



HAL
open science

Metacarpophalangeal impairment in hand osteoarthritis is not rare and is associated with mechanical factors: Results from the DIGICOD hand osteoarthritis cohort

Inès Kouki, Sophie Tuffet, Michel D. Crema, Alexandra Rousseau, Pascal Richette, Maxime Dougados, Francis Berenbaum, Jérémie Sellam, Alice Courties

► To cite this version:

Inès Kouki, Sophie Tuffet, Michel D. Crema, Alexandra Rousseau, Pascal Richette, et al.. Metacarpophalangeal impairment in hand osteoarthritis is not rare and is associated with mechanical factors: Results from the DIGICOD hand osteoarthritis cohort. *Arthritis Care & Research = Arthritis Care and Research*, In press, 10.1002/acr.24642 . hal-03228453

HAL Id: hal-03228453

<https://hal.sorbonne-universite.fr/hal-03228453v1>

Submitted on 18 May 2021

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

1 **Metacarpophalangeal impairment in hand osteoarthritis is not rare and is associated**
2 **with mechanical factors: Results from the DIGICOD hand osteoarthritis cohort**

3 Running head: **Metacarpophalangeal joints in hand osteoarthritis**

4

5 Inès Kouki¹, MD, Sophie Tuffet², MsC, Michel D. Crema³, MD, Alexandra Rousseau², PhD,
6 Pascal Richette⁴, MD, PhD, Maxime Dougados⁵, MD, PhD, Francis Berenbaum¹, MD, PhD,
7 Jérémie Sellam*¹, MD, PhD, Alice Courties*¹, MD, PhD

8

9 ¹Sorbonne University, Department of Rheumatology, Saint-Antoine Hospital, Assistance
10 Publique – Hôpitaux de Paris, CRSA Inserm UMRS_938, Paris, 75012, France

11 ²Sorbonne University, Service de Pharmacologie Clinique et Plateforme de Recherche
12 Clinique de l'Est Parisien (URCEST, CRB, CRC), Saint-Antoine Hospital, Assistance
13 Publique – Hôpitaux de Paris, Paris, 75012, France

14 ³Institut d'Imagerie du Sport, Institut National du Sport, de l'Expertise et de la Performance
15 (INSEP), Paris, 75012, France

16 ⁴University of Paris, Department of Rheumatology, Lariboisière Hospital, Assistance
17 Publique – Hôpitaux de Paris, Paris, 75010, France

18 ⁵University of Paris, Department of Rheumatology, Cochin Hospital, Assistance Publique –
19 Hôpitaux de Paris, INSERM (U1153) : Clinical epidemiology and biostatistics, PRES
20 Sorbonne Paris-Cité, Paris, 75014, France

21 * These two authors contributed equally to the study

22

23 Word count for the manuscript: 3269

24 Sponsor : DIGICOD is sponsored by Assistance Publique-Hôpitaux de Paris

25 Grant research: unrestricted grant from TRB Chemedica, which did not take part in the
26 study design, collection, analysis and interpretation of data, writing of the report or the
27 decision to submit the article for publication.

28 Conflict of interest statement: The authors declare that there is no conflict of interest.

29

30 **Corresponding author:**

31 Prof. Francis Berenbaum

32 Hôpital Saint-Antoine, Service de Rhumatologie

33 184 Rue du Faubourg Saint-Antoine

34 75012 Paris, FRANCE

35 Tel: + 33 1 49 28 25 20; Fax: + 33 149 28 25 13; Email: francis.berenbaum@aphp.fr

36

37

38 **Abstract**

39 **Objective:** To determine the prevalence, distribution and characteristics associated with
40 radiographic metacarpophalangeal (MCP) osteoarthritis (OA).

41 **Methods:** This is a cross-sectional study of baseline data from the DIGItal Cohort
42 Osteoarthritis Design, a French monocentric cohort including patients with symptomatic hand
43 OA (HOA). We evaluated the prevalence of radiographic MCP OA defined as ≥ 2 MCP joints
44 with a Kellgren and Lawrence score ≥ 2 . We compared the prevalence of MCP OA in the
45 dominant and non-dominant hands. Associations between radiographic MCP OA and patient
46 characteristics were studied using univariable and multivariable logistic regression.

47 **Results:** Radiographic MCP OA was present in 138 of the 425 patients (32.5%) but was not
48 severe. Patients with MCP OA had a mean age of 69.2 ± 6.9 years, a BMI of 25 ± 4.2 kg/m²,
49 and 86.2% were women. MCP OA was more frequent in the dominant hand and
50 predominated at the 1st and 2nd MCP joints. In the multivariable analysis, MCP OA was
51 associated with older age (OR 1.05, 95%CI [1.01,1.10] for each year), manual occupation
52 (OR 3.74, 95%CI [1.21,11.54]), scaphotrapezial OA (OR 2.18, 95%CI [1.27,3.72]), and a
53 high number of proximal interphalangeal joints with radiographic OA. MCP OA was not
54 associated with metabolic syndrome or HOA symptoms.

55 **Conclusion:** In this cross-sectional study using a hospital-based HOA cohort, radiographic
56 MCP OA was frequent and associated with structural HOA features rather than with symptom
57 severity. Our results suggest that the involvement of MCP joints in HOA is predominantly
58 related to mechanical rather than systemic factors in this population.

59

60

61
62
63
64
65
66
67
68
69
70
71
72
73
74
75
76
77
78
79
80
81
82
83

Significance and innovation

- Radiographic MCP OA was found in one-third of the present cases but was not clinically and radiographically severe.
- Radiographic MCP OA is associated with radiographic severity of the proximal interphalangeal OA and with the presence of scaphotrapezial OA.
- Involvement of MCP in HOA is associated more with mechanical factors than with systemic factors in this cohort of patients with symptomatic hand OA.
- The presence of MCP OA without proximal interphalangeal OA involvement should lead us to reconsider the diagnosis of HOA.

