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Abstract

Background:

Carpal Tunnel Syndrome (CTS) is the most common entrapment neuropathy of the upper limb. Research has shown that associative factors for CTS include occupational and biomechanical elements, gender and age. To date no systematic review has been undertaken to determine specifically whether there are any psychosocial risk factors in developing CTS. The objective, to determine whether psychosocial factors are associated with and/or predicts the development CTS.

Methods:

A systematic review was conducted including searches of PubMed (MEDLINE), EMBASE and CINAHL from inception to May 30th 2017. Quantitative studies must have investigated a minimum of one or more psychosocial factors; cognitive, affective, behavioural, vocational or interpersonal processes (e.g., social support) and include a point or risk estimate. One reviewer conducted the search and two reviewers independently assessed eligibility and completed methodological quality assessment using a modified Downs and Black checklist. Data was analysed narratively.

Results:

Six moderate to high quality studies were included in the final review. Five studies reported a positive association between psychosocial factors and CTS, where psychosocial factors was more in those who reported CTS. One study reported no positive or negative

24 association with CTS development. Four studies reported a negative association between
25 psychosocial factors and CTS where psychosocial factors was less in those who reported
26 CTS.

27

28 Conclusions:

29 There is limited evidence for a positive association between psychosocial factors and CTS.
30 However this was not a consistent finding across all included. Further research is indicated
31 in standardising CTS diagnostic criteria and investigating other working environments.

32

33 **PROSPERO REGISTRATION NUMBER: CRD42016039900**

34

35

INTRODUCTION

36 Carpal Tunnel Syndrome (CTS) is a compressive neuropathy of the median nerve at the
37 carpal tunnel [1]. It is the most common upper limb entrapment neuropathy and can
38 negatively impact function and work ability [12, 17]. CTS features can include paresthesia
39 and hypoesthesia in the distribution of the median nerve, particularly at night hand and
40 arm pain median motor deficit [17], allodynia [12], dysesthesia and hyperesthesia [24].

41

42 The reported incidence of CTS can vary between countries and it is reported that CTS
43 affects more women than men [5, 8, 13]. Atroshi et al [5] reported an estimated incidence
44 of 324 per 100,000 women in Sweden compared with 524 per 100,000 in United States of
45 America (USA). The estimated prevalence of CTS among men is 166 per 100,000 in
46 Sweden and 303 per 100,000 in USA [20]. As a result, in 1995 the estimated societal costs
47 for undergoing a CTS decompression surgery were over \$2bn in USA [26]. There are no
48 cost estimates to individuals or healthcare providers in the United Kingdom (UK) [18],
49 however, there is an estimated incidence of 87 and 192 per 100,000 for men and women
50 respectively [20]. The variance may be attributed to health beliefs and behaviours,
51 occupation and co-morbidities such as diabetes [20].

52

53 Understanding risk factors are important for patients, clinicians and policy makers to
54 identify, predict and prevent risks associated with CTS [26]. The occupational risk factors
55 for the development of CTS have been reported by Kozak et al [19]. Kozak et al [19]
56 synthesized systematic reviews and primary studies reporting occupational biomechanical
57 risk factors and concluded that there was high quality evidence supporting repetitive wrist

58 and hand movements, forceful exertion and vibration as risk factors for developing CTS.
59 However, the authors recognize that there are other factors such as age, gender, co-
60 morbidities and psychosocial factors that may interact with the occupational biomechanical
61 elements in the development of CTS [22]. Harris-Adamson et al [15] in their cohort study
62 reported how biomechanical and work psychosocial exposures, such as job strain, are
63 independent risk factors for incidence of CTS [19]. In addition, the risk of developing CTS
64 may be further attributed to high psychological work demand for women and low skill
65 discretion for men [28].

66

67 Psychosocial risk factors associated with the development of musculoskeletal disorders
68 are well documented [6, 10, 19, 22, 28]. Somatization and adverse health beliefs around
69 diagnosis and prognosis are known to be associated with chronic musculoskeletal disorders
70 [28]. High work demands [10], work stress [22] and distress [22] are related to the
71 development of low back pain. The lack of work variation, low control over work time [30]
72 and anxiety [16] are related to the development of neck pain and shoulder pain. Targeting
73 these potential risk factors can potentially reduce the incidence of CTS and as a result
74 lessen the healthcare and societal costs to individuals and families.

