

ORIGINAL ARTICLE

Movements of the wrist and the risk of carpal tunnel syndrome: a nationwide cohort study using objective exposure measurements

Christina Bach Lund, ¹ Sigurd Mikkelsen, ¹ Lau Caspar Thygesen, ² Gert-Åke Hansson, ^{3,4} Jane Frølund Thomsen ¹

► Additional material is published online only. To view please visit the journal online (http://dx.doi.org/10.1136/ oemed-2018-105619).

¹Department of Occupational and Environmental Medicine, Bispebjerg Hospital, Copenhagen, Denmark ²National Institute of Public Health, University of Southern Denmark, Copenhagen, Denmark ³Occupational and Environmental Medicine, University and Regional Laboratories Region Scania, Lund, Sweden ⁴Division of Occupational and Environmental Medicine, Lund University, Lund, Sweden

Correspondence to

Christina Bach Lund, Depatment of Occupational and Environmental Medicine, Bispebjerg Hospital, Copenhagen 2400, Denmark; bach.christina@gmail.com

Received 4 December 2018 Revised 17 May 2019 Accepted 28 May 2019 Published Online First 12 June 2019

ABSTRACT

Objectives We conducted a large cohort study to investigate the association between work-related wrist movements and carpal tunnel syndrome (CTS).

Methods Electro-goniometric measurements of wrist movements were performed for 30 jobs (eg, office work, child care, laundry work and slaughterhouse work). We measured wrist angular velocity, mean power frequency (MPF) and range of motion (ROM). We established a cohort of Danish citizens born 1940-1979 who held one of these jobs from age 18-80 years, using Danish national registers with annual employment information from 1992 to 2014. We updated the cohort by calendar year with job-specific and sex-specific means of measured exposures. Dates of a first diagnosis or operation because of CTS were retrieved from the Danish National Patient Register. The risk of CTS by guintiles of preceding exposure levels was assessed by adjusted incidence rate ratios (IRR^{adj}) using Poisson regression models.

Results We found a clear exposure—response association between wrist angular velocity and CTS with an IRR^{adj} of 2.31 (95% CI 2.09 to 2.56) when exposed to the highest level compared with the lowest. MPF also showed an exposure—response pattern, although less clear, with an IRR^{adj} of 1.83 (1.68 to 1.98) for the highest compared with the lowest exposure level. ROM showed no clear pattern. Exposure—response patterns were different for men and women.

Conclusions High levels of wrist movement were associated with an increased risk of CTS. Preventive strategies should be aimed at jobs with high levels of wrist movements such as cleaning, laundry work and slaughterhouse work.

INTRODUCTION



© Author(s) (or their employer(s)) 2019. Re-use permitted under CC BY-NC. No commercial re-use. See rights and permissions. Published by BMJ.

To cite: Lund CB, Mikkelsen S, Thygesen LC, et al. Occup Environ Med 2019;**76**:519–526. Carpal tunnel syndrome (CTS) has consistently been shown to be associated with work-related biomechanical factors. The specific pathogenesis is not clearly understood, but symptoms relate to compression of the median nerve at the wrist, and several studies have demonstrated an increased pressure in the carpal tunnel when performing hand straining tasks. Symptoms include pain and paraesthesia of the three radial fingers and the radial part of the fourth finger and may involve atrophy of the thenar muscles. In severe cases, the ability to perform manual work may be compromised.

Key messages

What is already known about this subject?

Previous studies have found an association between work-related repetitive work and carpal tunnel syndrome, but these studies have often been limited by a cross-sectional study design and self-reported outcome and exposure assessment.

What are the new findings?

▶ This study found a clear exposure—response relationship between wrist angular velocity and the risk of carpal tunnel syndrome, identified through register-based diagnoses, in a large population-based cohort with exposures based on technical measurements within 30 different jobs.

How might this impact on policy or clinical practice in the foreseeable future?

➤ The results of this study suggest that preventive strategies should be aimed at jobs with high levels of wrist movements such as cleaners, laundry workers and slaughterhouse workers.

CTS is the most common nerve entrapment disorder and a common cause of work absence and private earning losses.² In a large Swedish general population-based study (25–74 years) based on both clinical and electrophysiological diagnosis of CTS, the overall prevalence was 2.1% (95% CI 1.3% to 3.0%) among men and 3.0% (2.1% to 3.9%) among women.³ Pregnancy, wrist fractures, hypothyroidism, rheumatoid arthritis, diabetes mellitus and obesity are known risk factors.^{4 5} The best documented work-related risk factors are repetition, force and hand–arm vibration.^{6 7}

Self-reported exposure is widely used in studies on biomechanical risk factors in CTS increasing the risk of reporting bias compared with objectively assessed exposure. So Several studies included job titles or job function to define exposure, but both are difficult to interpret in terms of specific risk factors. Some studies have used expert ratings and observation, either alone or in combination, and some have used video analyses or technical measurements. Cross-sectional designs dominate the studies investigating biomechanical risk factors in

CTS, but longitudinal designs have also been used.^{6 7 16 17} Exposure–response relationships for force and repetition has been determined in studies using observed measures^{6 7}; however, exposure–response relationships for force and repetition in men and women based on technically obtained exposure measures have not yet been determined. To our knowledge, no large prospective cohort studies based on technical measurements within several different jobs have been conducted.

The aim of this study was to investigate the associations between movements of the wrist and the risk of CTS in a large prospective study, using representative whole day electro-goniometrical measurements of wrist movements in 30 specific jobs to assess exposure in a national register-based cohort of persons who ever held any of these jobs.