84

85

86

87

88 **Introduction**

89 Hand osteoarthritis (HOA) affects a large part of the population. The prevalence of
90 radiographic HOA is estimated to be between 27% and 50% in the general population (1,2).
91 Moreover, symptomatic HOA affects approximately 13% of men and 26% of women over 70
92 years of age (3). The burden of HOA is important for individual-level pain and disability as
93 well as public health (4). Among HOA risk factors, some are well established (older age,
94 female, family history), while others such as obesity or metabolic syndrome remain
95 controversial (5,6) .

96 The commonly used clinical definition of HOA proposed by the American College of
97 Rheumatology (ACR) has a good radiographic correlation (94% sensitivity and 87%
98 specificity, with 78% agreement) demonstrated in secondary care population. The definition
99 takes into account the distal and proximal interphalangeal (DIP/PIP) joint involvement but
100 excludes metacarpophalangeal (MCP) swellings (7). The consensus among rheumatologists is
101 that MCP joints are not affected by OA but by inflammatory rheumatic diseases, such as
102 rheumatoid arthritis, psoriatic arthritis, or crystal arthropathies.

103 However, previous evidence suggests that MCP joints can also be affected in OA. In the
104 Framingham cohort (n=2301), radiographic MCP OA, defined by one or more MCP joints
105 with a Kellgren and Lawrence score (KL) ≥ 2 , was observed in 11.9% of men and 6.8% of
106 women (2). In the Rotterdam study (n=3906), 8.2% of people aged over 55 years and 14.8%

107 of people aged over 85 years were affected (8). In contrast, symptomatic MCP OA is less
108 common and affects about 2% of people aged over 60 years (9).

109 Very few studies have addressed the prevalence of MCP OA in individuals with OA and its
110 determinants. The role of mechanical stress may be critical because high manual strain at
111 work and high grip strength are associated with radiographic and symptomatic MCP OA (i.e.
112 farmers, dockers, pneumatic drillers, or jackhammer operators) (10–12). However, its
113 association with systemic factors has never been studied. Finally, MCP OA might be
114 associated with a more severe HOA impairment since it has been associated with hand
115 disability (2,8,13) and accelerated OA (14). The aim of this study was to determine the
116 prevalence, distribution, and features associated with radiographic MCP OA in a large
117 hospital-based HOA cohort.

118

119 **Patients and Methods**

120 Study population

121 The DIGItal Cohort Osteoarthritis Design (DIGICOD, NCT01831570) is a French
122 monocentric prospective cohort, including patients with symptomatic HOA. A detailed
123 description of the study design and data collection has been described elsewhere (15). For
124 inclusion, people aged over 35 years with symptomatic HOA defined as symptomatic HOA
125 (pain or nodes) on at least 2 joints among PIP or DIP joints or first IP joint with $KL \geq 2$,
126 and/or symptomatic OA (pain or deformation) at the thumb base with $KL \geq 2$, were eligible to
127 participate.

128 Patients with known polyarticular chondrocalcinosis or other inflammatory rheumatic
129 diseases, such as psoriatic arthritis or rheumatoid arthritis, were excluded. At baseline,
130 patients were clinically examined by a rheumatologist, completed self-reported questionnaires

131 (the Australian Canadian Osteoarthritis Hand Index [AUSCAN], Functional Index for Hand
132 Osteoarthritis [FIHOA], and the Cochin Hand Function Scale), provided a blood sample, and
133 underwent a radiograph of both hands. As previously described, this radiograph was
134 performed in a postero-anterior view, with both hands in the same film (18x24 cassette), with
135 the same device, and wrists, MCP, PIP and PID had to be well visualized. Both hands are
136 pronated and hands and wrists had to be flat, the fingers well-extended, very little apart, and
137 the second metacarpal had to be in the extension of the radius. The focal-film distance is 1
138 meter and technical parameters were 40 to 65 kV, 5-10 mAs (15).

139 Each hand joint of both hands (DIP, PIP, MCP, trapeziometacarpal [TMC], and
140 scaphotrapezial [ST]) was radiographically scored by one trained musculoskeletal radiologist
141 (MDC) blinded to the clinical data to obtain the KL (0–4 for each hand joint) and Verbruggen
142 anatomical scores (16,17). The protocol for this cross-sectional analysis of the baseline visit
143 assessments was approved by the local Ethics Board (Comité de Protection des Personnes
144 Paris Ile de France IV). All participants provided written informed consent.

145

146 Clinical and biological data collection

147 At baseline, the following demographic and biological variables were collected: age, gender,
148 occupation, body mass index (BMI), alcohol consumption, smoking, diabetes, hypertension
149 (HTN), dyslipidemia, cardiovascular diseases, metabolic syndrome as defined by the National
150 Cholesterol Education Program–Adult Treatment Panel III (NCEP ATP III) criteria (18),
151 menopausal status, family history of OA, biological inflammation (C-reactive protein [CRP]
152 ≥ 5 mg/L), hyperferritinemia (defined by ferritinemia > 300 $\mu\text{g/L}$ in men or > 200 time of
153 evolution of HOA, and use of analgesics or non-steroidal anti-inflammatory drugs (NSAIDs).
154 All the definitions used for these characteristics are shown in **Supplementary Table 1**.
155 Current or previous occupations were classified using the *Institut National de la Statistique et*

156 *des Etudes Economiques* classification and grouped into three categories: (1) manual
157 professions; (2) intermediate professions and employees; and, (3) intellectual professions. In
158 this study, we call “manual workers” farmers, craftsmen, traders and workers.

159

160 Radiographic metacarpophalangeal osteoarthritis

161 A MCP joint with OA was defined by a KL score ≥ 2 and radiographic MCP OA was defined
162 by at least two MCP joints with a KL score ≥ 2 among the MCP joints of both hands.

