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Andrews, Pamela; Steultjens, M.; Riskowski, J.

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Chronic Widespread Pain Prevalence in the General Population: A Systematic Review

P. Andrews<sup>1</sup>, M. Steultjens<sup>1</sup>, J. Riskowski<sup>1</sup>,

1. Institute for Allied Health Research, Glasgow Caledonian University, Glasgow, UK

Correspondence

Pamela Andrews

Room A262

Cowcaddens Road, G4 0BA

Email: pamela.andrews@gcu.ac.uk

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### Abstract

**Background and Objective:** Chronic widespread pain (CWP) is a significant burden in communities. Understanding the impact of population-dependent (e.g., age, gender) and context-dependent (e.g., survey method, region, inequality level) factors have on CWP prevalence may provide a foundation for population-based strategies to address CWP. Therefore, the purpose of this study was to estimate the global prevalence of CWP and evaluate the population and contextual factors associated with CWP.

**Databases and Data Treatment:** A systematic review of CWP prevalence studies (1990-2016) in the general population was undertaken. Meta-analyses were conducted to determine CWP prevalence, and study population data and contextual factors were evaluated using a meta-regression.

**Results:** Thirty-nine manuscripts met the inclusion criteria. Study CWP prevalence ranged from 1.4%-24.0%, with CWP prevalence in men ranging from 0.8%-15.3% and 1.7%-22.1% in women. Estimated overall CWP prevalence was 9.6% (8.0-11.2%). Meta-regression analyses showed gender, United Nations country development status, and human development index (HDI) influenced CWP prevalence, while survey method, region, methodological and reporting quality, and inequality showed no significant effect on the CWP estimate.

**Conclusion:** Globally CWP affects one in ten individuals within the general population, with women more likely to experience CWP than men. HDI was noted to be the socioeconomic factor related to CWP prevalence, with those in more developed countries having a lower CWP prevalence than those in less developed countries. Most CWP estimates were from developed countries, and CWP estimates from

countries with a lower socioeconomic position is needed to further refine the global estimate of CWP.

What does this study add? This systematic review and meta-analysis updates the current global CWP prevalence by examining the population-level (e.g., age, gender) and contextual (e.g., country development status; survey style; reporting and methodologic quality) factors associated with CWP prevalence. This analyses provides evidence to support higher levels of CWP in countries with a lower socioeconomic position relative to countries with a higher socioeconomic position.

**Key Words:** Chronic widespread pain, general population, class, socioeconomic position, systematic review, epidemiology.

## 1 **1. Introduction**

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The estimated prevalence of chronic pain, defined as pain lasting more than three 3 months, is between 35% and 50% worldwide (Elzahaf et al., 2012). Epidemiologic 4 studies of chronic pain have tended to centre on one joint, such as the foot, knee, low 5 back and shoulder (Freburger et al., 2009; Hiller et al., 2012; Hurley et al., 2012; Roh 6 et al., 2012). However, some individuals experience pain all over the body, and in 7 1990, the term "chronic widespread pain" (CWP) was defined as pain lasting longer 8 9 than 3 months, with pain being on the left and right sides of the body, above and below the waist, and on the axial skeleton (Wolfe et al., 1990). With the formal 1990 10 definition of CWP, a recent review suggested the worldwide estimate of CWP ranges 11 12 from 10.6% to 11.8% (Mansfield et al., 2016).

CWP adversely affects quality of life, mobility and physical function (Nicholl 13 et al., 2009). Further, CWP is a common condition associated with fibromyalgia 14 syndrome (FMS) and is noted to be an early indicator of FMS (Forseth et al., 1999; 15 Toda 2011). CWP and FMS can place significant challenges onto the healthcare 16 17 system, and inconsistent messages exist within the literature with regard to the most effective diagnosis and management strategies (Lee et al., 2014). Living with CWP 18 can have significant cost implications to not only the government but also the 19 20 individual patient in terms of lost work, benefits and medical costs (Barham 2012; Gaskin and Richard 2012; Henschke et al., 2015). In Europe approximately 1.5-3.0% 21 of their annual gross domestic product (GDP) is spent on chronic pain (Barham 2012; 22 Gaskin and Richard 2012). In the United States (US), chronic pain costs between 23 \$560 and \$635 billion annually, a cost higher than heart disease (\$309b), cancer 24

(\$243b) and diabetes (\$188b). Further, direct and indirect annual costs of CWP per
patient in the US are estimated to be \$12,428 (Schaefer et al., 2015).