METHODS

Exposure measurements in specific jobs

Whole day measurements of movements and position of the wrist were performed within 33 different jobs, classified by their occupational title according to the Danish version of the International Standard Classification of Occupations (DISCO). The majority of jobs (n=23) was measured in Denmark (Copenhagen and Aarhus) as part of two previous studies of work-related musculoskeletal upper extremity disorders. Ten jobs were measured in earlier studies in Sweden and Norway and were part of an existing job exposure database at the Department of Occupational and Environmental Medicine, University of Lund, Sweden. The methods were the same for all measurements and were supervised by one of the authors (G-ÅH). For each job, a description of the work tasks performed during the measurements was recorded. Only healthy, right-handed individuals with no musculoskeletal complaints were measured.

The jobs were chosen to represent large exposure contrasts and jobs of both sexes. We aimed to obtain measurements of: 10 men in male-dominated jobs, 10 women in female-dominated jobs and 10 men *and* 10 women in jobs with a similar proportion of both men and women. House painters, cleaners and office workers had measurements performed as part of other studies and therefore had more measurements performed.^{15 21}

Wrist measurements were performed using biaxial goniometers (SG75, Biometrics, LTD, Newport, UK), placed on the dorsal side of the right and left wrist with the proximal part in the midline between radius and ulna and the distal part over the third metacarpal bone, and data were recorded by personworn data loggers (Logger Teknologi HB, Åkarp, Sweden) with a sampling frequency of 20 Hz for a full work day, excluding time for instruction, mounting and dismounting the equipment. The mean measuring time was 5.5 hours (3.1-6.9). Analyses were made bilaterally. For each person, the flexion/extension median angular velocity (°/s) and the mean power frequency (MPF) (Hz) was calculated as a measure of wrist movements. Furthermore, the 10th and 90th percentile of the wrist angular deviation from the neutral position was calculated and used the difference between these percentiles as a measure of range of motion (ROM) (°). For each job and sex, the means of these measures were calculated to represent the group and sex average angular velocity, MPF and ROM to create a job exposure matrix for subsequent use in epidemiological studies.

Study population

From the national Danish Civil Registration System, ²² we established a cohort of persons born in Denmark between 1 January 1940 and 31 December 1979 including information on sex, date

of birth, immigration, emigration and death. Each person was identified by their unique Danish personal identification number allowing linkage to personal information in other national registers in Statistics Denmark. We retrieved individual annual information from 1 January 1992 to 1 January 2015 on their main occupational title by DISCO codes, the industry where the person worked, coded by the Danish Industrial Classification of All Economic Activities, ²³ and the highest fulfilled educational level coded by the Danish version of the International Standard Classification of Education. ²⁴

We used this information on occupational title, industry and education to identify persons who had ever held a job corresponding to the previously established job matrix of 33 jobs with measurements. Since each DISCO code can refer to several job titles, industry and education codes were in some cases used to identify the specific job. The specific codes used for each job are listed in online supplementary appendix 1. We excluded three jobs (assembly worker, packing worker and wood parquet industry worker) that could not be defined with sufficient accuracy in the national registers, thus 30 jobs were finally included.

We retrieved information on the proportion of time with salaried work in each calendar year, defining the job-specific exposure time for each calendar year in the 30 jobs. The annual proportions of working time were multiplied with the job-specific exposure for angular velocity, MPF and ROM, creating an annual measure of exposure level (duration*intensity) for that calendar year for each of the three exposures. We further calculated total cumulative exposures for each calendar year by adding the exposure of all previous years. We excluded persons without information on annual proportion of working time and persons who immigrated after 1 January 1992, if they were more than 18 years old at immigration (figure 1).

Outcome and health-related confounders

The Danish National Patient Register (DNPR) holds information about all patient contacts to all Danish hospitals since 1977.²⁵ Cases were identified in DNPR by primary CTS diagnosis or CTS operation. Diagnoses were coded by the International Classification of Diseases (ICD) 8th (ICD-8) (1971-1993) and 10th (ICD-10)(1994-2014) revisions, respectively, and CTS operations were coded by a specific Danish classification (1977-1995) and the NOMESCO Classification of Surgical Procedures (NCSP) (1996–2014), respectively.²⁶ We used ICD-8 code 357.99 and ICD-10 code G56.0 to identify CTS diagnoses, and the specific Danish classification code 3680 and NCSP codes KACC51 and KACC61 to identify CTS operations. Both diagnosis and operation codes were hospital discharge codes. Similar information was recorded for pregnancies, wrist-near fractures, hypothyroidism, rheumatoid arthritis, diabetes mellitus and obesity (see online supplementary appendix 2). Pregnancy was defined as 7 months prior to and 5 months after giving birth. Date of occurrence of hypothyroidism, rheumatoid arthritis, diabetes and obesity was set to 1 January the year of diagnosis and they were included as risk factors from that year and onwards.