163

164 Radiographic and clinical severity

165 Radiographic severity excluded the MCP joints since they contributed to the outcome and was
166 defined as the sum of the KL score at the PIP, DIP, ST, and TMC joints (total score between 0
167 and 88); the number of PIP, DIP, ST, and TMC joints with KL ≥ 2 , each and together; and the
168 number of erosive joints among the PIP and DIP joints defined by phase E or R of the
169 Verbruggen radiographic score.

170 Clinical severity was defined by an AUSCAN pain score $\geq 40/100$, FIHOA score $> 5/30$,
171 Cochin Hand Function Scale score between 0 and 90, the visual analog scale (VAS) score for
172 hand pain $\geq 40/100$, number of painful joints on applying pressure, swelling joints among the
173 PIP, DIP, MCP joints and thumb base, and grip and pinch strengths on the dominant hand.

174

175 Statistical analysis

176 The prevalence of radiographic MCP OA defined by at least two MCP joints with a KL score
177 ≥ 2 was evaluated as well as the prevalence of patients with at least two MCP joints with a
178 KL score ≥ 3 or 4. The prevalence of painful MCP joints spontaneously or under pressure was
179 assessed too. Proportions of patients with MCP OA were compared between the dominant and
180 non-dominant hand using a McNemar test for paired data.

181
182 Baseline characteristics were reported according to the presence or absence of radiographic
183 MCP OA using frequencies and percentages for categorical variables and mean \pm standard
184 deviation (SD) or median and interquartile range (IQR) for continuous variables (according to
185 the distribution). Differences in proportions between groups were reported with 95%
186 confidence intervals (CIs), estimated by the exact method, and the difference of means
187 between groups was reported with a 95% CI estimated by the pooled or Satterthwaite method
188 depending on whether variances were equal or unequal between groups, respectively.

189
190 Associations between the presence of radiographic MCP OA and demographic characteristics
191 and radiographic and clinical severity were analyzed using logistic regression. A
192 multivariable model was built using covariates selected from the univariable analysis
193 ($p < 0.20$) and was systematically adjusted for age, gender, BMI, and family history of OA.
194 The final model was determined using a backward elimination method. Results were
195 expressed as odds ratios (OR) with 95% CIs. All analyses were performed using SAS version
196 9.4 statistical software (SAS Institute Inc.). All tests were two-sided, and a p -value < 0.05
197 indicated statistical significance.

198

199 **Results**

200 In total, 436 patients were included in the DIGICOD cohort between April 2013 and June
201 2017 from the rheumatology department of Saint-Antoine Hospital (Assistance Publique-
202 Hôpitaux de Paris). Eleven participants were excluded (one withdrew consent, 3 had
203 unavailable radiographs, 6 did not meet the selection criteria, one with missing KL score).
204 Finally, data from 425 patients were analyzed in the present study.

205

206 MCP OA is frequent, predominates on the dominant hand but is radiographically and
207 clinically not severe.

208

209 Among the 425 patients, 49.4% had at least one MCP KL ≥ 2 , 16.7% had at least one MCP
210 joint KL ≥ 3 , and only 3.3% KL ≥ 4 . The median MCP KL total score (extremes 0–40) was
211 3.0 (IQR: 1.0 to 6.0). MCP OA was more frequent in the dominant hand (40.6% vs. 34.7%,
212 $p < 0.05$) and was predominant in the thumb and index finger (20.2% and 20.7%, respectively)
213 (**Figure 1**). Spontaneous painful MCP joint was rare ($n=29/424$, 6.8%), while 139/424
214 (32.8%) patients had at least one painful MCP joint under pressure. Only 2 patients (0.5%)
215 had one erosive MCP joint and none had more. In total, 138 patients (32.5%) had at least 2
216 MCP joints with a KL ≥ 2 and were considered as patients with radiographic MCP OA for
217 subsequent analyses.

218

219 Patients with MCP OA are older, have a more manual job, and are radiographically more
220 severe

221

222 Patients with or without radiographic MCP OA were similar in their BMI (25.0 ± 4.2 vs.
223 25.2 ± 4.4) and OA family history (66.9% vs. 70.5%). In addition, the percentages of patients
224 with metabolic syndrome or its components were similar between the two groups. Patients
225 with MCP OA were slightly older than those without MCP OA (69.2 ± 6.9 vs. 65.5 ± 7.3
226 years). Although there was a predominance of women in both groups, proportion of women
227 appeared to be slightly more frequent among individuals with MCP OA (86.2% vs. 82.6%).
228 There were twice as many patients with past or present work involving hands among the
229 patients with MCP OA (10.4% vs. 4.6%) compared to patients without MCP OA. In contrast,
230 intellectual professions were less frequent among patients with MCP OA. There were no

231 differences between the two groups in terms of iron parameters, CRP level, and calcemia
232 (**Table 1**).

233

234 In the MCP OA group, the mean KL score excluding MCP joints (0–88) was 50.3 ± 11.9
235 compared to 37.9 ± 15.3 in the group without MCP OA. The presence of a high number of
236 PIP joints with OA (9 or 10) was much more frequent in patients with radiographic MCP OA,
237 while the presence of a low number of PIP joints with OA (0 to 4) was much more frequent in
238 patients without MCP OA. In addition, 56.2% of the patients with MCP OA had erosive OA
239 versus 39.2% of patients without MCP OA.

240 Concerning the clinical severity, 59.1% of patients with MCP OA had a VAS score $\geq 40/100$
241 compared to 57.2% of patients without MCP OA. There was no evident difference in the
242 function scores on the FIHOA and Cochin scales (**Table 2**).

243

244 Age, socio-professional category, number of PIP and ST with OA are independently
245 associated with MCP OA

246

247 Duration of HOA, number of erosive joints, DIP or TMC joints with OA, painful joints on
248 applying pressure, number of joints with synovitis, and the prehension strength on the
249 dominant hand were significantly associated with MCP OA in the univariable analysis
250 ($p < 0.20$) but the association did not remain in multivariable analysis. No
251 biological factors were associated with MCP OA (calcemia, CRP, hyperferritinemia, or high
252 TSC). Neither BMI nor metabolic syndrome was associated with MCP OA (**Table 3**).