75

76 To date no systematic review has been undertaken to determine specifically whether there
77 are any psychosocial risk factors in developing CTS amongst adults. Accordingly, the
78 primary aims of this review are to investigate the incidence of CTS in association to
79 psychosocial factors and whether psychosocial factors may predict the development of
80 CTS.

81

METHODS

82 The systematic review was registered with PROSPERO review database (Ref:
83 CRD42016039900), and completed following the PRISMA guidelines of reporting [23].

84

Search Strategy

86 A systematic search of electronic databases PubMed, CINAHL and MEDLINE from
87 inception to May 30th 2017 was completed by reviewer (MM). An example of the
88 MEDLINE search strategy can be viewed in Figure 1. An unpublished (grey) literature
89 search and trial registry search was also completed. A hand search was completed of the
90 reference lists of the records screened for potential inclusion. Corresponding authors from
91 all included studies were contacted to determine if there were any pending article
92 publications in this area or unpublished work. An assessment of reliability (between-
93 reviewer) for the eligibility criteria was performed for a random sample of 10 potentially
94 eligible papers using a weighted Kappa statistic. This indicated that the between-reviewer
95 agreement ranged from 0.80 to 1.00 across the criteria, with perfect (Kappa: 1.00) for
96 overall agreement on eligibility of individual papers (available on request).

97

Eligibility Criteria

99 Studies were included if they met the following criteria:

100 a) Any quantitative study type

- 101 b) Adult subjects (over 18 years) with clinically diagnosed Carpal Tunnel Syndrome
102 (CTS) with or without electrophysiological testing
- 103 c) Study must have investigated a minimum of one or more psychosocial factors;
104 cognitive (e.g. neuropsychological functioning), affective (e.g., distress, mood),
105 behavioural (e.g., coping strategies), vocational (e.g. employment status, job satisfaction,
106 self – perceived work ability) or interpersonal processes (e.g., social support) and include
107 a point or risk estimate.

108

109 No limitation of publication date was applied. All considered articles had to be in the
110 English language. Articles were excluded if psychosocial factors were not measured or if
111 the participants' CTS was related to systemic pathology, fracture, radiculopathy,
112 myelopathy or upper motor neuron pathology.

113

114 Study Identification

115 Two reviewers (MM, FS) independently reviewed article titles and abstracts of all search
116 results against the inclusion criteria. From this, full text articles from potentially eligible
117 articles were retrieved and independent assessment was completed by two reviewers
118 (MM, FS). Final eligibility was decided based on full-text assessment.

119

120

121

122 Data Extraction

123 Data were extracted onto a pre-defined data extraction table independently by two
124 reviewers (MM, FS). Data extracted included: Lead author, study design, participant
125 demographics, gender, psychosocial measure, CTS diagnosis classification and strength
126 of association of CTS development (risk estimate with confidence intervals).

127

128 Quality Assessment

129 Two reviewers (MM and FS) independently assessed each included study using a
130 modified Downs and Black [9]. This tool is reported to be a valid and a reliable critical
131 appraisal tool to assess methodological quality of non-randomised control studies, which
132 was the predominant study design amongst our eligible papers [9]. The two reviewers
133 discussed their scoring and any disagreement in respect of study eligibility, data extraction
134 or critical appraisal was discussed and agreed between the two reviewers (MM, FS). If an
135 agreement could not be reached a third reviewer (MT) acted as adjudicator. Items 4, 8, 13-
136 15, 19 and 23-24 were removed from our quality assessment because the items did not
137 address our research question and aim of review.

138

139 The scoring between the two reviewers of the included studies had an agreement rate of
140 87% (109/126). Disagreements were around items 20-22 and 25-27 which were all
141 resolved through discussion and consensus was achieved.

142

143 Data Analysis

144 The study heterogeneity of the included studies was assessed by the two reviewers (MM,
145 FS) through examination of the data extraction table. This demonstrated significant
146 heterogeneity in respect of subject characteristics, co-interventions, exposure and the
147 method of assessing CTS. Based on these factors, a meta-analysis was not appropriate
148 and a narrative analysis was completed to answer our question.