Analysis

The cohort was followed from the first calendar year the person entered one of the 30 jobs if they were at least ≥18 years of age until the date of first-time CTS, emigration, death or 1 January 2015, whichever came first. We excluded persons with a CTS diagnosis or CTS operation prior to start of follow-up (figure 1). Each person obtained risk time from 1 January the year after they entered the cohort until the end of follow-up

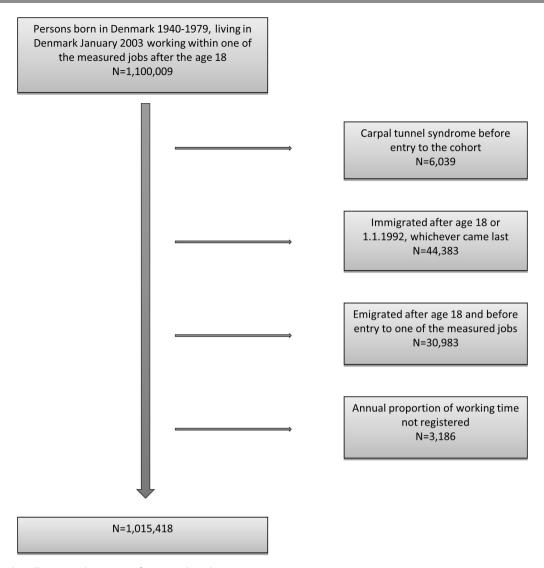


Figure 1 Flow chart illustrating the process of creating the cohort.

time. Risk time was divided into age-specific risk time, based on the proportion of time before and after the date of birth using the Lexis-SAS macro.²⁷ In case that a wrist-near fracture had occurred, the risk time was split into risk time before and after the date of fracture and was then included as a risk factor for the rest of follow-up. In case of pregnancy, risk time was similarly split by dates of start and end of pregnancy. Regarding all other health-related confounders, risk time was split 1 January the year of the diagnosis.

Poisson regression models were used to examine incidence rate ratios (IRRs) of CTS in relation to angular velocity, MPF and ROM separately.²⁸ A logarithmic transformation of the risk time was used as offset value.

Our main analyses included the 1-year exposure from the previous calendar year (1-year lag) because of the potential risk of change in exposure due to symptoms preceding a CTS diagnosis. Exposure variables were examined by their quintile categories. Persons in jobs with unmeasured exposures in the relevant year were treated as a separate category. As sensitivity analyses, we examined the association with the total cumulative exposure with a 1-year lag to assess if the effect was different for recent exposure and longer lasting cumulative exposure. We also examined if the 1-year lag was sufficient to guard against effects of CTS on changes in exposure by using a 2-year lag instead.

Furthermore, we examined if the results of the main analyses were different for CTS diagnoses with or without operation.

Results were reported as crude and adjusted IRRs. We adjusted for sex, age (18–29, 30–39, 40–49, 50–59, 60–69 and 70–80 years), calendar year (<2000, 2000–2004, 2005–2009 and 2010–2015), pregnancy, wrist-near fracture, hypothyroidism, rheumatoid arthritis, diabetes and obesity. To assess whether exposure effects were different for men and women, we included a multiplicative interaction term between sex and angular velocity, MPF and ROM in the main analyses and further stratified analyses by sex.

Furthermore, we made Poisson regression models including exposure as a natural cubic spline with four knots and the same confounders as in the main adjusted analyses. Exposure quintiles were used as knots.

Data were analysed using SAS statistical software V.9.4 (English), except for the splines, which were made using R Studio version 1.1.456.

RESULTS

A total of 1 015 418 persons were included in the cohort, 43% men. Measured values of angular velocity ranged from 3.55° /s to 37.6° /s, MPF from 0.20 Hz to 0.45 Hz and ROM from 29.8° to

Table 1 Cohort characteristics on the 30 different jobs (eight with both sexes represented). Jobs are arranged according to angular wrist velocity. Number of persons (N) and age at entry are based on the total cohort. Total risk time and number of carpal tunnel syndrome cases are based on the dataset used in the analyses. Exposure intensities of wrist angular velocity, mean power frequency and range of motion are based on the electrogoniometric measurements

