe 48	
(Her skal De også regne med gang eller sykling til arbeidsteden, søndagstur m.m.)		Koffeinri kaffe 49	
Driver mosjonsidrett, tyngre hagearbeid e.l.? 27		Drikk ikke kaffe 50	
(Merk at aktiviteten skal vare i minst 4 timer i uken.)		H ARBEID	
Trener hardt eller driver konkurranseidrett regelmessig og flere ganger i uken? 28		Har De i de siste 12 månedene fått arbeidsledighetstrygd? 51	
E SALT/FETT		Er De for tiden sykemeldt, eller får De attføringspenger? 52	
Hvor ofte bruker De salt kjøtt eller salt fisk til middag?		Har De full eller delvis uførepensjon? 53	
Sett kryss i den ruten som passer best.		Har De vanligvis skiftarbeid eller nattarbeid 54	
Aldri eller sjeldnere enn en gang i måneden 25		Har De i det siste året hatt:	
Inntil en gang i uken 26		Sett kryss i den ruten som passer best.	
Inntil to ganger i uken 27		For det meste stillesittende arbeid? 55	
Mer enn to ganger i uken 28		(f.eks. skrivebordsarb., umakerarb., montering)	
Hvor ofte pleier De å strø ekstra salt på middagsmaten?		Arbeide som krever at De går mye? 56	
Sett kryss i den ruten som passer best.		(f.eks. ekspeditørb., lett industriarb., undervisn.)	
Sjelden eller aldri 29		Arbeide der De går og løfter mye? 57	
Av og til eller ofte 30		(f.eks. postbud, tyngre industriarb., bygningsarb.)	
Alltid eller nesten alltid 31		Tungt kroppsarbeid? 58	
Hva slags margarin eller smør bruker De vanligvis på brødet?		(f.eks. skogsarb., tungt jordbruksarb., tungt bygningarb.)	
Sett kryss i den ruten som passer best.		I ETTERUNDERSØKELSE	
Bruker ikke smør eller margarin på brød 32		Har noen i husstanden Deres (utenom Dem selv) vært innkalt til nærmere undersøkelse hos lege etter den siste hjerte-undersøkelsen? 59	
Smør 33		Dersom denne helseundersøkelsen viser at De bør undersøkes nærmere: Hvilken almenpraktiserende lege ønsker De da å bli henvist til?	
Hard margarin 34		Skriv navnet på legen her	
Myk (Soft) margarin 35		Ingen spesiell lege 60	
Smør/margarin blanding 36		Ikke skriv her	
Hva slags fett blir vanligvis brukt til matlaging i husholdningen Deres?		Ikke skriv her	
Sett kryss i den ruten som passer best.		61	
Smør eller hard margarin 37			
Myk (Soft) margarin eller olje 38			
Smør/margarin blanding 39			

ADDITIONAL QUESTIONS THE TROMSØ HEALTH SURVEY, 1986 - 87.

English translation; Mrs. Anne Clancy and Mr. Kevin McCafferty

Cardiovascular heart and circulatory diseases, on which the surveys of 1974 and 1979-80 focused, are a very varied category of diseases whose causes are still partly unknown. In Tromsø we are therefore trying to obtain a more complete description of factors which may be important for the course of these diseases, such as diet, psychological pressure, "stress", social conditions and the occurrence of disease in relatives. Such a description is also important in the search for factors that contribute to cancer, a group of diseases which we will also be trying to combat in the coming years.

When you were called in, you received a questionnaire which you handed in at the survey. The present questionnaire asks for further

GENERAL STATE OF HEALTH

How is your health? Tick the appropriate box. Yes

Very bad	<input type="checkbox"/>
Bad	<input type="checkbox"/>
Neither good nor bad. "middling"	<input type="checkbox"/>
Good	<input type="checkbox"/>
Excellent	<input type="checkbox"/>

information about your health and includes questions on various diseases and physical and psychological complaints. We have included questions on pregnancy, birth and menstruation.

In addition, we are interested in obtaining information on the public use of medical services in order to find out how to improve the health service.

We hope that you will take the trouble to fill in yet another questionnaire and return it to "Tromsø Board of Health" in the enclosed envelope. All information will be treated in strict confidence. If you have any comments to make on the survey, you may write them down in the space provided on the last page of the questionnaire.

Yours sincerely

*Tromsø Board Department of
of Health Medicine,
University of Tromsø*

ILLNESS

Have you/ have you had: Tick "yes" or "no" for each question. Yes No

The skin disease psoriasis?	<input type="checkbox"/>	<input type="checkbox"/>
Asthma?	<input type="checkbox"/>	<input type="checkbox"/>
Allergic eczema?	<input type="checkbox"/>	<input type="checkbox"/>
Hay fever?	<input type="checkbox"/>	<input type="checkbox"/>
Chronic bronchitis?	<input type="checkbox"/>	<input type="checkbox"/>
Stomach ulcer?	<input type="checkbox"/>	<input type="checkbox"/>
Duodenal ulcer?	<input type="checkbox"/>	<input type="checkbox"/>
Your appendix removed?	<input type="checkbox"/>	<input type="checkbox"/>
An operation for a stomach ulcer?	<input type="checkbox"/>	<input type="checkbox"/>
Chronic rheumatoid arthritis?	<input type="checkbox"/>	<input type="checkbox"/>
Cancer?	<input type="checkbox"/>	<input type="checkbox"/>
Epilepsy?	<input type="checkbox"/>	<input type="checkbox"/>
Migraine?	<input type="checkbox"/>	<input type="checkbox"/>

INFECTIONS

How many times in the last 6 months have you had infections like a cold, influenza (flu) diarrhoea/vomiting, or similar illnesses? *Number of times*

Have you had one of these infections in the past 14 days? *Yes No*

ILLNESS IN PARENTS OR SIBLINGS

Tick the appropriate box for relatives that have, or have had the following illnesses:

	Mother	Father	Brother	Sister
Cerebral stroke or brain haemorrhage:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Diabetes:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Rheumatoid arthritis:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Cancer:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Psoriasis:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Stomach or duodenal ulcer:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Asthma:	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

Tick the appropriate box if neither your parents nor siblings have or have had any of the above illnesses. *Yes No*

MEDICINES

Have you during the last year used tablets/sprays or had injections for asthma or allergies? *Yes No*

Have you used any of the following medicines in the past 14 days? *Yes No*

Painkillers:	<input type="checkbox"/>	<input type="checkbox"/>
Antipyretics (to reduce fever):	<input type="checkbox"/>	<input type="checkbox"/>
Eczema ointment:	<input type="checkbox"/>	<input type="checkbox"/>
Blood pressure medication:	<input type="checkbox"/>	<input type="checkbox"/>
Heart medication:	<input type="checkbox"/>	<input type="checkbox"/>
Sleeping tablets:	<input type="checkbox"/>	<input type="checkbox"/>
Nerve tablets:	<input type="checkbox"/>	<input type="checkbox"/>
Migraine medication:	<input type="checkbox"/>	<input type="checkbox"/>
Epilepsy medication:	<input type="checkbox"/>	<input type="checkbox"/>
Other medicines:	<input type="checkbox"/>	<input type="checkbox"/>

CONTACT DUE TO OWN HEALTH OR ILLNESS

How many visits have you made during the past year due to your own health or illness? *Number of visits*

To a GP (general practitioner):
To a specialist, (non hospital):
Emergency GP
Medical officer at work :
Physiotherapist:
Chiropractor:
Nature healer(homeopath etc.):
Hospital outpatient department
Number of hospital admissions in the past year:

DIET

How many slices of bread do you usually eat daily?

Tick the most appropriate box. *Yes*

Less than 2 slices	<input type="checkbox"/>
2 - 4 slices	<input type="checkbox"/>
5 - 6 slices	<input type="checkbox"/>
7 -12 slices	<input type="checkbox"/>
13 or more slices	<input type="checkbox"/>

What type of milk do you usually drink? *Tick the most appropriate box.* *Yes*

Do not drink milk	<input type="checkbox"/>
Full cream milk (ordinary or curdled)	<input type="checkbox"/>
Light milk	<input type="checkbox"/>
Skimmed milk (ordinary or curdled)	<input type="checkbox"/>

How many glasses/cups of milk do you usually drink daily? *Yes*

Less than 1 glass/cup	<input type="checkbox"/>
1 - 2 glasses/cups	<input type="checkbox"/>
3 - 4 glasses/cups	<input type="checkbox"/>
5 or more glasses/cups	<input type="checkbox"/>

FISH

How often do you eat cod, coal fish, red snapper or other lean fish for dinner or in a sandwich?

Tick the most appropriate box *Yes*

Less than once a week	<input type="checkbox"/>
Once a week	<input type="checkbox"/>
Twice a week	<input type="checkbox"/>
3 or more times a week	<input type="checkbox"/>

How often do you eat cod/pollock or other lean fish for dinner or in a sandwich?

- Tick the most appropriate box. Yes
- Less than once a week
- Once a week
- Twice a week
- 3 or more times a week

How often do you eat fat fish, such as herring, halibut, mackerel, salmon or trout for dinner or in a sandwich?

- Tick the most appropriate box Yes
- Less than once a week
- Once a week
- Twice a week
- 3 or more times a week

Do you take cod liver oil regularly?

- Tick the most appropriate box Yes
- No
- 'Dark-time' (mid-winter)
- All year

BREAKFAST

Do you usually eat breakfast every day? Yes No

-

DINNER

How often do you eat meat for dinner?

- Tick the appropriate box Yes
- Less than once a week
- Once or twice a week
- 3 - 4 times a week
- 5 or more times a week

How often do you use fat like butter, margarine, mayonnaise, etc. with your dinner?

- Tick the most appropriate box Yes
- Less than once a week
- Once or twice a week
- 3 - 4 times a week
- 5 or more times a week

Do you usually eat vegetables with your dinner? Yes No

-

FRUIT

How often do you usually eat fruit ?

- Tick the appropriate box.
- Less than once a week
- About once a week
- 2 - 3 times a week
- 4 - 5 times a week
- More or less

ALCOHOL

Are you a teetotaler? Yes No

-

If "not", how often do you drink beer?

- Tick the most appropriate box Yes
- Never or just a few times a year
- Once or twice a month
- About once a week
- 2 - 3 times a week
- More or less daily

How often do you drink wine ?

- Tick in the most appropriate box Yes
- Never or just a few times a year
- Once or twice a month
- About once a week
- 2 - 3 times a week
- More or less daily

How often do you drink spirits ?

- Tick the appropriate box Yes
- Never or just a few times a year
- Once or twice a month
- Approximately once a week
- 2 or 3 times a week
- More or less daily

Approximately how often in the past year have you drunk alcohol corresponding to at least 5 small bottles of beer, a bottle of wine, or a quarter bottle of spirits?

- Tick the most appropriate box Yes
- Not at all the past year
- A few times
- Once or twice a month
- 3 or more times a week

PHYSICAL ACTIVITY

How often do you take part in physical activity lasting at least 20 minutes, which makes you perspire or become breathless?

Tick the appropriate box. *Yes*

- Rarely or never
- Weekly
- Several times a week
- Daily

If you usually take part in this type of activity at least weekly, how much time do you spend exercising?

Tick the most appropriate box. *Yes*

- Less than 30 minutes a week
- Between 30 minutes and one hour weekly
- Between 1 and 2 hours a week
- More than 2 hours a week

CHANGE IN DIETARY HABITS AND OTHER HABITS

Have you changed any of the following habits during the last 5 years?