Since the inception of the ACR definition in 1990, researchers have estimated CWP at the local and country level in order to determine burden of CWP in the population (Mansfield et al., 2016). While this prior study of global CWP prevalence addressed a significant gap, the current review and analyses aims to build upon it to update the CWP estimate and to evaluate study population (e.g., age, gender) and contextual (e.g., country development status; survey style; methodology and reporting quality) factors associated with CWP prevalence.

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#### 36 **2.0 Methodology**

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## 38 2.1 Search Strategy

A primary literature search of electronic databases was performed to extract epidemiological studies of the global prevalence of chronic widespread pain (CWP) in the general adult population (1st January 1990 to 5th April 2017). The lower year limit of 1990 was applied to align with the seminal publication defining CWP (Wolfe et al., 1990).

Electronic databases included in the study were PSYCinfo, MEDLINE, 44 Cumulative Index to Nursing and Allied Health Literature (CINAHL), The Allied and 45 Complementary Database (AMED), Cochrane library, PubMed and OVID. To 46 identify publications related to the prevalence of CWP, there were three criterion 47 components of the search strategy: (1) outcome, (2) methodology, and (3) population, 48 which were combined using Boolean operators. Outcome search terms were 49 associated with 'chronic pain,' and methodology search terms were associated with 50 'prevalence,' and to limit the likelihood of sub-populations a 'NOT' operator of 51 'cancer' or 'diabetes' was used to reduce publications that were not focused on the 52 general population. No language restrictions were applied. 53

54

# 55 2.2. Selection Criteria and Data Extraction

Set inclusion and exclusion were specified a priori and applied in three steps (Table 1). In the first step, studies were eliminated if it was evident from the title that criteria regarding outcome, methodology, and population were not satisfied. At this title stage, one reviewer (PA) eliminated publications, with a second reviewer (JLR) verifying these results. In the second step, two reviewers (PA and JLR) independently

reviewed abstracts to determine if inclusion criteria were met. From the abstract stage, 61 full-texts of the manuscripts were obtained and reviewed for inclusion, with study 62 methods evaluated against the set criteria. Manuscripts written in languages other than 63 English were included and were reviewed by others comfortable with the language. 64 Prevalence data was recorded for CWP in the general population along with separate 65 figures for gender and age as well as weighted and unweighted where applicable. If 66 data from the same manuscript were reported in multiple publications, data are 67 reported as one study. 68

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# 70 2.3 Assessment of Study Quality

Two reviewers (JLR and PA) independently evaluated the included studies based on 71 72 the criteria in the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) checklist (Von Elm et al., 2007), a reliable method for 73 reporting observational studies (Tate and Douglas 2011). For this analysis, the 74 STROBE was modified to include 12 items. Each item was scored independently as 75 either 'Identified' (1 point) or 'Not Identified' (0 point), and scoring was discussed to 76 77 reach consensus. The points from the modified STROBE were summed (Table S1), and studies were considered as having low risk of bias if they were found to be of 78 high quality ( $\geq 9/12$ ) and high risk of bias if they were found to be of low quality 79 80  $(\leq 8/12)$ , with this cut-point near the 80% quality cut-point (Slavin 1995).

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#### 84 2.4 CWP Study Contextual Data

Additional contextual data was added to evaluate factors associated to the CWP prevalence. Contextual data included were the country's United Nations (UN) development status (i.e., developed and developing country) (UN 2012), World Health Organisation (WHO) region (WHO 2017), Human Development Index (HDI) (HDR 2016), and Gini index (TWB 2017).