			Age at entry,	Total risk time,	CTS cases,		Angular velocity, °/s	MPF, Hz	ROM, °
Job	Sex	N, at entry	mean (SD)	person years	N	N, measured	mean (SD)	mean (SD)	Mean*
Office worker	Men	66 741	33.2 (12.6)	148 535	50	18	3.55 (1.89)	0.20 (0.03)	50.30
Bank assistant	Men	15391	29.9 (9.9)	35 074	4	10	4.31 (1.05)	0.24 (0.03)	43.43
Truck/fork-lift operator	Men	23379	33.8 (11.2)	74 740	65	10	5.60 (2.55)	0.25 (0.06)	29.79
Office worker	Women	254224	34.8 (11.1)	1 182 381	1154	25	6.09 (3.51)	0.23 (0.05)	48.70
Truck driver	Men	29800	36.7 (11.0)	89 059	92	11	6.41 (3.76)	0.24 (0.04)	50.62
Dental hygienist	Women	6815	30.5 (9.5)	27 146	27	12	7.28 (1.61)	0.24 (0.02)	57.53
Electronical worker	Men	6437	35.2 (12.8)	13 229	18	10	8.04 (4.03)	0.24 (0.03)	48.01
Plumber	Men	6941	28.6 (7.7)	51 904	90	11	9.32 (2.22)	0.25 (0.01)	52.77
Smith	Men	42 743	32.1 (13.0)	108 399	129	12	10.21 (2.22)	0.25 (0.03)	51.22
Wood processing industry worker	Men	15 568	31.3 (11.5)	37511	42	10	10.39 (2.97)	0.26 (0.04)	50.64
Childcare worker	Women	72 414	33.0 (8.8)	402 829	554	11	10.72 (2.46)	0.27 (0.03)	63.71
Nurse, aide	Women	75 901	35.1 (10.4)	325 239	688	10	11.14 (3.41)	0.27 (0.03)	59.63
Carpenter	Men	34992	29.9 (9.7)	184 069	243	10	11.60 (2.29)	0.25 (0.04)	54.65
Gardener	Men	4348	30.5 (12.2)	10869	11	11	11.63 (2.43)	0.28 (0.04)	53.36
Cardboard worker	Women	513	30.3 (11.8)	1176	4	10	11.71 (3.85)	0.25 (0.04)	57.45
Electronical worker	Women	10678	34.3 (10.1)	37 155	78	11	12.48 (4.93)	0.27 (0.04)	56.93
Scaffolder	Men	2209	30.4 (8.6)	5913	9	10	12.88 (1.64)	0.30 (0.05)	56.93
Car mechanic	Men	37 020	27.5 (10.6)	175 281	188	10	13.77 (2.82)	0.27 (0.04)	51.43
Cardboard worker	Men	1581	33.8 (11.3)	6738	11	10	13.97 (3.59)	0.29 (0.04)	50.97
Construction worker	Men	20146	35.2 (11.4)	41 284	51	10	14.03 (3.76)	0.29 (0.03)	49.63
Bricklayer	Men	16848	33.0 (10.5)	1 03 240	142	10	14.18 (4.75)	0.29 (0.04)	50.88
Garbage collector	Men	2278	38.8 (11.2)	5747	5	11	14.31 (2.39)	0.34 (0.03)	51.30
House painter	Men	9905	33.3 (10.8)	73 044	66	25	14.51 (4.77)	0.27 (0.04)	55.24
House painter	Women	4095	27.1 (6.8)	20708	83	25	14.58 (4.47)	0.27 (0.04)	57.62
Farmer	Men	14375	42.8 (11.8)	63 005	76	10	14.64 (4.84)	0.28 (0.03)	57.00
Insulation worker	Men	2151	34.5 (10.6)	7268	4	10	15.98 (10.02)	0.35 (0.11)	57.22
Storage worker	Men	12 973	21.9 (9.3)	8038	7	10	17.14 (8.28)	0.34 (0.07)	48.57
Gardener	Women	801	28.8 (10.1)	1613	4	9	18.39 (7.19)	0.33 (0.07)	55.84
Storage worker	Women	8237	18.8 (5.9)	2047	6	10	18.92 (7.53)	0.35 (0.08)	50.24
Hairdresser	Women	16363	24.8 (9.3)	95 963	96	10	19.63 (4.76)	0.29 (0.05)	65.14
Postal worker	Men	42 345	27.5 (10.3)	151 636	88	10	20.81 (6.25)	0.34 (0.05)	60.61
Kitchen assistant	Women	84528	29.2 (12,0)	200 263	501	10	21.45 (4.00)	0.33 (0.03)	55.35
Postal worker	Women	22 000	26.4 (9.0)	74035	170	10	23.28 (5.52)	0.35 (0.04)	62.43
Fish industry worker	Men	2516	34.0 (12.1)	4408	4	8	24.69 (17.50)	0.41 (0.20)	47.00
Cleaners	Women	22 526	35.6 (12.8)	22 381	101	24	27.93 (6.49)	0.38 (0.05)	59.51
Laundry worker	Men	1522	30.8 (12.4)	3119	4	10	30.05 (8.49)	0.45 (0.10)	54.71
Laundry worker	Women	4275	34.3 (12.1)	12 625	36	13	31.76 (6.99)	0.42 (0.06)	56.40
Slaughterhouse worker	Men	19839	31.5 (10.9)	101 367	278	10	37.59 (4.86)	0.45 (0.05)	52.54
Not in one of the jobs above		0	,	1 467 886	1655		, ,	,	
Total		1 015 418		5 3 7 6 9 2 7	6834				

^{*}SD could not be measured for ROM, as we did not have access to the individual measurements.

65.1° (table 1). Angular velocity and MPF were highly correlated (Spearman correlation coefficient (CC)=0.94). The correlations of ROM and angular velocity and MPF, respectively, were considerably lower (Spearman CC=0.29 and 0.21, respectively). We identified 6834 cases of CTS of which 1546 also had an operation code. Mean age at entry ranged from 18.8 years to 42.8 years between jobs (table 1).

Sex varied considerably across exposure level, while age, wristnear fractures, hypothyroidism, rheumatoid arthritis, diabetes mellitus and obesity seemed evenly distributed across levels of exposure to angular velocity (table 2).

For angular velocity, the IRR and IRR^{adj} increased monotonously from the second to the fifth exposure quintile with IRR^{adj} 2.31 (2.09 to 2.56) for the highest, compared with the lowest levels (table 3). MPF also showed an increase in IRR with increasing exposure, but after adjustment, this pattern became somewhat irregular with a lower IRR^{adj} in the fourth quintile than in the adjacent quintiles. The highest exposure group had

CTS, carpal tunnel syndrome; MPF, mean power frequency; ROM, range of motion.

Distribution of potential confounders by 1-year exposure levels (intensity*duration) shown for wrist angular velocity. Numbers (N) and point prevalences (%) at end of follow-up (2014)?

Angular velocity, °/s	Sex	>0-<20 percentile 0.01≤-6.09 [*]	20≤–40 percentile 6.09≤–7.28*	40≤−60 percentile 7.28≤−11.1*	60≤−80 percentile 11.1≤−14.5*	80≤−≤100 percentile 14.5≤−≤37.7*
Confounders		<u> </u>	<u> </u>		<u> </u>	<u> </u>
Sex, N	Total	24200	53 039	51 527	44 265	46 043
	Men	18555	11 455	20718	31 480	23 189
	Women	5645	41 584	30809	12 785	22 854
Age, mean (SD)	Total	42 (14)	44 (12)	42 (12)	41 (13)	39 (15)
Wrist-near fracture, %	Total	10.6	7.4	9.3	12.0	9.7
Hypothyroidism, %	Total	1.0	1.3	1.1	1.0	1.0
Rheumatoid arthritis, %	Total	1.0	1.0	1.0	1.0	<1.0
Diabetes mellitus, %	Total	2.7	2.0	1.6	1.5	1.6
Obesity, %	Total	3.8	6.0	6.1	4.0	3.7

^{*}Quintile limits for angular velocity.