253 In the multivariable analysis, radiographic MCP OA was associated with higher age
254 (OR=1.05; 95% CI [1.01, 1.10]), manual works (OR=3.74, 95% CI [1.21, 11.54] for manual
255 workers versus intellectual professionals), and higher radiographic severity of HOA (**Table**

256 3). Patients with 5 to 8 PIP joints with OA, representing 34.1 % of the entire population, were
257 7.83 (95% CI [3.58, 17.16]) times more at risk for MCP OA than patients with 0 to 4 PIP
258 joints with OA. In addition, those with 9 to 10 PIP joints with OA (30% of the population)
259 were 14.29 (95% CI [6.46, 31.64]) times more at risk for MCP OA compared to patients with
260 0 to 4 PIP joints with OA. Lastly, ST impairment was also associated with radiographic MCP
261 OA with an OR of 2.18 (95% CI [1.27, 3.72]).

262

263 **Discussion**

264

265 Using a large hospital-based cohort of patients with symptomatic IP and/or thumb base HOA,
266 we found that radiographic MCP OA is frequent (32.5%) and predominates in the thumb and
267 the index finger of the dominant hand. Radiographic MCP OA was independently associated
268 with older age, manual work, PIP OA radiographic severity, and presence of ST OA, but not
269 with systemic or metabolic factors.

270

271 Here, we found that the prevalence of radiographic MCP OA is higher than that reported in
272 previous studies, such as the Framingham study (11.9% of men and 6.8% of women in
273 Framingham) (2). One explanation is that we studied MCP OA in a symptomatic HOA
274 population (defined by PIP/DIP or thumb OA) (i.e., hospital-based cohort), while the
275 Framingham cohort is a general population sample (i.e., a population-based cohort). Indeed,
276 in a recent study, MCP impairment was studied using magnetic resonance imaging (MRI) in
277 HOA patients compared to controls showing that 21/81 MCP joints had a loss of cartilage,
278 including 5 with areas of full-thickness loss (19). This prevalence is close to that observed in
279 our study. MCP OA is more frequent than expected and probably underestimated, especially
280 when it is studied with radiographs only. Although the prevalence of MCP OA was high, the

281 severity of the radiographic impairment was low (low prevalence of MCP joints at KL 3 or 4
282 and low median MCP KL sum) and so was the clinical burden (few spontaneously painful
283 MCP joints).

284

285 We found that older age was associated with MCP OA (5% increase in risk per year) but not
286 with gender. Conversely, in the Framingham cohort and the Zoetermeer study, MCP OA was
287 slightly more frequent in men and at a younger age than other HOA localizations. For
288 example, in the 40–44 year age group, 9.6% of men had MCP OA versus 8.2% of women
289 (2,20). These discrepancies can also be explained by the differences in the sample populations
290 (individuals with HOA vs. general population). The presence of confounding factors in the
291 Framingham study from a random general population would likely include more manual
292 workers among men, explaining this difference. Radiographic MCP OA did not seem to be
293 independently associated with disease duration, as may have been expected, probably because
294 disease duration is strongly associated with age.

295

296 In this population of symptomatic hand OA patients, MCP joints impairment was more
297 frequent and independently associated with manual jobs, which is consistent with the
298 literature (10–12). Indeed, some observations among samples up to 200 patients found an
299 association between manual works (farmers, cotton workers, workers using vibrating-tools)
300 and MCP OA in terms of frequency and of severity (10,21,22). Moreover, the distribution of
301 MCP OA was consistent with previous publications, since MCP OA was more frequently
302 described in the thumb, index and middle fingers probably because mechanical factors are
303 stronger at these fingers (2,8). The higher frequency of MCP OA in the dominant hand and
304 the association with manual occupations suggest that mechanical factors are strongly
305 implicated in the development of MCP OA, which is in accordance with studies that showed

306 that higher grip strengths can be associated with the occurrence of MCP OA later (11,12).
307 However, more specific mechanical factors such as quantitative data about the manual
308 occupation (mechanical loading or repetitive tasks), length of time in the employment or
309 manual hobbies were not available which limits the accuracy of manual activities assessment.

310

311 In contrast, systemic factors such as BMI, diabetes, metabolic syndrome, and elevated CRP
312 were not associated with MCP OA among these hand OA patients. The absence of such
313 associations reinforces the importance of mechanical factors in MCP OA and suggests that
314 MCP joints are more responsive to mechanical stress than PIP or DIP (5,23). There was no
315 association with hyperferritinemia, elevated TSC, or hypercalcemia, but it is expected because
316 the study population was primary OA patient. Some other systemic features may have
317 influenced MCP OA such as genetic factors but they were not investigated here..

318

319 Finally, MCP OA was associated with HOA radiographic severity, especially with PIP OA
320 severity and with the presence of ST OA. Although the ST joint is known to be related to
321 mechanical forces, it is independently associated with MCP OA, even when adjusted on
322 manual works. This is consistent with data showing that HOA of other joints co-occurred in
323 86% of patients with MCP involvement (8). Thus, OA in the MCP joints could be part of a
324 generalized and evolved form of HOA. Since only 2.7% and 9.5% of patients with MCP OA
325 have less than 3 DIP with OA and less than 4 PIP with OA, MCP OA without PIP or DIP OA
326 should lead us to reconsider the diagnosis of primary HOA.

327

328 While MCP OA was associated with symptoms in the univariable analysis, there was no
329 association between MCP OA and HOA clinical symptoms (pain, loss of function or loss of
330 grip strength) in the multivariable analysis. The reason may be because symptoms are closely

331 related to radiographic severity. This means that although radiographic MCP OA is common,
332 its clinical impact is limited and is mediated by the severity of the PIP or thumb base OA.