Tick the appropriate box. *Use now*

- | | <i>More</i> | <i>As before</i> | <i>Less</i> |
|-------------------------|--------------------------|--------------------------|--------------------------|
| Dietary fat | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Soya margarine or oil | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Skimmed or low fat milk | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Coffee intake | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Alcohol intake | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |
| Physical activity | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> |

MARRIAGE / PARTNER

Are you married or 'living together'? *Yes No*

How old were you when you first married or moved in with a partner? *age:*

HOUSEHOLD

How many persons live in your household?
Number of persons :

Is anyone in your household 10 years or younger? *Yes No*

Does anyone in your household need special care/assistance? *Yes No*
(Other than the children)

SCHOOLING

How many years schooling have you had? (include secondary and folk high schools)

Number of years :

EMPLOYMENT

Have you had paid work this past year?

Tick the appropriate box *Yes*

- Full-time work
- Part-time work
- Unpaid work

How much house work do you normally do yourself?

Tick the appropriate box *Yes*

- All or almost all
- At least half
- More than a quarter
- Less than a quarter

BACK AND JOINTS CONDITIONS

During this last year have you suffered from backache that has lasted longer than 4 weeks?

Yes No

If "yes", does the pain improve when you exercise? *Yes No*

Have you suffered from morning stiffness in your back lasting more than 30 minutes?

Yes No

During the past 3 years have you suffered from pain in any of the following joints lasting more than 30 minutes? *Yes No*

- Knees
- Elbows
- Innermost finger joints
- Other joints

If "yes", have you suffered from stiff joints in the mornings lasting more than 30 minutes? *Yes No*

NECK HEAD AND SHOULDER COMPLAINTS

How often do you suffer headache?

- Tick the appropriate box *Yes*
- Rarely or never
- Once or twice a month
- Once or twice a week
- Daily

How often do you suffer pain in the neck or shoulder?

- Tick the appropriate box *Yes*
- Rarely or never
- Once or twice a month
- Once or twice a week
- Daily

Do these complaints inhibit your ability to work?

- Tick the appropriate box. *Yes*
- Little or no effect
- To some degree
- To a large degree
- Cannot do ordinary work

Have your back, shoulders, and /or neck ever been x-rayed?

- Yes No*
-

SLEEPLESSNESS / LOSS OF CONSCIOUSNESS

Have you ever suffered from sleeplessness? *Yes No*

-

If "yes", at what time of the year do you usually suffer from sleeplessness?

- Tick the appropriate box *Yes*
- No particular time
- Especially during the "dark time"
- Especially during the arctic summer (midnight sun)
- Especially in spring and autumn

Have you at any time during the last 12 twelve months suffered from tiredness that has affected your work performance? *Yes No*

-

Have you suffered from sudden loss of consciousness in the past year? *Yes No*

-

Have you noticed sudden changes in your pulse rate or heartbeat in the past year? *Yes No*

-

REACTION TO PROBLEMS

If you have major personal problems, do you expect to get help and support from your spouse or family? *Yes No*

-

In the last year, have you long felt a need to seek help with personal problems, without doing so? *Yes No*

-

During the past 2 weeks have you felt unable to cope with your problems?

- Tick the appropriate box *Yes*
- Seldom or never
- Sometimes
- Often
- Nearly always

During the past 2 weeks have you felt unhappy or depressed?

- Tick the appropriate box *Yes*
- Seldom or never
- Sometimes
- Often
- Nearly always

Do you ever feel lonely?

- Tick the appropriate box *Yes*
- Very often
- Sometimes
- Rarely or never

THE REMAINING SECTION OF THE QUESTIONNAIRE APPLIES TO WOMEN ONLY.

MENSTRUATION

How old were you when you started menstruating? *age:*

When did you start (date, month, year) your last period?

How many days usually pass from the first day of one period to the first day of your next period (the time lapsed between the start of two periods)?

Number of days:

Do/did you menstruate regularly? *Yes No*

Do you usually need pain-killers during menstruation? *Yes No*

PRE-MENSTRUAL TENSION

Do you have any of the following complaints before your period?

Are you depressed or irritable? *Yes*

Tick the appropriate box

Hardly at all

Noticeably

Very much so

Are your breasts painful? *Yes*

Tick the appropriate box

Hardly at all

Noticeably

Very much so

Do you have swollen hands/feet, put on weight, or feel bloated?

Tick the appropriate box. Yes

Hardly at all

Noticeably

Very much so

Do the complaints disappear when you get your period? *Yes No*

What type of medication do you use for these complaints?

Tick the appropriate box. Yes

Diuretics

Other medicines

PREGNANCY

How many children have you had?

Number of children:

How old were you when you got pregnant for the first time? *Age:*

CONTRACEPTION

Do you now use or have you ever used the contraceptive pill or an intrauterine device?

Yes No

If "yes", for how many years altogether have you used: *Number of years*

The pill:

An intrauterine device:

How old were you when you started using:

The pill: *age:*

An intrauterine device: *age:*

If you stopped taking the pill, did 6 months or more pass without menstruating (having a period), without your being pregnant?

Yes No

Did you have to stop taking the pill due to high blood pressure? *Yes No*

CERVICAL SMEAR TEST

How many times have you had a cervical smear test in the last 3 years?

Number of times:

How many years is it since you had your last cervical smear test?

Number of years:

Comments

.....

Thank you for your help! Remember to post the questionnaire today!

The Tromso survey 1986-1987.

Tilleggsspørsmål til Helseundersøkelsen i Tromsø 1986-87.

Hjerte-karsykdommene, som Hjerte-karundersøkelsene i 1974 og 1979-80 spesielt tok opp, er en mangeartet sykdomsgruppe med tildels dårlig kjente årsaksforhold. I Tromsø vil vi derfor forsøke å få en mer fullstendig kartlegging av forhold som kan være av betydning for sykdommens forløp, f.eks. kosthold, psykisk press «stress», sosiale forhold og sykdomsforekomst blant slektninger. En slik kartlegging er også viktig for å finne fram til sykdomsskapende forhold for kreftsykdommene, som er en sykdomsgruppe vi også vil prøve å bekjempe i årene som kommer.