IRR^{adj} 1.83 (1.68 to 1.98). The analyses regarding ROM showed no clear pattern.

Sex differences

The incidence rate of CTS was 10.69 cases/10 000 personyears for men and 14.03 cases/10 000 person-years for women. For women, the risk of CTS increased with increasing angular velocity from the third to the fifth quintile (figure 2). For men, the risk of CTS increased more steeply than for women from the first to the third quintile and then became slightly smaller in the fourth and the fifth quintiles for both angular velocity and MPF. For women, the results for MPF were somewhat similar to the main analyses. For women, stratified analyses of ROM showed a steep increase from the second to the third quintile, levelling off in the fourth and decreasing a bit in the fifth quintile. We found no clear exposure-response pattern for men. The differences between men and women were significant for all three exposures (p < 0.0001).

Sensitivity analyses

One-year exposure 2 years prior to outcome showed a pattern like the main analyses but with stronger IRRs reflecting an even clearer exposure-response pattern. Analyses of total cumulative exposure showed a pattern like the main analyses for angular velocity but with lower IRRs. The results showed no effect of MPF and ROM on CTS. The analyses including natural cubic splines also showed a similar pattern of the association between

Number of persons with carpal tunnel syndrome (CTS) (diagnosis or surgery), incidence rate (IR), crude and adjusted incidence rate ratio (IRR) of CTS cases by 1-year exposure levels (intensity*duration) shown for wrist angular velocity, mean power frequency and range of motion

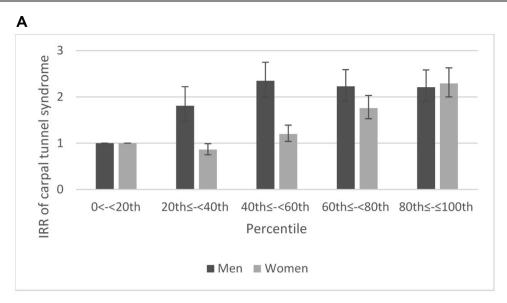
Exposure variable						
Angular velocity, °/s	0-<20th percentile 0.01≤-<6.09	20th≤-40th percentile 6.09≤-7.28	40th≤-60th percentile 7.28≤-11.1	60th≤-80th percentile 11.1≤-14.5	80th≤-≤100th percentile 14.5≤-≤37.6	Total
CTS, number of cases	472	1408	1265	1748	1941	6834
Person-years	686 215	1 460 532	983 846	1118501	1127833	5 376 927
IR*	6.9	9.6	12.9	15.6	17.2	12.7
IRR, crude	1.00 (ref.)	1.40 (1.26 to 1.56)	1.87 (1.68 to 2.08)	2.27 (2.05 to 2.52)	2.50 (2.26 to 2.77)	
IRR, adjusted†	1.00 (ref.)	1.04 (0.93 to 1.16)	1.58 (1.42 to 1.76)	2.02 (1.82 to 2.24)	2.31 (2.09 to 2.56)	
Mean power frequency, Hz	0-<20th percentile <0.001≤-<0.23	20th≤-40th percentile 0.23≤-0.24	40th≤-60th percentile 0.24≤-0.27	60th≤-80th percentile 0.27≤-0.29	80th≤-≤100th percentile 0.29≤-0.45	Total
CTS, number of cases	897	1117	1068	1941	1811	6834
Person years	936 938	1 179 089	924313	1 258 279	1 078 308	5 376 927
IR*	9.6	9.5	11.6	15.4	16.8	12.7
IRR, crude	1.00 (ref.)	0.99 (0.91 to 1.08)	1.21 (1.10 to 1.32)	1.61 (1.49 to 1.74)	1.75 (1.62 to 1.90)	
IRR, adjusted†	1.00 (ref.)	0.78 (0.72 to 0.86)	1.51 (1.37 to 1.66)	1.33 (1.23 to 1.44)	1.83 (1.68 to 1.98)	
Range of motion, °	0-<20th percentile 0.05≤-<48.7	20th≤-40th percentile 48.7≤-49.6	40th≤-60th percentile 49.6≤-52.8	60th≤-80th percentile 52.8≤-59.6	80th≤-≤100th percentile 59.6≤-65.1	Total
CTS, number of cases	974	1196	1229	1711	1724	6834
Person years	881 074	1 241 358	1 073 460	963 779	1 217 256	5 376 927
IR*	11.1	9.6	11.5	17.8	14.2	12.7
IRR, crude	1.00 (ref.)	0.87 (0.80 to 0.95)	1.04 (0.95 to 1.13)	1.61 (1.48 to 1.74)	1.28 (1.18 to 1.39)	
IRR, adjusted†	1.00 (ref.)	0.62 (0.57 to 0.68)	1.33 (1.21 to 1.45)	1.44 (1.33 to 1.55)	0.97 (0.90 to 1.06)	

Numbers in brackets are 95% CI. Persons with unknown exposures are not included in the lowest exposure category.

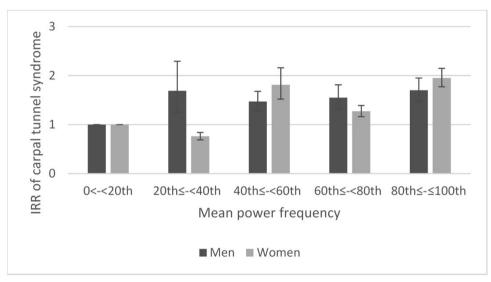
CTS, carpal tunnel syndrome.