149

150

RESULTS

Search Strategy

152 Seven studies met the selection criteria (Figure 2). However, on further inspection one
153 study was excluded as the study did not report risk factor [11] . Accordingly, six papers
154 were included in the final review [2, 14, 21, 25, 29, 32].

155

Study Characteristics

157 The characteristics of the included studies are presented in Table 1. Three studies were
158 cohort study designs [2, 14, 19]. One study was a matched cohort study [32]. There was
159 one case control study design [25] and one cross sectional survey [21] . Four studies
160 recruited participants from industrial assembly line factory workers; two in France [21, 29]
161 and two in United States of America [14, 32]. Two studies recruited participants of mixed
162 occupational background including manual work, administration, professional services and
163 office based occupations [2, 25]. A total of 12, 773 participants were recruited across the
164 six included studies.

165

Quality Assessment

167 The quality assessment scoring of the six included studies was very good, the mean score
168 over the eight included studies was 83% (100% score meaning all criteria met) with a
169 range of 72% to 100%. The most common criteria that included studies met were *Is the*
170 *hypothesis/aim/objective of the study clearly described?* (Criteria 1), *Are the main*

171 *outcomes to be measured clearly described in the Introduction or Methods section?*
172 *(Criteria 2) and Are the characteristics of the patients included in the study clearly*
173 *described? (Criteria 3). The most common criteria that studies scored least favourably on*
174 *were criteria 12, Were those subjects who were prepared to participate representative of*
175 *the entire population from which they were recruited? Criteria 20; were the main outcome*
176 *measures used accurate (valid and reliable)? And Criteria 22, were study subjects in*
177 *different intervention groups (trials and cohort studies) or were the cases and controls*
178 *(case-control studies) recruited over the same time?*

179

180 Carpal Tunnel Syndrome (CTS) Diagnosis Classification

181 The six included studies had variance on the diagnosis classification of CTS through a mix
182 of self-reported symptoms, clinical findings and electrophysiological testing. One study
183 confirmed CTS through a positive Tinel's sign or Phalen's test *or* if a definite diagnosis
184 based on nerve conduction velocity [21]. Werner et al [32] utilised a positive hand diagram
185 for numbness, tingling, burning, *or* pain in the median distribution, *and* a prolongation of
186 the median sensory-evoked response that was 0.5 msec longer than the ipsilateral ulnar
187 sensory response for their inclusion criteria.

188

189 One study reported CTS diagnosis through sensory and motor electrophysiological testing
190 of the median nerve and sensory testing of the ulnar nerve [2]. One study included
191 participants with a CTS diagnosis through physician examination or previous CTS
192 treatment *and* numbness, tingling, pain, or paraesthesia in the hand, wrist, arm, or forearm
193 within one month of the date of diagnosis of CTS [25]. Anderson et al [2] combined self-

194 reported symptoms in a median nerve distribution and physician interview for CTS
195 diagnosis. One study utilised clinical assessment findings only, patients were included if
196 there were symptoms related to median nerve distribution of paraesthesia for one week or
197 intermittently 10 months over a 12 month period, a positive Tinels, Phalens test or
198 diminished sensation to pin prick in median nerve distribution and an absence of
199 symptoms related to cervical radiculopathy, thoracic outlet syndrome or pronator teres
200 [29].

201

202 Psychosocial Factors Measurement

203 Two studies assessed job control using the Karasek's Job Control Questionnaire [2, 32].
204 Furthermore, Anderson et al [2] also assessed personal characteristics (negative affect
205 and "type A" behaviour) through self-administered questionnaires. Roquelaure et al [29]
206 used a self-assessment of psychological demand and social support alongside the
207 General Health Questionnaire (GHQ-12) to measure psychological status. One study
208 measured psychological job demand, work decision latitude scales and social support was
209 measured using the Job Content Questionnaire (JCQ) [14]. One study collected
210 information on psychosocial risk factors through participants' medical records and a
211 telephone interview [25]. LeClerc et al [21] assessed psychological and psychosomatic
212 wellbeing using Langner's screening questionnaire and job control through a self-
213 assessment Likert scale questionnaire.