Sammen med innkallingen fikk De et spørreskjema som De leverte ved undersøkelsen. Dette spørreskjema kartlegger helseforholdene bedre og inkluderer spørsmål om noen forskjellige sykdommer og fysiske/psykiske plager. Spesielt er det tatt med spørsmål vedrørende svangerskap, fødsel og menstruasjon.

Dessuten er vi interessert i å få oversikt over hvordan folk bruker helsetjenesten, for å få kunnskap om hvordan helsetjenesten kan bedres.

Vi håper De vil være brydd med å fylle ut også dette skjemaet, og sende det tilbake til Tromsø Helseråd i den utleverte konvolutt. Alle opplysninger i forbindelse med Helseundersøkelsen vil bli behandlet strengt konfidensielt. Har De noen kommentarer til undersøkelsen kan De skrive dem i kommentarfeltet på siste side.

Med hilsen

Tromsø Helseråd Fagområdet medisin

HELSETILSTAND	
Hvordan er Deres helsetilstand?	
Sett kryss i den ruten der «Ja» passer best.	
Meget dårlig 12	<input type="checkbox"/> 1
Dårlig 2	<input type="checkbox"/> 2
Hverken god eller dårlig, middels 3	<input type="checkbox"/> 3
Bra 4	<input type="checkbox"/> 4
Utmerket 5	<input type="checkbox"/> 5
SYKDOM	
Har De, eller har De hatt	
Kryss av «Ja» eller «Nei» for hvert spørsmål	
Hudsykdommen psoriasis 13	<input type="checkbox"/> Ja <input type="checkbox"/> Nei
Astma 14	<input type="checkbox"/> <input type="checkbox"/>
Allergisk eksem 15	<input type="checkbox"/> <input type="checkbox"/>
Høysnue 16	<input type="checkbox"/> <input type="checkbox"/>
Kronisk bronkitt 17	<input type="checkbox"/> <input type="checkbox"/>
Sår på magesekken 18	<input type="checkbox"/> <input type="checkbox"/>
Sår på tolvfingertarmen 19	<input type="checkbox"/> <input type="checkbox"/>
Blindtarms-operasjon 20	<input type="checkbox"/> <input type="checkbox"/>
Magesårs-operasjon 21	<input type="checkbox"/> <input type="checkbox"/>
Leddgikt (kronisk revmatoid artritt) 22	<input type="checkbox"/> <input type="checkbox"/>
Kreftsykdom 23	<input type="checkbox"/> <input type="checkbox"/>
Epilepsi (fallesyke) 24	<input type="checkbox"/> <input type="checkbox"/>
Migrene 25	<input type="checkbox"/> <input type="checkbox"/>
INFEKSJON	
Hvor mange ganger har De hatt infeksjon slik som forkjølelse, influensa, «ræksjuka» og lignende siste halvår? 26	
Antall <input type="checkbox"/>	
Har De hatt slik infeksjon siste 14 dager? 27	<input type="checkbox"/> Ja <input type="checkbox"/> Nei

SYKDOM HOS FORELDRE OG SØSKEN	
Kryss av for de slektningene som har eller har hatt noen av sykdommene:	
Hjerneslag eller hjerneblødning 28	<input type="checkbox"/> Mor <input type="checkbox"/> Far <input type="checkbox"/> Bror <input type="checkbox"/> Søster
Sukkersyke 32	<input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/>
Leddgikt (revmatoid artritt) 36	<input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/>
Kreft 40	<input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/>
Psoriasis 44	<input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/>
Magesår eller tolvfingertarmsår 48	<input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/>
Astma 52	<input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/> <input type="checkbox"/>
Kryss av dersom slektningene ikke har eller har hatt noen av disse sykdommene 56	
<input type="checkbox"/> Ja <input type="checkbox"/> Nei	
MEDISINER	
Har De siste år brukt tabletter, sprøyter eller astmaspray mot astma eller allergi 60	
<input type="checkbox"/> Ja <input type="checkbox"/> Nei	
Har De brukt følgende medisiner siste 14 dager?	
Smertestillende 61	<input type="checkbox"/> Ja <input type="checkbox"/> Nei
Febersenkende 62	<input type="checkbox"/> <input type="checkbox"/>
Eksemsalve 63	<input type="checkbox"/> <input type="checkbox"/>
Blodtrykksmedisin 64	<input type="checkbox"/> <input type="checkbox"/>
Hjertemedisin 65	<input type="checkbox"/> <input type="checkbox"/>
Sovemedisin 66	<input type="checkbox"/> <input type="checkbox"/>
Nervemedisin 67	<input type="checkbox"/> <input type="checkbox"/>
Migrenemedisin 68	<input type="checkbox"/> <input type="checkbox"/>
Medisin mot epilepsi (fallesyke) 69	<input type="checkbox"/> <input type="checkbox"/>
Annen medisin 70	<input type="checkbox"/> <input type="checkbox"/>

KONTAKT PGA. EGEN HELSE ELLER SYKDOM

Hvor mange besøk har De hatt siste år på grunn av egen helse eller sykdom?

Hos vanlig lege 71
 Hos spesialist utenfor sykehuset 72
 På legevakta 85
 Hos bedriftslege 87
 Hos fysioterapeut 89
 Hos kiropraktor 81
 Hos naturmedisinere (homeopat, soneterapeut o.l.) 83
 På sykehusets poliklinikk 85

Antall innleggelse på sykehus siste år 87

Antall besøk

KOSTHOLD

Hvor mange brødskeer spiser De vanligvis daglig?

Sett kryss i den ruten der «Ja» passer best

Mindre enn 2 skiver 88
 2 - 4 skiver
 5 - 6 skiver
 7 - 12 skiver
 13 eller flere skiver

Ja
 1
 2
 3
 4
 5

Hva slags melk drikker de vanligvis?