^{*}Number of cases pr. 10 000 person-years.

[†]Adjusted for sex, age, calendar year, pregnancy, wrist-near fracture, hypothyroidism, rheumatoid arthritis, diabetes mellitus and obesity.



В



С

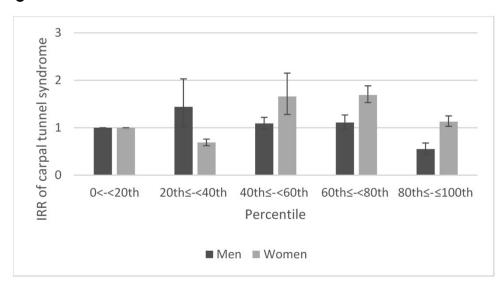


Figure 2 The association between angular velocity (A), mean power frequency (B) and range of motion (C) and carpal tunnel syndrome stratified by sex. Exposure ranges in percentiles as in table 3. Estimates are adjusted incidence rate ratios (IRRs), including marks reflecting the CIs. Adjustments are similar to adjustments described in table 3.

wrist velocity and CTS as the main analyses, as did analyses based on CTS operation alone (data not shown).

DISCUSSION

We found a clear exposure–response association between angular velocity and CTS. The association between MPF and CTS was similar but with smaller risk estimates and a less consistent pattern, whereas ROM showed no clear association with CTS. Separate analyses for men and women showed different patterns for angular velocity and MPF. The increase in risk of CTS for women followed the same pattern as the main analyses, while for men, the increase in risk of CTS was initially steeper followed by a plateau. For ROM, analyses for men and women showed no clear pattern. The sensitivity analyses testing different time frames for exposure showed similar results for angular velocity, supporting that our main findings did not depend on the specific analysis. The results for MPF and ROM were more vulnerable to the choice of analysis. For all three exposures, analyses with 1-year exposure and 2-year lag resulted in higher IRRs.

Our findings correspond well with findings in other longitudinal studies. Dale *et al*¹³ recently found a positive exposureresponse association between repetition and force defined by both an expert-based job exposure matrix and by observation of hand activity level (HAL) and CTS. Gell *et al* investigated clerical workers and found an increased risk of CTS when peak force in combination with HAL was above a predefined limit which, however, was not statistically significant. Werner *et al* found video-observed radio/ulnar wrist deviation and elbow posture to be an ergonomic risk factor in CTS among auto assembly workers. The study did not find any association between flexion/extension, force, repetition or HAL on CTS risk.

Few have studied exposure assessed by technical measurements on the risk of CTS. Nordander et al¹⁴ and Heilskov-Hansen et al¹⁵ studied movements of the wrist using electro-goniometric measurements as risk factors of developing CTS. Nordander et al investigated a range of different work places and jobs in a cross-sectional design, whereas Heilskov-Hansen et al investigated house painters with a task-based approach in a longitudinal design. Both studies showed a positive association between wrist angular velocity and CTS. One study also found a positive association with MPF, but none of the studies found an association between position and CTS. 15 No prior study has, to our knowledge, applied objective technical measurements to a large nationwide cohort with the use of a job exposure matrix. Cross-sectional studies using objective assessment of wrist exposures have also found positive associations between repetition and CTS and, contrary to our results, between position and CTS.³¹ The evidence is however still limited regarding position because of divergent results and large heterogeneity in the definition of position. 16 32

We constructed the ROM variable based on the 90th and 10th interpercentile for flexion/extension. A true effect of ROM, if any, might be detected more easily using a larger inter-percentile range. The interpercentile range holds, however, no information about whether the position is primarily flexion, extension or both. If the potential association between position and CTS is correlated to either flexion or extension, it would be difficult to demonstrate in our setting. A study of Heilskov-Hansen et al¹⁵ defined position as percentage of time with angles exceeding 45° flexion/extension or 20° radial/ulnar deviation and found no association between position and CTS. A study by Nordander et al¹⁴ likewise found no association between wrist position and CTS. Unfortunately, we could not use absolute measures of flexion or extension because the reference

value changed from functional zero in the older measurements to anatomical zero in the newer. We believe, however, that the ROMs used in this study, compared with, for instance, the absolute measures of time spent in flexed/extended positions used in the study by Heilskov-Hansen *et al*, are essentially expressing the same risk factor, non-neutral position.

Force is generally accepted as a risk factor in CTS, both alone and in combination with repetition and position. ¹⁶ Force might explain some of the association between angular velocity and MPF and CTS, as some of the measured jobs are also traditionally seen as forceful. Unfortunately, we had no measure of force in this study; thus, it is unclear if this correlation can explain some of the observed association of CTS with velocity and MPF.

Women generally have a higher risk of CTS than men.³ ³³ Our analyses showed a multiplicative interaction between sex and exposure. The differences in exposure–response patterns may, however, partly reflect sex differences in the proportion of high-force jobs in the cohort. A large part of male jobs included forceful work tasks (eg, constructions workers and scaffolders), while most female jobs did not. In a recent study of house painters by Heilskov-Hansen *et al*, ³⁴ evidence of a multiplicative interaction between exposure and sex was not found.