214

215

216

217 Psychosocial Risk Factors and Association to CTS

218 The 6 included studies reported both positive and negative associations within each paper.
219 Five studies reported a positive association between psychosocial factors and CTS where
220 psychosocial factors was more in those who reported CTS. A GHQ-12 score over the 90th
221 percentile (i.e. over 18.5), indicating high psychological distress, (Odds Ratio (OR) 4.3;
222 95% CI: 1.0 to 18.6) [29] and “Psychological problems” (OR 2.34; 95% CI: 1.42 to 3.85)
223 [21] were more frequent and statistically significant in workers with CTS. Low social
224 support was reported as a positive association in CTS (OR 1.2; 95% CI: 0.90 to 1.80) [2].
225 Furthermore, a poor social network was also positively associated with CTS development
226 (OR 1.2; 95% CI: 0.7 to 2.2) [2]. There was a small (non-significant) positive association
227 between “type A behaviour” and CTS symptoms (OR 1.1; 95% CI: 0.70 to 1.80) [2].

228

229 A high psychological work demand score (Hazard Ratio (HR) 1.57; 95% CI: 1.06 to 2.33)
230 [14] and a high job strain (high demand and low control) was positively associated with
231 CTS (HR 1.86; 95% CI: 1.11 to 3.14) [14]. Furthermore, a high job demand (OR 1.3; 95%
232 CI: 0.9-1.8) [2], low level of job control and dissatisfaction (OR 1.59; 95% CI: 1.04 to 2.43)
233 [21] were positively associated with CTS. Workers reporting the least influence over their
234 work were also positively associated with CTS (OR 2.86; 95% CI: 1.10 to 7.14).

235

236 4 studies reported a negative association between psychosocial factors and CTS where
237 psychosocial factors was less in those who reported CTS. High social support (HR 0.54;

238 95% CI: 0.31 to 0.95) [14], high hierarchical control of work performed (OR 0.5; 95% CI:
239 0.20 to 1.30) [29], more co-worker support (OR 0.69; 95% CI: 0.48 to 0.99) [32] and a high
240 decision latitude (HR 0.73; 95% CI: 0.51 to 1.04) [14] were negatively associated with
241 CTS. Whereas, Anderson et al [2] reported low job control was negatively associated with
242 CTS (OR 0.9; 95% CI: 0.70 to 1.40).

243

244 One study reported that time pressures at work had no positive or negative association
245 with CTS development (OR 1.0; 95% CI: 0.7-1.6) [2].

246

247

DISCUSSION

248 This is the first systematic review investigating the incidence of psychosocial risk factors in
249 association with CTS and whether psychosocial risk factors predict the development of
250 CTS. Five moderate to high quality studies reported a positive association between
251 psychosocial factors; high psychological work demand, high job strain, least influence over
252 their work, a high job demand, low level of job control, high psychological distress, low
253 social support, poor social network and “type A behaviour” and CTS. Four moderate to
254 high quality studies reported a negative association between psychosocial factors; high
255 decision latitude, high hierarchical control of work, more co-worker support and high social
256 support and CTS. One study reported that time pressures at work had no positive or
257 negative associations with CTS. There was a wide variance of the working environments
258 and occupations of the recruited participants. Four studies recruited participants from
259 industrial assembly line factory workers. Two studies recruited participants of mixed
260 occupational background including manual work, administration, professional services and
261 office based occupations. This variance may impact the external validity to other
262 occupations and working environments.

263

264 The diagnostic criteria for CTS varied considerably between each study, and included a
265 combination of subjective reported symptoms, participant self-reported symptoms, clinical
266 assessment testing and/or sensory and motor electrophysiological testing. This may
267 question the reliability and external validity of findings. Furthermore, all included studies
268 used varying psychosocial measurements including Karasek’s Job Control Questionnaire,
269 personal characteristics (negative affect and “type A” behaviour) through self-administered

270 questionnaires, General Health Questionnaire (GHQ-12), Job Content Questionnaire
271 (JCQ) and Langner's screening questionnaire. There are no universally agreed diagnostic
272 criteria for CTS which can be used as a comparative consistently within both clinical and
273 research fields [32]. The standardisation of CTS diagnostic criteria is essential for
274 clinicians and researchers alike in order to generate research where results can be cross
275 compared and pooled to make meaningful conclusions regarding this common and
276 disabling condition.