Sett kryss i den ruten der «Ja» passer best.

Drikker ikke melk 89
 Melk (helmelk), søt, sur
 Lettmelk
 Skummet melk, søt, sur

Ja
 1
 2
 3
 4

Hvor mange glass/kopper melk drikker De vanligvis daglig?

Mindre enn ett glass/kopp 90
 1 - 2 glass/kopper
 3 - 4 glass/kopper
 5 eller flere glass/kopper

Ja
 1
 2
 3
 4

FISKEMAT

Hvor ofte spiser De torsk/sei eller annen mager fisk til middag eller som pålegg?

Sett kryss i den ruten der «Ja» passer best.

Sjeldnere enn en gang i uken 91
 1 gang i uken
 2 ganger i uken
 3 eller flere ganger i uken

Ja
 1
 2
 3
 4

Hvor ofte spiser De fet fisk som sild, kveite, uer, makrell, laks, ørret til middag eller som pålegg?

Sett kryss i ruten der «Ja» passer best.

Sjeldnere enn en gang i uken 92
 1 gang i uken
 2 ganger i uken
 3 eller flere ganger i uken

Ja
 1
 2
 3
 4

Bruker De tran regelmessig?

Sett kryss i ruten der «Ja» passer best.

Nei 93
 I mørketida
 Hele året

Ja
 1
 2
 3

FROKOST

Spiser De vanligvis frokost daglig? 94

Ja Nei

MIDDAGSMAT

Hvor ofte spiser De vanligvis kjøtt til middagen?

Sett kryss i ruten der «Ja» passer best.

Sjeldnere enn en gang i uken 95
 1 - 2 ganger i uken
 3 - 4 ganger i uken
 5 eller flere ganger i uken

Ja
 1
 2
 3
 4

Hvor ofte bruker De fett (smør, margarin, remulade, majones og lignende) til eller på middagsmaten?

Sett kryss i ruten der «Ja» passer best.

Sjeldnere enn en gang i uken 96
 1 - 2 ganger i uken
 3 - 4 ganger i uken
 5 eller flere ganger i uken

Ja
 1
 2
 3
 4

Bruker De vanligvis grønnsaker som del av middagsmaten? 97

Ja Nei

FRUKT

Hvor ofte spiser De vanligvis frukt?

Sett kryss i ruten der «Ja» passer best.

Sjeldnere enn en gang i uken 98
 Omtrent en gang i uken
 2 - 3 ganger i uken
 4 - 5 eller flere ganger i uken
 Omtrent daglig

Ja
 1
 2
 3
 4
 5

ALKOHOL

Er De total avholdsmann/-kvinne

Ja Nei

Hvis nei, - Hvor ofte pleier De å drikke øl?

Sett kryss i ruten der «Ja» passer best.

Aldri, eller noen få ganger i året 100
 1 - 2 ganger i måneden
 Omtrent 1 gang i uken
 2 - 3 ganger i uken
 Omtrent hver dag

Ja
 1
 2
 3
 4
 5

Hvor ofte pleier De å drikke vin?

Sett kryss i ruten der «Ja» passer best.

Aldri, eller noen få ganger i året 101
 1 - 2 ganger i måneden
 Omtrent 1 gang i uken
 2 - 3 ganger i uken
 Omtrent hver dag

Ja
 1
 2
 3
 4
 5

- Hvor ofte pleier De å drikke brennevin?

Sett kryss i ruten der «Ja» passer best.

Aldri, eller noen få ganger i året 102
 1 - 2 ganger i måneden
 Omtrent 1 gang i uken
 2 - 3 ganger i uken
 Omtrent hver dag

Ja
 1
 2
 3
 4
 5

Omtrent hvor ofte har De i løpet av siste år drukket alkohol tilsvarende minst 5 halvflasker ol, en helflaske vin eller ¼ flaske brennevin?

Sett kryss i ruten der «Ja» passer best.

Ikke siste år 103
 Noen få ganger
 1 - 2 ganger i måneden
 3 eller flere ganger i uken

Ja
 1
 2
 3
 4

FYSISK AKTIVITET

Hvor ofte utfører De fysisk aktivitet av minst 20 minutters varighet og som fører til at De blir svett eller andpusten?

Sett kryss i ruten der «Ja» passer best.

- | | | | | |
|---------------------|-----|----|--------------------------|---|
| Sjelden eller aldri | 104 | Ja | <input type="checkbox"/> | 1 |
| Ukentlig | | | <input type="checkbox"/> | 2 |
| Flere ganger i uka | | | <input type="checkbox"/> | 3 |
| Daglig | | | <input type="checkbox"/> | 4 |

Dersom De vanligvis utfører slik aktivitet minst en gang i uka, hvor mye tid bruker De ukentlig til slik aktivitet?

Sett kryss i ruten der «Ja» passer best.

- | | | | | |
|------------------------------------|-----|----|--------------------------|---|
| Mindre enn 30 minutter i uka | 105 | Ja | <input type="checkbox"/> | 1 |
| Mellom 30 minutter og 1 time i uka | | | <input type="checkbox"/> | 2 |
| Mellom 1 og 2 timer i uka | | | <input type="checkbox"/> | 3 |
| Mer enn 2 timer i uka | | | <input type="checkbox"/> | 4 |

VANE- OG KOSTENDRINGER

Har De endret Deres vaner/kosthold i løpet av de siste 5 år når det gjelder:(Sett kryss for hvert spørsmål)

- | | | | | | |
|-----------------------------|-----|----|--------------------------|-----|--------------------------|
| Fett i kosten | 106 | Ja | <input type="checkbox"/> | Nei | <input type="checkbox"/> |
| Soyamargarin eller matoljer | 107 | | <input type="checkbox"/> | | <input type="checkbox"/> |
| Skummet melk eller lettmeik | 108 | | <input type="checkbox"/> | | <input type="checkbox"/> |
| Kaffe-forbruk | 109 | | <input type="checkbox"/> | | <input type="checkbox"/> |
| Alkohol-forbruk | 110 | | <input type="checkbox"/> | | <input type="checkbox"/> |
| Fysisk aktivitet | 111 | | <input type="checkbox"/> | | <input type="checkbox"/> |

EKTESKAPS-/SAMBO-FORHOLD

Er De gift eller samboende

Ja Nei

Hvor gammel var De da De første gang giftet Dem eller innledet et samboerforhold?