333

334 The strengths of this study should be considered. First, DIGICOD is a large cohort of
335 individuals with symptomatic HOA, defined by worldwide validated criteria. The radiographs
336 were analyzed by a trained radiologist blinded to the patient characteristics. Second, the
337 definition of MCP OA was stringent compared to other studies because we used the presence
338 of at least 2 MCP joints with $KL \geq 2$ (and not at least 1 MCP joints with $KL \geq 2$) to be more
339 specific in the association studied. Finally, patient characteristics were defined using
340 composite criteria (patient interview, clinical examination, and biological samples) thereby
341 strengthening the results.

342

343 However, our study had some limitations. First, it was cross-sectional, that limits the
344 possibility to demonstrate a causality link between MCP OA and manual activity. Second, it
345 was conducted in a symptomatic HOA population, that limits the generalizability of the
346 findings to asymptomatic HOA patients. It is particularly true for the high prevalence, which
347 is not what is observed in general population. Then, it would have been pertinent to conduct
348 sensitivity analysis using more stringent criteria, but it was technically impossible because of
349 the low number of patients with MCP joints KL 3 and 4. Moreover, because it is a
350 symptomatic hand OA cohort, it was predominantly female population, which prevented us to
351 describe MCP OA among men. Finally, because of missing data, the analyzed population was
352 smaller than the included population, with a slight overrepresentation of diabetic and
353 hypertensive individuals as well as those with metabolic syndrome. However, since these
354 factors were not associated with MCP OA, their influence over the results would be unlikely.

355

356 In conclusion, radiographic MCP OA is frequent in patients with symptomatic PIP/DIP or
357 thumb base OA but is clinically rarely severe. Among these patients, the presence of
358 radiographic MCP OA is cross-sectionally associated with mechanical factors rather than
359 systemic factors such as metabolic factors and is especially associated with severe
360 radiographic PIP OA. The presence of MCP OA without PIP OA involvement should lead us
361 to reconsider the diagnosis of HOA, which is consistent with the literature. Further
362 prospective studies are necessary to confirm longitudinally these cross-sectional results.

363

364 Acknowledgments: All the patients from the DIGICOD cohort who participated. All the
365 investigators who managed the patients' recruitments, and the monitoring of clinical visits
366 (Dr. Camille Deprouw, Dr. Sandra Desouches, Dr. Ariane Do, Dr. Emeline Gaigneux, Dr.
367 Camille Glanowski, Dr. Karine Louati, Dr. Stéphanie Malbos, Dr. Sabine Trelu, all from the
368 rheumatology department of AP-HP Saint-Antoine Hospital), the staff members of the
369 URCEST and the Centre des Ressources Biologiques, AP-HP Saint-Antoine Hospital.

370 References

- 371 1. Lawrence RC, Felson DT, Helmick CG, Arnold LM, Choi H, Deyo RA, et al. Estimates of
372 the prevalence of arthritis and other rheumatic conditions in the United States. Part II.
373 *Arthritis Rheum* 2008;58:26–35.
- 374 2. Haugen IK, Englund M, Aliabadi P, Niu J, Clancy M, Kvien TK, et al. Prevalence,
375 incidence and progression of hand osteoarthritis in the general population: the Framingham
376 Osteoarthritis Study. *Ann Rheum Dis* 2011;70:1581–1586.
- 377 3. Zhang Y, Niu J, Kelly-Hayes M, Chaisson CE, Aliabadi P, Felson DT. Prevalence of
378 symptomatic hand osteoarthritis and its impact on functional status among the elderly: The
379 Framingham Study. *Am J Epidemiol* 2002;156:1021–1027.
- 380 4. Michon M, Maheu E, Berenbaum F. Assessing health-related quality of life in hand

381 osteoarthritis: a literature review. *Ann Rheum Dis* 2011;70:921–928.

382 5. Yusuf E, Nelissen RG, Ioan-Facsinay A, Stojanovic-Susulic V, DeGroot J, Osch G van, et
383 al. Association between weight or body mass index and hand osteoarthritis: a systematic
384 review. *Ann Rheum Dis* 2010;69:761–765.

385 6. Haugen IK, Magnusson K, Turkiewicz A, Englund M. The Prevalence, Incidence, and
386 Progression of Hand Osteoarthritis in Relation to Body Mass Index, Smoking, and Alcohol
387 Consumption. *J Rheumatol* 2017;44:1402–1409.

388 7. Altman R, Alarcón G, Appelrouth D, Bloch D, Borenstein D, Brandt K, et al. The
389 American College of Rheumatology criteria for the classification and reporting of
390 osteoarthritis of the hand. *Arthritis Rheum* 1990;33:1601–1610.

391 8. Dahaghin S, Bierma-Zeinstra SMA, Ginai AZ, Pols H a. P, Hazes JMW, Koes BW.
392 Prevalence and pattern of radiographic hand osteoarthritis and association with pain and
393 disability (the Rotterdam study). *Ann Rheum Dis* 2005;64:682–687.

394 9. Zhang Y, Xu L, Nevitt MC, Niu J, Goggins JP, Aliabadi P, et al. Lower prevalence of hand
395 osteoarthritis among Chinese subjects in Beijing compared with white subjects in the United
396 States: the Beijing Osteoarthritis Study. *Arthritis Rheum* 2003;48:1034–1040.

397 10. Williams WV, Cope R, Gaunt WD, Adelstein EH, Hoyt TS, Singh A, et al.
398 Metacarpophalangeal arthropathy associated with manual labor (Missouri metacarpal
399 syndrome). Clinical radiographic, and pathologic characteristics of an unusual degeneration
400 process. *Arthritis Rheum* 1987;30:1362–1371.

401 11. Chaisson CE, Zhang Y, Sharma L, Kannel W, Felson DT. Grip strength and the risk of
402 developing radiographic hand osteoarthritis: results from the Framingham Study. *Arthritis*
403 *Rheum* 1999;42:33–38.