The HDI is a composite measure of three basic dimensions: life expectancy, 90 education, and per capita income, and it is an indicator of the country's support 91 92 systems and its citizen's health, personal, social, and political freedom, and wellbeing. The GINI index is a measure of statistical dispersion used to represent the net 93 income distribution within a country, and it can define a country's level of rich-to-94 poor inequality. GINI index values can range between 0 and 1, with 0 representing 95 perfect equality and 1 representing perfect inequality, but in practice, it ranges from 96 approximately 0.2 to 0.7 (TWB 2017). The HDI and Gini values and the country's 97 development status were based on the year of the data collection, and when an 98 estimate was not available for the study year, the closest year to the study collection 99 100 period was used.

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#### 102 2.5 Data Analysis

A meta-analysis combined the CWP prevalences of the individual studies to estimate the prevalence of CWP for the overall population sample as well as by gender, age, WHO region and survey method. Univariate meta-analyses were performed on all individual and contextual variables to determine if there was a significant effect of the variable on CWP prevalence. Statistical significance was set to p<0.05. There was no multiple testing correction, which may increase the likelihood of false positive;

109	however it is a valid means for exploring value of each variable in regression
110	modelling (Bender and Lange 2001). $I^2$ statistical calculations were conducted to
111	examine the heterogeneity between all studies and subgroups. The 95% confidence
112	intervals were calculated using the Wilson score method with continuity corrections.
113	All statistical analyses were performed using R version 3.3.1.
114	

- 116 **3.0 Results**
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#### 118 *3.1 Study Selection*

Implementation of the search strategy yielded 12,097 records, of which 5,768 were 119 duplicates (Figure 1). Screening of titles excluded 6,038 manuscripts, leaving 291 120 records for the abstract stage. At the abstract stage an additional 120 titles were 121 122 excluded, leaving 171 for the full-text stage. Full-text screening excluded 132 manuscripts, leaving 39 manuscripts (30 unique studies with 41 CWP population-123 level estimates) with a total of 632,937 participants. Study sample size ranged from 124 361 (Santos et al., 2010) to 501,733 (Walker-Bone et al., 2016). Twenty-six studies 125 included both genders, whereas three studies included only women (Abusdal et al., 126 1997a; Abusdal et al., 1997b; Schochat and Beckmann 2003; Topbas et al., 2005) and 127 one study included only men (Lee et al., 2010; Macfarlane et al., 2009b). Six studies 128 failed to report gender characteristics (Bergman et al., 2001; Gerdle et al., 2008; 129 Hagen et al., 2005; Lindell et al., 2000; Papageorgiou et al., 2002; Scudds et al., 2006; 130 Wolfe et al., 1995). Participants' age in the included studies ranged from 15-94 years. 131 132

133 Figure 1: Flow chart of included studies

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135 *3.2 Study Characteristics* 

Country CWP prevalence data (Table 2) is from the UK (N=9) (Benjamin et al., 2000;
Carnes et al., 2007; Choudhury et al., 2013; Croft et al., 1993; Flüß et al., 2015; Hunt
et al., 1999; Lee et al., 2010; Macfarlane et al., 1999; Macfarlane et al., 2009a;
Macfarlane et al., 2009b; Pang et al., 2010; Papageorgiou et al., 2002; Vandenkerkhof
et al., 2011; Walker-Bone et al., 2016), Spain (N=4) (Bannwarth et al., 2009; Branco