år

HUSSTAND

Hvor mange personer bor det i deres husstand?

Antall

Er noen i Deres husstand 10 år eller yngre?

Ja Nei

Trenger noen i Deres husstand spesielt tilsyn/pleie - utenom barna?

Ja Nei

SKOLEGANG

Hvor mange års skolegang har De (ta også med folkeskole og ungdomsskole)?

år

ARBEID

Har De hatt lønnet arbeid hele siste år?

Sett kryss i ruten der «Ja» passer best.

- | | | | | |
|--------------------|-----|----|--------------------------|---|
| Fulltidsarbeid | 121 | Ja | <input type="checkbox"/> | 1 |
| Deltidsarbeid | | | <input type="checkbox"/> | 2 |
| Ikke lønnet arbeid | | | <input type="checkbox"/> | 3 |

Hvor stor del av det daglige arbeid i hjemmet gjør De vanligvis selv?

Sett kryss i ruten der «Ja» passer best.

- | | | | |
|-------------------------|-----|--------------------------|---|
| Alt eller nesten alt | 122 | <input type="checkbox"/> | 1 |
| Minst halvparten | | <input type="checkbox"/> | 2 |
| Mer enn en fjerdedel | | <input type="checkbox"/> | 3 |
| Mindre enn en fjerdedel | | <input type="checkbox"/> | 4 |

RYGG- OG LEDDPLAGER

Har De i løpet av siste år vært plaget av smerter i ryggen som har vart lenger enn 4 uker?

Ja Nei

Har De vært plaget av stivhet i ryggen om morgenen som varte lenger enn 30 minutter?

Har De i løpet av siste 3 år vært plaget av smerter i noen av de følgende ledd i mer enn 3 måneder?

Ja Nei

- | | | | |
|---------------------------|-----|--------------------------|--------------------------|
| Kneleddene | 126 | <input type="checkbox"/> | <input type="checkbox"/> |
| Albueleddene | 127 | <input type="checkbox"/> | <input type="checkbox"/> |
| De innerste fingerleddene | 128 | <input type="checkbox"/> | <input type="checkbox"/> |
| Andre ledd | 129 | <input type="checkbox"/> | <input type="checkbox"/> |

Hvis ja, merket De stivhet i leddene om morgenen av mer enn 30 minutters varighet?

PLAGER I HODE, NAKKE OG SKULDRE

Hvor ofte er De plaget av hodepine?

Sett kryss i ruten der «Ja» passer best.

Ja

- | | | | |
|---------------------------------|-----|--------------------------|---|
| Sjelden eller aldri | 131 | <input type="checkbox"/> | 1 |
| En eller flere ganger i måneden | | <input type="checkbox"/> | 2 |
| En eller flere ganger i uken | | <input type="checkbox"/> | 3 |
| Daglig | | <input type="checkbox"/> | 4 |

Hvor ofte er De plaget av smerter i nakke eller skuldre?

Sett kryss i ruten der «Ja» passer best.

Ja

- | | | | |
|---------------------------------|-----|--------------------------|---|
| Sjelden eller aldri | 132 | <input type="checkbox"/> | 1 |
| En eller flere ganger i måneden | | <input type="checkbox"/> | 2 |
| En eller flere ganger i uken | | <input type="checkbox"/> | 3 |
| Daglig | | <input type="checkbox"/> | 4 |

Reduserer plagene i hodet, nakken eller skuldrene Deres arbeidsevne?

Sett kryss i ruten der «Ja» passer best.

Ja

- | | | | |
|--------------------------------|-----|--------------------------|---|
| Aldri, eller i ubetydelig grad | 133 | <input type="checkbox"/> | 1 |
| I noen grad | | <input type="checkbox"/> | 2 |
| I betydelig grad | | <input type="checkbox"/> | 3 |
| Klarer ikke vanlig arbeid | | <input type="checkbox"/> | 4 |

Har De noen gang fått røntgenundersøkt ryggen, nakken og/eller skuldre?

Ja Nei

SØVNLØSHET/BEVISSTLØSHET

Hender det at De er plaget av søvnløshet?

Ja Nei

Hvis ja, når på året er De mest plaget?

Sett kryss i ruten der «Ja» passer best.

Ja

- | | | | |
|------------------------|-----|--------------------------|---|
| Ingen spesiell tid | 136 | <input type="checkbox"/> | 1 |
| Særlig mørketiden | | <input type="checkbox"/> | 2 |
| Særlig i midnattstiden | | <input type="checkbox"/> | 3 |
| Særlig høst og vår | | <input type="checkbox"/> | 4 |

Har De gjennom hele siste år vært plaget av søvnløshet slik at det går ut over arbeidsevnen?

Ja Nei

Har De siste år hatt anfall med plutselig tap av bevissthet?

Ja Nei

Har De merket anfall med plutselig endring i pulsen eller hjerterytmen siste år?