404 12. Chaisson CE, Zhang Y, Sharma L, Felson DT. Higher grip strength increases the risk of
405 incident radiographic osteoarthritis in proximal hand joints. *Osteoarthr Cartil* 2000;8 Suppl

406 A:S29-32.

407 13. Schaefer LF, McAlindon TE, Eaton CB, Roberts MB, Haugen IK, Smith SE, et al. The
408 associations between radiographic hand osteoarthritis definitions and hand pain: data from the
409 osteoarthritis initiative. *Rheumatol Int* 2018;38:403–413.

410 14. Davis JE, Schaefer LF, McAlindon TE, Eaton CB, Roberts MB, Haugen IK, et al.
411 Characteristics of Accelerated Hand Osteoarthritis: Data from the Osteoarthritis Initiative. *J*
412 *Rheumatol* 2019;46:422–428.

413 15. Sellam J, Maheu E, Crema MD, Touati A, Courties A, Tuffet S, et al. The DIGICOD
414 cohort: A hospital-based observational prospective cohort of patients with hand
415 osteoarthritis - methodology and baseline characteristics of the population. *Joint Bone Spine*
416 2021;88:105171.

417 16. Kellgren JH, Lawrence JS. Radiological assessment of osteo-arthrosis. *Ann Rheum Dis*
418 1957;16:494–502.

419 17. Verbruggen G, Veys EM. Numerical scoring systems for the anatomic evolution of
420 osteoarthritis of the finger joints. *Arthritis Rheum* 1996;39:308–320.

421 18. Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in
422 Adults. Executive Summary of The Third Report of The National Cholesterol Education
423 Program (NCEP) Expert Panel on Detection, Evaluation, And Treatment of High Blood
424 Cholesterol In Adults (Adult Treatment Panel III). *JAMA* 2001;285:2486–2497.

425 19. Saltzherr MS, Muradin GSR, Haugen IK, Selles RW, Neck JW van, Coert JH, et al.
426 Cartilage evaluation in finger joints in healthy controls and early hand osteoarthritis patients
427 using high-resolution MRI. *Osteoarthr Cartil* 2019;27:1148–1151.

428 20. Saase JL van, Romunde LK van, Cats A, Vandenbroucke JP, Valkenburg HA.
429 Epidemiology of osteoarthritis: Zoetermeer survey. Comparison of radiological osteoarthritis
430 in a Dutch population with that in 10 other populations. *Ann Rheum Dis* 1989;48:271–280.

- 431 21. Lawrence JS. Rheumatism in cotton operatives. *Br J Ind Med* 1961;18:270–276.
- 432 22. Gemne G, Saraste H. Bone and joint pathology in workers using hand-held vibrating
- 433 tools. An overview. *Scand J Work Environ Health* 1987;13:290–300.
- 434 23. Puenpatom RA, Victor TW. Increased prevalence of metabolic syndrome in individuals
- 435 with osteoarthritis: an analysis of NHANES III data. *Postgrad Med* 2009;121:9–20.

436 **Tables and figures**

437 **Table 1.** Characteristics of patients according to the presence or absence of MCP OA

Variables	No MCP		MCP osteoarthritis		Difference (95% CI)
	osteoarthritis		n=138		
	n*	n=287	n*		
Age at inclusion (years), mean±SD	287	65.5±7.3	138	69.2±6.9	3.7 (2.2 to 5.1)
Women, No. (%)	287	237 (82.6)	138	119 (86.2)	3.7 (-3.6 to 10.9)
Body mass index (kg/m ²), mean±SD	285	25.2±4.4	133	25.0±4.2	-0.1 (-1.0 to 0.8)
Family history of osteoarthritis, No. (%)	278	196 (70.5)	133	89 (66.9)	-3.6 (-13.2 to 6.0)
Socio-professional category, No. (%)	282		135		
Manual works		13 (4.6)		14 (10.4)	5.8 (0.06 to 11.5)
Intermediate professions and employees		107 (37.9)		63 (46.7)	8.7 (-1.4 to 18.9)
Intellectual professions		162 (57.4)		58 (43.0)	-14.5 (-24.6 to -4.3)
Time of evolution of hand osteoarthritis (years), No. (%)	285		136		
< 5 years		70 (24.6)		14 (10.3)	-14.3 (-21.4 to -7.1)
5-15 years		143 (50.2)		68 (50.0)	-0.2 (-10.4 to 10.0)
> 15 years		72 (25.3)		54 (39.7)	14.4 (4.8 to 24.1)
Current tobacco consumption, No. (%)	285	21 (7.4)	134	8 (6.0)	-1.4 (-6.4 to 3.6)
Current alcohol consumption, No. (%)	285	215 (75.4)	134	109 (81.3)	5.9 (-2.4 to 14.2)
Diabetes, No. (%)	287	24 (8.4)	138	9 (6.5)	-1.8 (-7.1 to 3.4)
Hypertension (HTN), No. (%)	287	155 (54.0)	138	82 (59.4)	5.4 (-4.6 to 15.4)
Cardiovascular disease except AHT, No. (%)	287	13 (4.5)	138	8 (5.8)	1.3 (-3.3 to 5.8)
LDL (g/L), mean±SD	277	1.4±0.4	134	1.4±0.4	0.009 (-0.07 to 0.09)
Metabolic syndrome, No. (%)	279	99 (35.5)	134	51 (38.1)	2.6 (-7.4 to 12.5)
CRP, No. (%)	237		112		
< 5 mg/L		213 (89.9)		101 (90.2)	0.3 (-6.4 to 7.0)
≥5 mg/L		24 (10.1)		11 (9.8)	-0.3 (-7.0 to 6.4)
Hyper-ferritinemia, No. (%)	253	28 (11.1)	121	12 (9.9)	-1.1 (-7.7 to 5.4)
TSC>45%, No. (%)	277	8 (2.9)	134	5 (3.7)	0.8 (-2.9 to 4.6)
Calcemia (mmol/L), mean±SD	279	2.4±0.1	135	2.4±0.1	0.003 (-0.02 to 0.02)
Current analgesics or NSAIDs consumption, No. (%)	287	149 (51.9)	138	80 (58.0)	6.1 (-4.0 to 16.1)

438 * number of patients with available data. MCP = metacarpophalangeal joints. HTN =
439 Hypertension. LDL = Low density lipid. CRP = C reactive protein. NSAID = non-steroidal
440 anti-inflammatory drugs. TSC = transferrin saturation coefficient

441

442

443 **Table 2.** Clinical and radiographic severity of patients according to the presence or absence of
 444 MCP OA.