141	et al., 2010; Dueñas et al., 2016; Dueñas et al., 2015; Lee et al., 2010; Macfarlane et
142	al., 2009b; Mas et al., 2008), Brazil (N=3) (Assumpção et al., 2009; Cabral et al.,
143	2014; Santos et al., 2010), Sweden (N=4) (Bergman et al., 2001; Dragioti et al., 2016;
144	Gerdle et al., 2008; Lee et al., 2010; Lindell et al., 2000; Macfarlane et al., 2009b),
145	US (N=2) (Riskowski 2014; Wolfe et al., 1995), France (N=2) (Bannwarth et al.,
146	2009; Branco et al., 2010; Perrot et al., 2011), Germany (N=2) (Bannwarth et al.,
147	2009; Branco et al., 2010; Schochat and Raspe 2003), Israel (N=2) (Ablin et al., 2012;
148	Buskila et al., 2000), Italy (N=2) (Bannwarth et al., 2009; Branco et al., 2010; Lee et
149	al., 2010; Macfarlane et al., 2009b), Norway (N=2) (Abusdal et al., 1997a; Abusdal et
150	al., 1997b; Hagen et al., 2005), Belgium (N=1) (Lee et al., 2010; Macfarlane et al.,
151	2009b), Canada (N=1) (White et al., 1999), Estonia (N=1) (Lee et al., 2010;
152	Macfarlane et al., 2009b), Hong Kong (N=1) (Scudds et al., 2006), Hungary (N=1)
153	(Lee et al., 2010; Macfarlane et al., 2009b), Netherlands (N=1) (Picavet and Schouten
154	2003), Poland (N=1) (Lee et al., 2010; Macfarlane et al., 2009b), Portugal (N=1)
155	(Bannwarth et al., 2009; Branco et al., 2010) and Turkey (N=1) (Topbas et al., 2005).
156	The included studies varied in terms of CWP definition, survey method and
157	measurement processes (Table S2). CWP was identified using the ACR criteria
158	(N=24) (Wolfe et al., 1990), the Manchester definition (N=3), and a study-specific
159	definition (N=5). CWP data were collected by postal survey (N=10) (Abusdal et al.,
160	1997a; Abusdal et al., 1997b; Bergman et al., 2001; Carnes et al., 2007; Croft et al.,
161	1993; Dragioti et al., 2016; Flüß et al., 2015; Gerdle et al., 2008; Hagen et al., 2005;
162	Lindell et al., 2000; Papageorgiou et al., 2002; Picavet and Schouten 2003), telephone
163	(N=1) (Dueñas et al., 2016; Dueñas et al., 2015) face-to-face interviews (N=2)
164	(Cabral et al., 2014; Mas et al., 2008), clinical examination (N=1) (Santos et al.,
165	2010), touch screen questionnaire (N=1) (Walker-Bone et al., 2016) or combined

methods (N=15) (Ablin et al., 2012; Assumpção et al., 2009; Bannwarth et al., 2009;
Benjamin et al., 2000; Branco et al., 2010; Buskila et al., 2000; Choudhury et al.,
2013; Hunt et al., 1999; Lee et al., 2010; Macfarlane et al., 1999; Macfarlane et al.,
2009a; Macfarlane et al., 2009b; Pang et al., 2010; Perrot et al., 2011; Riskowski
2014; Schochat and Raspe 2003; Scudds et al., 2006; Topbas et al., 2005;
Vandenkerkhof et al., 2011; White et al., 1999; Wolfe et al., 1995).

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### 173 *3.3 Chronic Widespread Pain Prevalence*

The included 30 studies provided 41 prevalence estimates of CWP. From the included manuscripts, overall CWP sample prevalence, ranged from 1.4% in the UK (Walker-Bone et al., 2016) to 24.0% in Brazil (Assumpção et al., 2009). In combining the studies where sample prevalence data was available and excluding any studies with single gender analysis, a total of 622,169 participants across 26 studies were included in the analysis, and the estimated overall CWP prevalence was 9.6% (95% confidence interval [CI]: 8.0-11.2%).

181

182 *3.4 Gender* 

Four studies were of a single gender, one of men only (Lee et al., 2010; Macfarlane et al., 2009b) and three of women only (Abusdal et al., 1997a; Abusdal et al., 1997b; Schochat and Raspe 2003; Topbas et al., 2005), while eleven studies provided estimates from both genders in the general population. In the single gender studies, CWP prevalence in men was estimated at 8.3%, with data only available from a CWP study in Europe (Lee et al., 2010; Macfarlane et al., 2009b), while in women CWP prevalence ranged from 13.5% in Germany (Schochat and Raspe 2003) to 22.1% in

190	Norway (Abusdal et al., 1997a; Abusdal et al., 1997b). When combining women-only
191	studies (n=6805), the CWP prevalence in women was 17.3% (16.4-18.1%).