Ja Nei

REAKSJONER PÅ PROBLEMER			
Hvis De får store personlige problemer, regner De da med å få hjelp og støtte fra ektefelle, samboer eller familie? 140	Ja <input type="checkbox"/> Nei <input type="checkbox"/>	Har De i de siste 14 dager følt Dem ulykkelig og nedtrykt (deprimert)? Sett kryss i ruten der «Ja» passer best.	Ja <input type="checkbox"/>
Har De i lengere tid følt behov for å oppsøke noen på grunn av personlige problem siste år, uten at De har tatt slik kontakt? 141	Ja <input type="checkbox"/> Nei <input type="checkbox"/>	Aldri eller sjelden 143	<input type="checkbox"/> 1
Har De i de siste 14 dager følt Dem ute av stand til å takle Deres vanskeligheter? Sett kryss i ruten der «Ja» passer best.	Ja <input type="checkbox"/>	Av og til	<input type="checkbox"/> 2
Aldri eller sjelden 142	<input type="checkbox"/> 1	Ofte	<input type="checkbox"/> 3
Av og til	<input type="checkbox"/> 2	Nesten hele tida	<input type="checkbox"/> 4
Ofte	<input type="checkbox"/> 3	Hender det ofte at De føler Dem ensom? Sett kryss i ruten der «Ja» passer best.	Ja <input type="checkbox"/>
Nesten hele tida	<input type="checkbox"/> 4	Meget ofte 144	<input type="checkbox"/> 1
		Av og til	<input type="checkbox"/> 2
		Aldri eller nesten aldri	<input type="checkbox"/> 3
RESTEN AV SKJEMAET BESVARES BARE AV KVINNER			
MENSTRUASJON Hvor gammel var De da De fikk menstruasjon første gang? 145	<input type="checkbox"/> år	Forsvinner plagene når menstruasjonen kommer? 160	Ja <input type="checkbox"/> Nei <input type="checkbox"/>
Når begynte Deres siste menstruasjon? 147	dag mnd. år / /	Bruker De mot slike plager: - vanndrivende tabletter? 161	Ja <input type="checkbox"/> Nei <input type="checkbox"/>
Hvor mange dager er/var det vanligvis fra menstruasjonens 1. blødningsdag til neste menstruasjons 1. blødningsdag (= tiden mellom to menstruasjoners begynnelse)? 153	<input type="checkbox"/> dager	- andre medisiner? 162	<input type="checkbox"/> <input type="checkbox"/>
Pleier/pleide menstruasjonen å være regelmessig 155	Ja <input type="checkbox"/> Nei <input type="checkbox"/>	SVANGERSKAP	Antall <input type="checkbox"/>
Bruker De vanligvis smertestillende tabletter under menstruasjonen? 156	Ja <input type="checkbox"/> Nei <input type="checkbox"/>	Hvor mange barn har De født? 163	<input type="checkbox"/>
PLAGER FOR MENSTRUASJON		Hvor gammel var De første gang De var gravid? 164	<input type="checkbox"/> år
Har De før menstruasjon noen av disse plagene: - Er De nedtrykt (deprimert) eller irritable? Sett kryss i ruten der «Ja» passer best	Ja <input type="checkbox"/>	PREVENSJON	Ja <input type="checkbox"/> Nei <input type="checkbox"/>
Ubetydelig 157	<input type="checkbox"/> 1	Bruker eller har De brukt P-piller eller spiral? 166	<input type="checkbox"/> <input type="checkbox"/>
Merkbart	<input type="checkbox"/> 2	Hvis ja, hvor mange år har De tilsammen brukt: P-piller? 167	<input type="checkbox"/> år
Plagsomt	<input type="checkbox"/> 3	Spiral? 169	<input type="checkbox"/> år
- Har De smertefulle bryster? Sett kryss i ruten der «Ja» passer best	Ja <input type="checkbox"/>	Hvor gammel var De da De begynte med: P-piller? 171	<input type="checkbox"/> år
Ubetydelig 158	<input type="checkbox"/> 1	Spiral? 173	<input type="checkbox"/> år
Merkbart	<input type="checkbox"/> 2	Hvis De har sluttet med P-piller, uteble da menstruasjonen i mer enn 6 måneder uten at De var gravid? 175	Ja <input type="checkbox"/> Nei <input type="checkbox"/>
Plagsomt	<input type="checkbox"/> 3	Har de måttet slutte med P-piller fordi De fikk høyt blodtrykk? 176	Ja <input type="checkbox"/> Nei <input type="checkbox"/>
- Har De hovne hender/føtter, vektøkning, eller følelse av å «ese ut»? Sett kryss i ruten der «Ja» passer best	Ja <input type="checkbox"/>	KREFTPROVE	Antall prøver <input type="checkbox"/>
Ubetydelig 159	<input type="checkbox"/> 1	Hvor mange ganger har De fått tatt kreftprøve (celleprøve) fra livmorhalsen siste 3 år? 177	<input type="checkbox"/>
Merkbart	<input type="checkbox"/> 2	Hvor mange år siden er det siden siste prøve? 178	<input type="checkbox"/> år
Plagsomt	<input type="checkbox"/> 3		
Deres kommentarer: 179			

Takk for hjelpen! Husk å postlegge skjemaet idag!
 Tromsundersøkelsen 1986-7

ISM SKRIFTSERIE - FØR UTGITT:

1. Bidrag til belysning av medisinske og sosiale forhold i Finnmark fylke, med særlig vekt på forholdene blant finskattede i Sør-Varanger kommune.
Av Anders Forsdahl, 1976. (nytt opplag 1990)
2. Sunnhetstilstanden, hygieniske og sosiale forhold i Sør-Varanger kommune 1869-1975 belyst ved medisinalberetningene.
Av Anders Forsdahl, 1977.
3. Hjerter-karundersøkelsen i Finnmark - et eksempel på en populasjonsundersøkelse rettet mot cardiovasculære sykdommer. Beskrivelse og analyse av etterundersøkelsesgruppen.
Av Jan-Ivar Kvamme og Trond Haider, 1979.
4. The Tromsø Heart Study: Population studies of coronary risk factors with special emphasis on high density lipoprotein and the family occurrence of myocardial infarction.
Av Olav Helge Førde og Dag Steinar Thelle, 1979.
5. Reformen i distriktshelsetjenesten III: Hypertensjon i distriktshelsetjenesten.
Av Jan-Ivar Kvamme, 1980.
6. Til professor Knut Westlund på hans 60-års dag, 1983.
- 7.* Blodtrykksovervåkning og blodtrykksmåling.
Av Jan-Ivar Kvamme, Bernt Nesje og Anders Forsdahl, 1983.
- 8.* Merkesteiner i norsk medisin reist av allmennpraktikere - og enkelte utdrag av medisinalberetninger av kulturhistorisk verdi.
Av Anders Forsdahl, 1984.
9. "Balsfjordsystemet." EDB-basert journal, arkiv og statistikk-system for primærhelsetjenesten.
Av Toralf Hasvold, 1984.
10. Tvunget psykisk helsevern i Norge. Rettsikkerheten ved slikt helsevern med særlig vurdering av kontrollkommisjonsordningen.
Av Georg Høyer, 1986.