277

278 Contrasting this review's results to other populations with entrapment neuropathies may
279 enhance knowledge and understanding of assessment and management strategies.
280 However, there are a limited number of studies published in this area of research. A
281 systematic review reporting the prognostic role of psychological factors in adults with
282 conservatively treated 'sciatica' [4], reported depression, avoidance behaviour, 'nonverbal
283 pain behaviour' and social support significant in pain intensity prognostic outcomes. The
284 psychosocial factors reported in this study were similar to our findings, however, caution
285 should be taken as this is based on only one longitudinal study with a small sample size.

286

287 Psychosocial stressors may have a synergistic effect on pathophysiological at the level of
288 the person leading to poor tolerance of minor symptomology consistent with being at risk
289 of CTS; catastrophising and associated illness behaviours such as over protection of
290 and/or avoidance of movement using may be related to developing symptoms of
291 CTS. Equally, because CTS has been linked to conditions known to have high levels of
292 psychological distress e.g. fibromyalgia, care may be indicated to prevent the diagnosis of

293 CTS based purely on clinical signs and symptoms, which may in fact be due to the
294 preexisting condition [30].

295

296 Psychosocial factors have been widely linked to the presentation and development of
297 persistent musculoskeletal pain, although few studies have attempted to assess their
298 impact on compression neuropathies. At present there has been a paucity of research on
299 how these factors may interact with specific pathophysiological mechanisms implicated in
300 the development of musculoskeletal pain. The accepted view is that these factors act
301 secondary to the primary 'physical' pathology acting in an adjunct capacity. There is
302 however a growing focus within contemporary research to assess the potential for these
303 factors to directly interact and influence with the pathophysiological mechanisms [7] and
304 this is likely to be both revealing and informative.

305

306 Following this systematic review, further research is warranted to identify the association
307 and prediction of psychosocial risk factors and CTS. The consistency of CTS diagnostic
308 criteria needs to be established in future studies, this will enhance the analysis of results
309 when this review is updated. There should be a research priority to undertake prospective
310 studies with longer term follow up across multiple professions, working environments and
311 healthcare settings. This would improve the generalisability of results and enhance our
312 assessment strategies in clinical practice.

313

314 There are potential limitations to this review which is a result of the current available
315 literature. Firstly, six studies were identified and included which were highly
316 heterogeneous. This can question the strength of the narrative analysis and how
317 generalisable our findings are to clinical practice. The occupations and working
318 environments of recruited participants varied across the included studies. Therefore
319 making it challenging to interpret the results and apply the analysis to specific populations.
320 Whilst it is recognised that psychosocial factors are multi-dimensional complex
321 interactions, there was variability of the psychosocial measurement tool used across the
322 included studies, adopting a more standardised approach in future research may enable a
323 meta-analysis to be completed.

324

325

CONCLUSION

326 This review indicates a positive association between psychosocial factors (high
327 psychological work demand, high job strain, least influence over their work, a high job
328 demand, low level of job control, high psychological distress, low social support, poor
329 social network and “type A behaviour”) and CTS, where these factors were present in
330 those who reported CTS. In addition, a negative association between psychosocial factors
331 (high decision latitude, high hierarchical control of work, more co-worker support and high
332 social support) and CTS, where these psychosocial factors was less likely to be
333 associated with CTS has been highlighted. However, these conclusions should be
334 interpreted with caution as the results were based on highly heterogeneous studies.
335 Further prospective studies across multiple working environments and professions are
336 indicated to enhance understanding between the association and prediction of
337 psychosocial risk factors and CTS.

338

339 Word Count: 3, 293

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341 Conflict of Interest Statement: The authors declare that they have no conflict of interest.

342 Statement of Human and Animal Rights: This article does not contain any studies with
343 human or animal subjects

344 Statement of Informed Consent: This article is systematic review of literature, as such, it
345 does information relating to or with human or animal subjects is not applicable

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349

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437 Figure Legends:

438 Figure I – MEDLINE Search Strategy

439 Figure II – Flow Diagram

440