Where studies included data for both genders, the prevalence in men ranged 192 from 0.8% in Sweden (Dragioti et al., 2016) to 15.3% in Estonia (Lee et al., 2010; 193 Macfarlane et al., 2009b), and in women it ranged from 1.7% (Walker-Bone et al., 194 2016) to 15.6% (Croft et al., 1993), with both study CWP estimates coming from the 195 UK. When combining the data for men (n=242,808), the estimated overall CWP 196 prevalence was 7.2% (5.5-8.9%), while in women, (n=291,129) the estimated overall 197 CWP prevalence was 11.2% (8.3-14.2%). Univariate regression analysis by gender 198 found women had a significantly higher CWP prevalence relative to men (p<0.01; 199 Table 3). 200

201

202 *3.5 Age* 

Age-specific data was provided in 14 studies (Abusdal et al., 1997a; Abusdal et al., 203 1997b; Bannwarth et al., 2009; Benjamin et al., 2000; Bergman et al., 2001; Branco et 204 al., 2010; Buskila et al., 2000; Carnes et al., 2007; Croft et al., 1993; Dragioti et al., 205 2016; Dueñas et al., 2016; Dueñas et al., 2015; Gerdle et al., 2008; Hunt et al., 1999; 206 Lee et al., 2010; Lindell et al., 2000; Macfarlane et al., 1999; Macfarlane et al., 207 2009b; Mas et al., 2008; Picavet and Schouten 2003; Walker-Bone et al., 2016). Due 208 209 to the variability in each of the available studies age bandings it was not possible to combine the data for further analysis. Of studies evaluating CWP by age, nine 210 reported an increase in pain prevalence with age (Abusdal et al., 1997a; Abusdal et 211 al., 1997b; Benjamin et al., 2000; Bergman et al., 2001; Buskila et al., 2000; Croft et 212 al., 1993; Dueñas et al., 2016; Dueñas et al., 2015; Gerdle et al., 2008; Hunt et al., 213 1999; Lindell et al., 2000; Macfarlane et al., 1999; Picavet and Schouten 2003), while 214

four reported a decrease or levelling out of pain prevalence from 50-60 years only to
increase again from 60 years (Croft et al., 1993; Dragioti et al., 2016; Lee et al., 2010;
Macfarlane et al., 2009b; Mas et al., 2008).

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219 *3.6 Survey Method* 

Methods of data collection varied between studies, with sixteen studies using a single 220 style of data collection (i.e., telephone, face-to-face or clinical/physical exam) and 221 fourteen using a combined method (postal or telephone with clinical/physical 222 examination). The method of survey was further grouped into a personal (face-to-223 face, telephone and clinical examination) and non-personal (postal survey) approach 224 for further analysis. Seventeen studies with 21 CWP estimates (n=546,553) were 225 226 included in the personal group, while nine studies (n=75,616) were included in the non-personal survey method. Random-effects CWP prevalence estimates between 227 personal and non-personal were similar (9.9% [7.5-12.3%] v 7.6% [4.7-10.4%,], 228 p=0.981). 229

230

231 3.7 Region

By WHO regions (Figure 2), there were nineteen studies of CWP prevalence in Europe, five of the Americas, and one in Western Pacific. Combining country data for Europe and the Americas revealed overall CWP prevalence estimates were similar (8.9% [6.9-10.9%] v 10.9% [5.1-16.7%], p=0.497).