11. The use of self-administered questionnaires about food habits. Relationships with risk factors for coronary heart disease and associations between coffee drinking and mortality and cancer incidence.
Av Bjarne Koster Jacobsen, 1988.
- 12.* Helse og ulikhet. Vi trenger et handlingsprogram for Finnmark.
Av Anders Forsdahl, Atle Svendal, Aslak Syse og Dag Thelle, 1989.
13. Health education and self-care in dentistry - surveys and interventions.
Av Anne Johanne Søgaard, 1989.
14. Helsekontroller i praksis. Erfaringer fra prosjektet helsekontroller i Troms 1983-1985.
Av Harald Siem og Arild Johansen, 1989.
15. Til Anders Forsdahls 60-års dag, 1990.
16. Diagnosis of cancer in general practice. A study of delay problems and warning signals of cancer, with implications for public cancer information and for cancer diagnostic strategies in general practice.
Av Knut Holtedahl, 1991.
17. The Tromsø Survey. The family intervention study. Feasibility of using a family approach to intervention on coronary heart disease. The effect of lifestyle intervention of coronary risk factors.
Av Synnøve Fønne Knutsen, 1991.
18. Helhetsforståelse og kommunikasjon. Filosofi for klinikere.
Av Åge Wifstad, 1991.
19. Factors affecting self-evaluated general health status - and the use of professional health care services.
Av Knut Fylkesnes, 1991.
20. Serum gamma-glutamyltransferase: Population determinants and diagnostic characteristics in relation to intervention on risk drinkers.
Av Odd Nilssen, 1992.

21. The Healthy Faith. Pregnancy outcome, risk of disease, cancer morbidity and mortality in Norwegian Seventh-Day-Adventists.
Av Vinjar Fønnebo, 1992.
22. Aspects of breast and cervical cancer screening.
Av Inger Torhild Gram, 1992.
23. Population studies on dyspepsia and peptic ulcer disease: Occurrence, aetiology, and diagnosis. From The Tromsø Heart Study and The Sørreisa Gastrointestinal Disorder Studie.
Av Roar Johnsen, 1992.
24. Diagnosis of pneumonia in adults in general practice.
Av Hasse Melbye, 1992.
25. Relationship between hemodynamics and blood lipids in population surveys, and effects of n-3 fatty acids.
Av Kaare Bønaa, 1992.
26. Risk factors for, and 13-year mortality from cardiovascular disease by socioeconomic status. A study of 44690 men and 17540 women, ages 40-49.
Av Hanne Thürmer, 1993.
27. Utdrag av medisinalberetninger fra Sulitjelma 1891-1990.
Av Anders Forsdahl, 1993.
28. Helse, livsstil og levekår i Finnmark. Resultater fra Hjerte-karundersøkelsen i 1987-88. Finnmark III.
Av Knut Westlund og Anne Johanne Søgaard, 1993.
29. Patterns and predictors of drug use. A pharmacoepidemiologic study, linking the analgesic drug prescriptions to a population health survey in Tromsø, Norway.
Av Anne Elise Eggen, 1994.
30. ECG in health and disease. ECG findings in relation to CHD risk factors, constitutional variables and 16-year mortality in 2990 asymptomatic Oslo men aged 40-49 years in 1972.
Av Per G. Lund-Larsen, 1994.

31. Arrhythmia, electrocardiographic signs, and physical activity in relation to coronary heart risk factors and disease. The Tromsø Study.
Av Maja-Lisa Løchen, 1995.
32. The Military service: mental distress and changes in health behaviours among Norwegian army conscript.
Av Edvin Schei, 1995.
33. The Harstad injury prevention study: Hospital-based injury recording and community-based intervention.
Av Børge Ytterstad, 1995.
- 34.* Vilkår for begrepsdannelse og praksis i psykiatri. En filosofisk undersøkelse.
Av Åge Wifstad, 1996. (utgitt Tano Aschehoug forlag 1997)
35. Dialog og refleksjon. Festskrift til professor Tom Andersen på hans 60-års dag, 1996.
36. Factors affecting doctors' decision making.
Av Ivar Sønbo Kristiansen, 1996.
37. The Sørreisa gastrointestinal disorder study. Dyspepsia, peptic ulcer and endoscopic findings in a population.
Av Bjørn Bernersen, 1996.
38. Headache and neck or shoulder pain. An analysis of musculoskeletal problems in three comprehensive population studies in Northern Norway.
Av Toralf Hasvold, 1996.
39. Senfølger av kjernefysiske prøvespreninger på øygruppen Novaya Semlya i perioden 1955 til 1962. Rapport etter programmet "Liv". Arkangelsk 1994.
Av A.V. Tkatchev, L.K. Dobrodeeva, A.I. Isaev, T.S. Podjakova, 1996.
40. Helse og livskvalitet på 78 grader nord. Rapport fra en befolkningsstudie på Svalbard høsten 1988.
Av Helge Schirmer, Georg Høyer, Odd Nilssen, Tormod Brenn og Siri Steine, 1997.

De som er merket med * har vi dessverre ikke flere eksemplarer av.