This study did not include measures of acceleration. Previous studies using the same technical method have found a constant ratio between acceleration and velocity of around 10, but due to methodological challenges, velocity measures have been the preferred goniometric measure, although acceleration, as well as force, is clinically relevant risk factors in work-related MSDs. 35 36

Our cohort consisted of persons born 1940–1979. Persons who turned 18 years before we could register their jobs in 1992 were included, which means that their work-related exposure before 1992 remains unknown. Analyses assessing total cumulative exposures therefore are at risk of truncation bias as persons without a full job history would be the oldest and therefore potentially a group exposed differently, most likely more, than the rest. However, mechanistic studies have shown that regarding CTS, the relevant exposures are the more recent, ³⁷ and since we identified the first outcomes in 1993 and only used updated exposure 1 year prior (with 1-year lag) in the main analysis, the influence of truncation bias is minimal in the main analyses.

Sensitivity analyses with 1-year exposure and 2-year lag generally resulted in higher IRRs and showed an even clearer exposure–response association between angular velocity and CTS. This could reflect a selection out of the most exposed jobs among symptomatic persons before being diagnosed. The sensitivity analyses including cumulated exposures have limitations, since risk time spent in one of the unmeasured jobs did not contribute to the exposure. This misclassification was independent of the outcome, and it therefore seems most likely that it will bias results of sensitivity analyses towards the null. This is in accordance with the less pronounced exposure–response relation for total cumulated exposure in comparison with the main results. The main analyses included only 1-year exposures and consequently misclassification was not a problem for these results.

The CTS diagnosis registered in DNPR is considered to be valid as the majority of patients admitted for assessment of CTS at the hospital have electroneuronography performed routinely to confirm the diagnosis.³⁸ A certain part of CTS cases is diagnosed and conservatively treated by general practitioners, especially CTS during pregnancy. These CTS cases were not included in our data because of lack of valid coding in primary care, which explains the relatively low prevalences found in this study compared with other studies based on screening in

Workplace

general/working populations. ⁶ ³⁹ ⁴⁰ CTS cases diagnosed in the healthcare system are most likely more severe than undiagnosed CTS cases in the population. Our results, therefore, may not be generalisable to all CTS cases. Furthermore, CTS cases with hand demanding work may experience more trouble than CTS cases with less hand demanding work and therefore become diagnosed earlier than CTS cases with less hand demanding work. If so, exposure–response associations could be inflated or spurious. The register-based design was further limited by a lack of information on handedness.

CONCLUSION

With the use of technical measurements and register-based outcome measures, this study found an increasing risk of CTS with increasing levels of wrist angular velocity. A similar association, although less clear, was found between repetition assessed by MPF and CTS. Exposure–response patterns differed for men and women perhaps reflecting different force exposure. No clear association was found for wrist position assessed by ROM, defined as the 90th and 10th interpercentile for flexion/extension, and CTS. Our results indicate that a reduction of repetitive and rapid movements may prevent the development of new CTS cases.

Acknowledgements The authors would like to thank Thomas Heilskov-Hansen and coworkers at the Department of Occupational Medicine in Herning, especially Annett Dalbøge, who took part in performance of the measurements. Furthermore, we would like to thank coworkers at The University Hospital in Lund, especially research and associate professor engineer Henrik Enquist for assistance during management of data from the electro-goniometric measurements.

Contributors CBL, SM, LCT, G-ÅH and JFT planned the study. CBL, SM, LCT and JFT made the analyses. CBL drafted the manuscript. All authors critically reviewed and approved the manuscript.

Funding This study was supported by grants from the Danish Working Environment Research Fund (grant #43-2010-03).

Competing interests None declared.

Patient consent for publication Not required.

Ethics approval The study protocol was accepted by the Danish Data Protection Agency (j.nr. 2013-41-2555). Approval from the Ethics Committee of the Capital Region of Denmark was not necessary according to Danish law.

Provenance and peer review Not commissioned; externally peer reviewed.

Open access This is an open access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited, appropriate credit is given, any changes made indicated, and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/.

REFERENCES

- 1 Viikari-Juntura E, Silverstein B. Role of physical load factors in carpal tunnel syndrome. Scand J Work Environ Health 1999:25:163–85.
- 2 Foley M, Silverstein B, Polissar N. The economic burden of carpal tunnel syndrome: long-term earnings of CTS claimants in Washington State. Am J Ind Med 2007;50:155–72.
- 3 Atroshi I, Gummesson C, Johnsson R, et al. Prevalence of carpal tunnel syndrome in a general population. JAMA 1999;282:153–8.
- 4 Geoghegan JM, Clark DI, Bainbridge LC, et al. Risk factors in carpal tunnel syndrome. J Hand Surg Br 2004;29:315–20.
- 5 Padua L, Di Pasquale A, Pazzaglia C, et al. Systematic review of pregnancy-related carpal tunnel syndrome. *Muscle Nerve* 2010;42:697–702.
- 6 Violante FS, Farioli A, Graziosi F, et al. Carpal tunnel syndrome and manual work: the OCTOPUS cohort, results of a ten-year longitudinal study. Scand J Work Environ Health 2016;42:280–90.
- 7 Harris-Adamson C, Eisen EA, Kapellusch J, et al. Biomechanical risk factors for carpal tunnel syndrome: a pooled study of 2474 workers. Occup Environ Med 2015:72:33–41.
- 8 Shiri R, Miranda H, Heliövaara M, et al. Physical work load factors and carpal tunnel syndrome: a population-based study. Occup Environ Med 2009;66:368–73.
- 9 Yagev Y, Gringolds M, Karakis I, et al. Carpal tunnel syndrome: under-recognition of occupational risk factors by clinicians. Ind Health 2007;45:820–2.