	No MCP osteoarthritis (n=287)		MCP osteoarthritis (n=138)		Differences (95% CI)
	n*		n*		
RADIOGRAPHIC SEVERITY					
Sum of KL score except MCP (0-88), mean±SD	276	37.9 ± 15.3	135	50.3 ± 11.9	12.4 (9.7 to 15.1)
Number of joints KL ≥ 2 excluding MCP (0-22), median (IQR)	276	13.0 (8.0 -17.0)	135	18.0 (14.0 -20.0)	
Number of erosive joints among PIP and DIP (0-18), No. (%)	281		137		
0		171 (60.9)		60 (43.8)	
1-3		82 (29.2)		47 (34.3)	
4 et +		28 (10.0)		30 (21.9)	
Number of DIP KL ≥ 2 (0-8), No. (%)	282		138		
0-3		48 (17.0)		4 (2.9)	
4-6		89 (31.6)		25 (18.1)	
7-8		145 (51.4)		109 (79.0)	
Number of PIP KL ≥ 2 (0-10), No. (%)	285		137		
0-4		138 (48.4)		13 (9.5)	
5-8		92 (32.3)		52 (38.0)	
9-10		55 (19.3)		72 (52.6)	
Number of ST KL ≥ 2 (0-2), No. (%)	284		137		
0		172 (60.6)		52 (38.0)	
1		49 (17.3)		32 (23.4)	
2		63 (22.2)		53 (38.7)	
Number of TMC KL ≥ 2 (0-2), No. (%)	284		135		
0		93 (32.7)		36 (26.7)	
1		56 (19.7)		17 (12.6)	
2		135 (47.5)		82 (60.7)	
Sum of KL score at TMC + ST (0-16), median (IQR)	283	5.0 (2.0 - 9.0)	135	8.0 (4.0 - 10.0)	
CLINICAL SEVERITY					
AUSCAN pain score ≥ 40/100, No. (%)	271	69 (25.5)	124	26 (21.0)	4.5 (-4.4 to 13.3)
FIHOA score > 5/30, No. (%)	272	129 (47.4)	132	73 (55.3)	-7.9 (-18.2 to 2.5)
Cochin function hand scale (0-90), median (IQR)	274	5.0 (1.0 - 13.0)	132	6.0 (2.0 - 15.0)	
Pain VAS (rest or activity) ≥ 40/100, No. (%)	285	163 (57.2)	137	81 (59.1)	-1.9 (-12.0 to 8.1)
Painful joints on pressure (0-30), median (IQR)	287	3.0 (2.0 - 6.0)	137	3.0 (1.0 - 7.0)	
Swelling joints (0-30), median (IQR)	287	0.0 (0.0 - 1.0)	136	1.0 (0.0 - 3.0)	
Grip strength on dominant hand (kg), median (IQR)	287	24.0 (20.0 - 31.0)	136	22.0 (18.0 - 28.0)	
Pinch strength on dominant hand (kg), median (IQR)	280	5.4 (4.3 - 6.7)	130	5.3 (4.4 - 6.5)	

445

446 * number of patients with available data. VAS = Visual analogic scale, DIP = distal inter-

447 phalangeal joints, PIP = proximal inter-phalangeal joints, ST = scaphotrapezial joints, TMC =

448 trapezometacarpal joint, KL = Kellgren and Lawrence, AUSCAN = Australian Canadian
449 osteoarthritis hand index, FIHOA = Functional Index for Hand osteoarthritis, KL = Kellgren
450 and Lawrence score.

451
452
453
454
455
456
457
458
459
460
461
462
463
464
465
466
467
468
469
470
471
472
473
474
475
476
477
478
479
480

481

482

483 **Table 3.** Univariable and multivariable analysis of factors associated with MCP OA (n=356,

484 population with available values for all selected variables in univariable analysis)

	Univariable analysis		Multivariable analysis*	
	OR [IC à 95%]	p-value	OR [IC à 95%]	p-value
Age at inclusion (by year)	1.09 [1.05-1.13]	<.0001	1.05 [1.01-1.10]	0.0194
Gender		0.2947		0.2505
Men	1	.	1	.
Women	1.40 [0.74-2.65]	.	1.59 [0.72-3.48]	.
Body Mass Index (kg/m²)	1.00 [0.95-1.05]	0.9912	0.99 [0.93-1.05]	0.7547
Family history of osteoarthritis		0.7893		0.8558
No	1	.	1	.
Yes	0.94 [0.58-1.51]	.	1.05 [0.59-1.87]	.
Socio-professional category		0.0114		0.0158
Intellectual professions	1	.	1	.
Manual works	3.16 [1.31-7.62]	0.0104	3.74 [1.21-11.54]	0.0218
Intermediate professions and employees	1.67 [1.05-2.68]	0.0320	1.85 [1.07-3.21]	0.0280
Time of evolution of hand osteoarthritis (years)		<.0001		
< 5 years	1	.		
5-15 years	4.64 [1.32-16.39]	0.0170		
> 15years	9.67 [2.91-32.14]	0.0002		
Number of erosive joints among PIP and DIP (0-18)		0.0027		
0	1	.		
1-3	1.58 [0.94-2.64]	0.0815		
4 and +	2.94 [1.57-5.52]	0.0008		
Number of DIP KL ≥ 2 (0-8)		<.0001		
0-3	1	.		
4-6	3.37 [1.11-10.25]	0.0323		
7-8	9.02 [3.16-25.77]	<.0001		
Number of PIP KL ≥ 2 (0-8)		<.0001		<.0001
0-4	1	.	1	.
5-8	6.78 [3.23-14.23]	<.0001	7.83 [3.58-17.16]	<.0001
9-10	15.34 [7.22-32.59]	<.0001	14.29 [6.46-31.64]	<.0001
Number of ST KL ≥ 2 (0-2)		<.0001		0.0045
0	1	.	1	.
1-2	2.60 [1.64-4.12]	.	2.18 [1.27-3.72]	.
Number of TMC KL ≥ 2 (0-2)		0.0430		
0-1	1	.		
2	1.59 [1.01-2.51]	.		
Painful joints at pressure (0-30)		0.0232		
0-1	1	.		
2-3	0.45 [0.24-0.82]	0.0097		
4-6	0.48 [0.25-0.93]	0.0289		
7 and +	0.86 [0.47-1.59]	0.6319		
Swelling joints (0-30)		0.0013		