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237 Figure 2: Geographical spread of CWP prevalence

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### 240 3.8 Development status, HDI Index and GINI Index

Contextual factors of socioeconomic position included the UN development status, 241 HDI and GINI Index. Based on the country's UN development status, there were 27 242 CWP estimates (n=620,214) from developed countries, and the CWP prevalence of 243 8.6% (6.9-10.3%). Three CWP estimates were from developing countries (n=1955), 244 and the CWP prevalence estimate for these countries was 14.5% (3.9-25.1%). The 245 meta-regression showed UN development status relating to CWP prevalence 246 (p=0.041), which was similar to the HDI results that countries with a higher the HDI 247 248 (i.e., more developed country) had a lower reported CWP prevalence (p=0.001).

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## 250 *3.9 Methodological Quality*

Quality scores ranged from 6/12 to 12/12 (Table S3), with 10 manuscripts being noted 251 as having low quality (≤8/12) (Abusdal et al., 1997a; Abusdal et al., 1997b; Gerdle et 252 al., 2008; Pang et al., 2010; Papageorgiou et al., 2002; Perrot et al., 2011; Picavet and 253 Schouten 2003; Scudds et al., 2006; Vandenkerkhof et al., 2011; White et al., 1999) 254 and 29 of high quality ( $\geq 9/12$ ) (Ablin et al., 2012; Assumpção et al., 2009; Bannwarth 255 et al., 2009; Benjamin et al., 2000; Bergman et al., 2001; Branco et al., 2010; Buskila 256 et al., 2000; Cabral et al., 2014; Carnes et al., 2007; Cho et al., 2012; Choudhury et 257 al., 2013; Croft et al., 1993; Dragioti et al., 2016; Dueñas et al., 2016; Dueñas et al., 258 259 2015; Flüß et al., 2015; Hagen et al., 2005; Hunt et al., 1999; Lee et al., 2010; Leveille et al., 2001; Lindell et al., 2000; Macfarlane et al., 1999; Macfarlane et al., 260 2009a; Macfarlane et al., 2009b; Mas et al., 2008; Riskowski 2014; Santos et al., 261 2010; Schochat and Raspe 2003; Topbas et al., 2005; Walker-Bone et al., 2016; 262 Wolfe et al., 1995). Most of the included manuscripts (N=37) consistently identified 263 the CWP outcome measure and study eligibility criteria. Lack of appropriate reporting 264

265	was in reporting bias and providing detailed methodology. When addressing bias,
266	only five manuscripts identified their methods for controlling bias, 18 failed to
267	provide their adjusted estimates and precision (e.g., 95% confidence interval) for
268	CWP prevalence, and 27 did not report data collection methods and participant
269	recruitment. The high-quality studies (n=592,034) had a CWP prevalence of 9.7%
270	(7.4-12.1%), while the low-quality studies (n=30,135) had a similar (p=0.242) CWP
271	prevalence estimate of 7.5% (5.3-9.6%).

275

The current review aimed to determine the global prevalence of CWP in the general 276 population. The review identified 30 studies with 41 estimates of CWP prevalence. 277 From these CWP studies, global CWP prevalence estimate was 9.6% (95% CI: 8.4-278 11.2%). Women were found to have a higher CWP prevalence than men (11.2% v 279 7.2%). In identifying other factors associated with CWP prevalence, data collection 280 style of personal or non-personal approach showed no significant effect, but the 281 282 personal approach (i.e., face-to-face, telephone, examination) tended to increase CWP prevalence compared to non-personal (9.9% v 7.6%). Additionally, countries with a 283 higher human development index (HDI) had a lower CWP prevalence compared to 284 lower HDI countries (8.6% v 14.5%). Results from this work suggest there is a 285 significant burden of CWP on the general population, particularly among women, and 286 that improving a country's standard of living, as indicated by the HDI, may influence 287 CWP prevalence. 288

289

290 4.1 Diagnostic Criteria

CWP diagnosis originally came from the ACR 1990 criteria of FMS. However, 291 recently the Manchester criteria, which requires pain to be found in two locations of 292 two contralateral limbs and also in the axial skeleton (Okifuji and Hare 2014), is 293 gaining traction. Although, the ACR 1990 and Manchester definitions allow 294 standardisation and comparisons to be made (Okifuji and Hare 2014), results of CWP 295 prevalence by these two definitions are not similar. Gerdle et al., 2008) 296 found CWP to be 7.4% when applying the Manchester criteria, while they only 297 recorded 4.8% with ACR. In contrast the Manchester cohort (Benjamin et al., 2000; 298