- 10 Bonfiglioli R, Mattioli S, Fiorentini C, et al. Relationship between repetitive work and the prevalence of carpal tunnel syndrome in part-time and full-time female supermarket cashiers: a quasi-experimental study. Int Arch Occup Environ Health 2007:80:248–53.
- 11 Bovenzi M, Zadini A, Franzinelli A, et al. Occupational musculoskeletal disorders in the neck and upper limbs of forestry workers exposed to hand-arm vibration. Ergonomics 1991:34:547–62.
- 12 Frost P, Andersen JH, Nielsen VK. Occurrence of carpal tunnel syndrome among slaughterhouse workers. Scand J Work Environ Health 1998;24:285–92.
- 13 Dale AM, Ekenga CC, Buckner-Petty S, et al. Incident CTS in a large pooled cohort study: associations obtained by a Job Exposure Matrix versus associations obtained from observed exposures. Occup Environ Med 2018;75:501–6.
- 14 Nordander C, Ohlsson K, Akesson I, et al. Exposure-response relationships in work-related musculoskeletal disorders in elbows and hands A synthesis of group-level data on exposure and response obtained using uniform methods of data collection. Appl Ergon 2013;44:241–53.
- 15 Heilskov-Hansen T, Mikkelsen S, Svendsen SW, et al. Exposure-response relationships between movements and postures of the wrist and carpal tunnel syndrome among male and female house painters: a retrospective cohort study. Occup Environ Med 2016:73:401–8.
- 16 Barcenilla A, March LM, Chen JS, et al. Carpal tunnel syndrome and its relationship to occupation: a meta-analysis. Rheumatology 2012;51:250–61.
- 17 Palmer KT, Harris EC, Coggon D. Carpal tunnel syndrome and its relation to occupation: a systematic literature review. Occup Med 2007;57:57–66.
- 18 http://www.ilo.org/public/english/bureau/stat/isco/ 2018.
- 19 Dalb'ge A. Occupational cumulative mechanical exposures as risk factors for surgery for subacromial impingement syndrome. PhD Thesis 2015.
- 20 Hansen TH. Physical work exposure and sex difference in work-related musculoskeletal disorders. *PhD Thesis* 2014.
- 21 Hansson GA, Balogh I, Byström JU, et al. Questionnaire versus direct technical measurements in assessing postures and movements of the head, upper back, arms and hands. Scand J Work Environ Health 2001;27:30–40.
- 22 Pedersen CB, G\u00f8tzsche H, M\u00f8ller JO, et al. The Danish Civil Registration System. A cohort of eight million persons. Dan Med Bull 2006;53:441–9.
- 23 Petersson F, Baadsgaard M, Thygesen LC. Danish registers on personal labour market affiliation. Scand J Public Health 2011;39:95–8.
- 24 Jensen VM, Rasmussen AW. Danish Education Registers. Scand J Public Health 2011;39:91–4.
- 25 Lynge E, Sandegaard JL, Rebolj M. The Danish National Patient Register. Scand J Public Health 2011;39:30–3.
- 26 https://norden.diva-portal.org/smash/get/diva2:970548/FULLTEXT01.pdf 2018.
- 27 http://bendixcarstensen.com/Lexis/Lexis.sas 2018.
- 28 Breslow NE, Day NE. Statistical methods in cancer research. Volume II–The design and analysis of cohort studies. IARC Sci Publ 1987;82:120–76.
- 29 Gell N, Werner RA, Franzblau A, et al. A longitudinal study of industrial and clerical workers: incidence of carpal tunnel syndrome and assessment of risk factors. J Occup Rehabil 2005:15:47–55.
- 30 Werner RA, Franzblau A, Gell N, et al. Incidence of carpal tunnel syndrome among automobile assembly workers and assessment of risk factors. J Occup Environ Med 2005;47:1044–50.
- 31 Maghsoudipour M, Moghimi S, Dehghaan F, et al. Association of occupational and non-occupational risk factors with the prevalence of work related carpal tunnel syndrome. J Occup Rehabil 2008;18:152–6.
- 32 NIOSH. Musculoskeletal disorders and workplace factors. A critical review of epedimiologic evidence for work-related musculoskeletal disorders of the neck, upper extremity, and low back. U.S. Department of Health and Human Services. 1997:97–141.
- 33 Atroshi I, Englund M, Turkiewicz A, et al. Incidence of physician-diagnosed carpal tunnel syndrome in the general population. Arch Intern Med 2011;171:941–4.
- 34 Heilskov-Hansen T, Svendsen SW, Frølund Thomsen J, et al. Sex differences in task distribution and task exposures among Danish house painters: an observational study combining questionnaire data with biomechanical measurements. PLoS One 2014:9:e110899.
- 35 Hansson GA, Balogh I, Ohlsson K, et al. Measurements of wrist and forearm positions and movements: effect of, and compensation for, goniometer crosstalk. J Electromyogr Kinesiol 2004;14:355–67.
- 36 Juul-Kristensen B, Hansson GA, Fallentin N, et al. Assessment of work postures and movements using a video-based observation method and direct technical measurements. Appl Ergon 2001;32:517–24.
- 37 Tabatabaeifar S, Svendsen SW, Johnsen B, et al. Reversible median nerve impairment after three weeks of repetitive work. Scand J Work Environ Health 2017;43:163–70.
- 38 http://neuro.dk/wordpress/nnbv/karpaltunnel/ 2018.
- 39 Franzblau A, Werner RA, Yihan J. Preplacement nerve testing for carpal tunnel syndrome: is it cost effective? J Occup Environ Med 2004;46:714–9.
- 40 Dale AM, Harris-Adamson C, Rempel D, et al. Prevalence and incidence of carpal tunnel syndrome in US working populations: pooled analysis of six prospective studies. Scand J Work Environ Health 2013;39:495–505.