0-2	1	.
3 and +	2.49 [1.43-4.35]	.
Grip strength on dominant hand		0.0143
> 30kg	1	
≤ 30kg	2.12 [1.16-3.87]	.

485

486 * Systematic adjustment on age, gender, BMI and family history of OA. DIP = distal inter-
487 phalangeal joints, PIP = proximal inter-phalangeal joints, ST = scaphotrapezial joints, TMC =
488 trapezometacarpal joint KL = Kellgren and Lawrence score

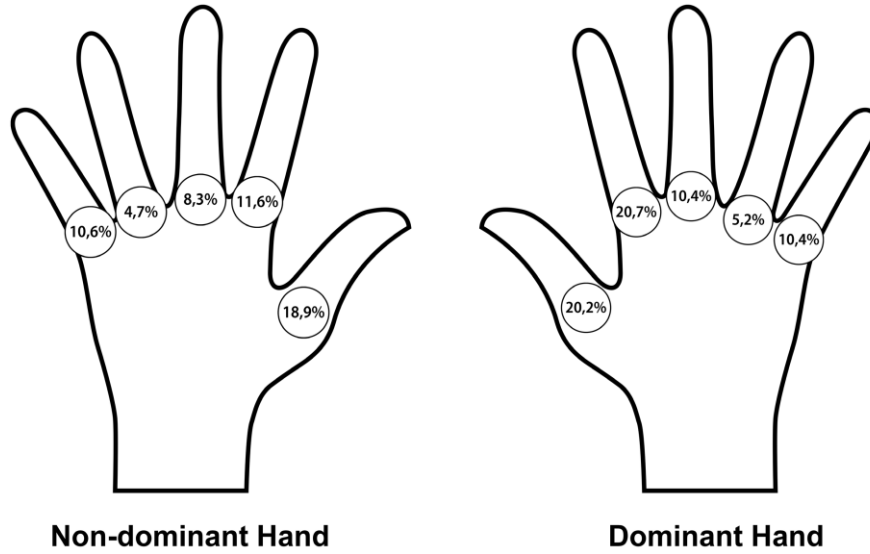
489
490
491
492
493
494
495
496
497
498
499
500
501
502
503
504
505
506
507
508
509
510
511
512
513
514
515
516
517
518
519
520
521
522
523
524
525

Socio-professional categories	<ul style="list-style-type: none"> - Manual workers : Farmers, craftsmen, traders and workers <ul style="list-style-type: none"> o Farmers o Craftsmen o Merchants and equivalent o Skilled workers o Unskilled workers o Agricultural workers - Intermediate professions and employees <ul style="list-style-type: none"> o Intermediate professions in education, health, public service and similar o Intermediate administrative and commercial professions of companies o Technicians o Foremen, supervisors o Public service employees o Commercial employees o Personal of direct services to individuals - Intellectual professions <ul style="list-style-type: none"> o Entrepreneurs with 10 or more employees o Corporate administrative employees o Liberal and assimilated professions o Public service executives, intellectual and artistic professions o Company executives
Diabetes	<ul style="list-style-type: none"> - declared diabetes, - anti-diabetic treatment, - fasting blood sugar > 1,26 g/L, or - HbA1c \geq 6,5%.
Hypertension	<ul style="list-style-type: none"> - declared high blood pressure, - anti-hypertensive treatment, - systolic blood pressure \geq 140 mmHg, or - diastolic blood pressure \geq 90 mmHg.
Cardiovascular diseases	<ul style="list-style-type: none"> - declared heart failure, - declared history of myocardial infarction or ischemic heart disease, - declared history of ischemic stroke or transient ischemic attack, or - declared obliterating arteriopathy of the lower limbs.
Metabolic syndrome is defined by Adult Treatment Panel III criteria, i.e. three or more of these components:	<ul style="list-style-type: none"> - Increased waist circumference (\geq102cm in men, >88cm in women) - Elevated triglycerides: \geq1,5 g/L or drug treatment for elevated triglycerides - Reduced HDL-cholesterol: < 0,4 g/L in men, < 0,5 g/L in women or drug treatment - Elevated blood pressure \geq 130/85 or drug treatment for elevated blood pressure - Elevated fasting glucose \geq 1g/L or drug treatment for elevated glucose
Dyslipidemia	LDL cholesterol in continuous variable.
Hyperferritinemia	<ul style="list-style-type: none"> - > 300 ug/L in men - > 200 ug/l in women.

Current consumption of analgesics or NSAID	Current consumption of acetaminophen, systemic or topic NSAIDs, and weak opioids.
--	---

526 **Supplementary Table 1. Definitions**

527 **Figure 1**



528

529 **Figure 1:** Distribution of MCP osteoarthritis (KL ≥ 2) according to the fingers and the
530 dominant hand.