Hunt et al., 1999; Macfarlane et al., 1999) found CWP to be 4.7% with the 299 Manchester criteria and 12.9% with the ACR definition. Between these studies 300 sample sizes were different (n=1953 (Benjamin et al., 2000; Hunt et al., 1999; 301 Macfarlane et al., 1999) v n=7637 (Gerdle et al., 2008)), but these two studies also 302 differed in the number of pain sites the study participant could select in the pain chart. 303 The Manchester group (Benjamin et al., 2000; Hunt et al., 1999; Macfarlane et al., 304 1999) had 26 pain sites available versus 17 in the Gerdle et al study (Gerdle et al., 305 2008). The number of pain sites available to select could not only lead to participant 306 307 confusion, but it could also lead to participants under or over-reporting the number of pain site depending on if their specific pain site is not provided. Research evaluating 308 style of pain reported has suggested that the most efficient method for assessing this is 309 through the completion of a body manikin (Croft 2002) or through the number of pain 310 sites rather than the location of pain (Beasley and Macfarlane 2014), which is what 311 the ACR 2010 definition does. As such, future research should aim to determine a 312 uniform diagnosis for CWP that utilises a body manikin with a set number of pain 313 sites to ensure prevalence figures are reliable and can be comparable across studies. 314

315

## 316 *4.2 Age and Gender*

The current review found no significant difference in CWP by age group. However, part of the lack of effect may be due to few studies reporting CWP prevalence by similar age group bandings. Given the inconsistencies in age group reporting it is difficult to determine accurate CWP prevalence estimates, and future studies should aim to report specific prevalence estimates for age using consistent age banding.

322 Studies have consistently shown that women experience more pain than men 323 (Bartley and Fillingim 2016; Fillingim et al., 2009; Pieretti et al., 2016). This review

found similar results, with the meta-analysis showing CWP was higher in women 324 compared to men (11.4% v 7.2%). Reasons for the gender differences in pain are 325 often hypothesised to be biological, but studies have also suggested that differences in 326 pain may relate to psychological or social factors (Wiesenfeld-Hallin 2005) as some 327 men may fear they will appear weak if they express their pain (Fillingim et al., 2009). 328 Researchers hypothesise that while women score higher on pain, they are often 329 encouraged to talk about their feelings and may be more comfortable than men at 330 indicating they are experiencing pain (Fillingim et al., 2009). 331

332

### 333 *4.3 Geographical region*

Although there were no regional variations of CWP prevalence noted by the meta-334 regression, these results should be viewed with caution, as regions other than Europe 335 and the Americas were not well represented. The prior CWP review (Mansfield et al., 336 2016) noted differences between Europe and America, with Europe having a higher 337 CWP prevalence than the Americas (12.8% v 7.1%). These prior results are in 338 contrast with the current study where it shows a non-significant difference between 339 regions (8.9% in Europe v 10.9% in Americas). A possible explanation for this CWP 340 difference could be the variation in the studies included between the two reviews. For 341 example, this review included only general population studies, not studies of specific 342 343 populations within the larger population. As such, the prior meta-analysis (Mansfield et al., 2016) included a CWP study of a Native American population (Jacobsson et al., 344 1996), a small sub-population within the US general population, but did not include a 345 large population-based National Health and Nutrition Examination Survey 346 (NHANES) study (Riskowski 2014), suggesting the current and prior CWP 347 systematic review focused on different study populations. 348

#### 349 4.4 Socioeconomics Position and CWP

The novelty of this meta-analysis was in the analyses of socioeconomic position 350 measures to CWP prevalence. The socioeconomic position contextual factors were the 351 HDI (HDR 2016), United Nations (UN) developed/developing country definition (UN 352 2012), and GINI index (HDR 2016). Although the HDI and UN definition of 353 developed/developing countries may appear similar, the HDI is a composite index 354 based on life expectancy, education level, and per capita income indicators, whereas 355 the UN definition of developed and developing countries is "intended to reflect basic 356 357 economic country conditions" rather than consideration of life within the country (United Nations United Nations Department of Economic and Social Affairs (US 358 DESA) 2012). 359

Within the work presented, the dichotomised UN-defined developed versus 360 developing country showed higher CWP prevalence within the developing countries 361 (p=0.04). The results of less developed countries having greater prevalence of CWP 362 aligned to results of the continuous HDI variable, which showed countries with a 363 higher HDI (more developed countries) having a lower CWP prevalence. The results 364 of a higher HDI (e.g., developed countries) associated with lower prevalence of pain 365 aligns with other studies evaluating socioeconomic position with chronic pain (Urwin 366 et al., 1998) and adds further evidence that socioeconomic position is associated with 367 health (Braveman et al., 2010b) and pain (Riskowski 2014). 368

Studies have suggested that financial strain and lower socioeconomic conditions can result in stress-induced muscular tension and pain (Soares and Jablonska 2004). At the individual-level, poor coping strategies to stress, leading to muscular tension, is believed to play a role in the higher rates of chronic pain in those in a lower socioeconomic position relative to their higher counterparts (Ridder De

2000; Roth and Geisser 2002). Extrapolating the individual-level measure of 374 socioeconomic position to the contextual population-level measure (e.g., HDI), poor 375 community and support structures that provide mechanisms to assist people in coping 376 with stress may explain the population-level association of CWP to lower 377 socioeconomic conditions. Given the low number of studies from developing and 378 lower socioeconomic countries, there is a need for epidemiological studies of chronic 379 pain in these regions to determine the global prevalence of CWP and to evaluate the 380 effect of the country's socioeconomic position with respect to CWP prevalence. 381

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## 383 **4.5 Strengths and limitations**

Although there is a relatively recent review examining global prevalence of CWP 384 (Mansfield et al., 2016), this review adds to their results by examining a number of 385 contextual factors that may impact the CWP prevalence, such as the HDI, survey 386 method, and WHO region. However, other factors not accounted in this review were 387 race and ethnicity, due to the lack of reported data. Future studies, where appropriate, 388 should include race and ethnicity information of participants as some studies have 389 suggested it may impact risk of CWP (Allison et al., 2002; Macfarlane et al., 2005). 390 Along these same lines, studies have also suggested that class or socioeconomic 391 position may also be an individual factor that relates to risk of chronic pain (Rios and 392 Zautra 2011; Urwin et al., 1998), but within the systematic review there was one 393 study that evaluated CWP by class or social position. Thus, the surrogate markers of 394 HDI and the WHO development status were used to evaluate the effect of social 395 deprivation at the country-level, with results suggesting that with greater social 396 deprivation there is greater risk of CWP. However, the country-level marker may not 397 truly represent the participants in the study, and future work should evaluate health 398

status along social strata in addition to racial and ethnic categories (Braveman et al.,2010a).

401

# 402 **4.6 Conclusion**

Results of this systematic review indicate that CWP affects one in ten individuals
globally within the general population. In 30 studies across 19 countries women were
found to have a higher CWP prevalence than men, and those in countries with a lower
HDI tended to be more likely to experience CWP than those in a high HDI country.
To further evaluate CWP, research is needed by other individual-level factors (e.g.,
race, ethnicity) with a greater range of developing and developed countries.

409

410 Authors Contributions

PA prepared the search strategy, and ran the initial search, which was confirmed by
JLR, both PA and JLR performed the search with MS resolving any issues in
decision. The manuscript was written and proofed by all three authors.

414

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# **Supporting Information**

Table S1: Strengthening the Reporting of Observational Studies in Epidemiology

(STROBE) checklist.

 Table S2: Additional study characteristics

Table S3: STROBE quality assessment for each included study

Figure F1: Search Strategy.