scientific reports



OPEN The impact of COVID-19 on patients with chronic pain seeking care at a tertiary pain clinic

Maisa S. Ziadni^{®™}, Dokyoung S. You[®], Eric M. Cramer[®], Steven R. Anderson, Gabrielle Hettie, Beth D. Darnall & Sean C. Mackey

Empirical data on the health impacts of the COVID-19 pandemic remain scarce, especially among patients with chronic pain. We conducted a cross-sectional study matched by season to examine patient-reported health symptoms among patients with chronic pain pre- and post-COVID-19 pandemic onset. Survey responses were analyzed from 7535 patients during their initial visit at a tertiary pain clinic between April 2017-October 2020. Surveys included measures of pain and painrelated physical, emotional, and social function. The post-COVID-19 onset cohort included 1798 initial evaluations, and the control pre-COVID-19 cohort included 5737 initial evaluations. Patients were majority female, White/Caucasian, and middle-aged. The results indicated that pain ratings remained unchanged among patients after the pandemic onset. However, pain catastrophizing scores were elevated when COVID-19 cases peaked in July 2020. Pain interference, physical function, sleep impairment, and emotional support were improved in the post-COVID-19 cohort. Depression, anxiety, anger, and social isolation remained unchanged. Our findings provide evidence of encouraging resilience among patients seeking treatment for pain conditions in the face of the COVID-19 pandemic. However, our findings that pain catastrophizing increased when COVID-19 cases peaked in July 2020 suggests that future monitoring and consideration of the impacts of the pandemic on patients' pain is warranted.

The COVID-19 pandemic has prompted healthcare organizations to make drastic changes in the delivery of pain care services to minimize virus exposure risks. For instance, elective medical procedures, in-person visits, and multidisciplinary services are being limited, while telehealth, digital treatment, and virtual reality expanded. These rapid changes require healthcare clinicians and patients to adjust to many unknowns. Understanding the impact of the pandemic on the health status of patients seeking treatment for pain conditions is critical to help identify areas of growing need, evaluate any gaps in services, and ensure optimal pain care.

To date, empirical data on the health impacts of the pandemic are scarce. Some evidence suggests that humans are overall resilient and are adjusting well during the pandemic²⁻⁴, while others show that the pandemic has adversely impacted adults' physical and mental health. Such adverse impacts include a significant reduction in daily physical activity⁵, increase in depression symptoms^{6,7}, and more frequent alcohol use⁸. These adverse impacts are expected to be even more evident in medically vulnerable populations. A survey study on patients with cancer found a significant reduction in cognitive and social function during the COVID-19 pandemic9. Patients with chronic pain conditions, which may be triggered or exacerbated by psychosocial stressors^{10,11}, are likely to be similarly vulnerable to the adverse impacts of the pandemic¹². Two Pan-Canadian studies found that adults living with chronic pain reported increased pain, increased psychological distress and lack of access to treatment 13,14. Similarly, a US-based study found increased pain intensity and interference among patients with chronic pain¹⁵, within a limited time-period following COVID-19 social distancing mandates.

A conceptual model has been proposed to explain how changes in social environment imposed by the COVID-19 pandemic influence the health status of patients with chronic pain 16. In this model, the pandemic is conceptualized as a social threat. People living with chronic pain face challenges in maintaining optimal levels of social well-being even before the pandemic. Then, rules and systems to control the spread of COVID-19 such as lockdown and social distancing may worsen patients' social isolation, increased child/family care burden, interpersonal conflict, access to pain care, and perceived injustice view of society and social inequality. Cumulative evidence suggests that these social threats worsen pain levels, pain-related symptoms, psychosocial function,

Systems Neuroscience and Pain Lab, Division of Pain Medicine, Stanford University School of Medicine, 1070 Arastradero Road, Suite 200, Palo Alto, CA 94304, USA. [™]email: mziadni@stanford.edu

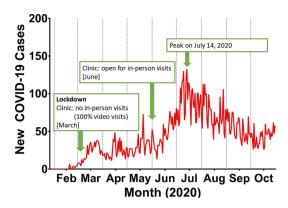


Figure 1. New COVID-19 cases in San Mateo county where the Stanford Pain Management. Center is located. Data were obtained from https://www.smchealth.org/coronavirus-health-data.

and pain interference^{17–22}. Evidence supporting this model was found in a cross-sectional online survey of community-dwelling adults with chronic pain who reported an increase in average pain intensity, pain flare-up episodes, distress, pain interference, and impaired sleep during the pandemic²³.

The aim of the current study was to evaluate the physical and psychosocial health impacts of the COVID-19 pandemic on patients seeking treatment for chronic pain. A secondary aim is to identify potential sex differences in the pandemic's health impact. Worse health status has been observed in female patients seeking treatment for chronic pain in primary care²⁴ and pain clinic²⁵ settings. Yet it is unknown whether the pandemic has a greater impact on female patients' health status. We hypothesized that pain and pain-related symptoms, psychological distress, and social function would worsen after the COVID-19 pandemic onset. Second, we hypothesized that female patients would report greater negative health impacts of the pandemic compared to male patients.

Methods

Study setting and participants. We conducted a cross-sectional study matched by season to examine patient-reported health symptoms among patients with chronic pain pre- and post-COVID-19 pandemic onset. Study procedures included a retrospective data analysis from patient surveys collected during initial visits to a tertiary pain clinic. Data was collected through CHOIR (Collaborative Health Outcomes Information Registry) to characterize patients' physical, psychological, and social function during the months of pandemic lockdown, which went into effect in March of 2020 in the California Bay Area, and compared it to a control pre-COVID-19 time period. Our clinic offered virtual visits only in the first three months of the active lockdown (March–May) and re-opened for in-person visits in June 2020. See Fig. 1 for a timeline of COVID-19 cases in San Mateo County with clinic lockdown and opening dates indicated.

The current study included a total of 7535 patients at the Stanford Pain Management Center between 2017 and 2020. The post-COVID-19 onset cohort included 1798 patients who completed initial surveys collected between April–October 2020, and the control pre-COVID-19 cohort included 5737 patients who completed initial surveys collected between April–October 2017, 2018, and 2019. To control for seasonal effects, we constrained our control cohort to the same months as the post-COVID-19 onset cohort for the preceding 3 years. Stanford University's Institutional Review Board (IRB) approved the study procedures. Informed consent was waived by the IRB, as CHOIR data were collected for clinical care and quality improvement purposes. The research was conducted in full compliance with the APA Ethical Principles in the treatment of human participants.

Measures. Demographic. Patient sex, race/ethnicity, disability status, employment status, marital status, and age were collected to assess demographic characteristics. We also included in analyses the number of doctor's office and ER visits over the past six months and the number of painful sites using a body map²⁶. Finally, we analyzed current drinking status and number of drinks per week for those endorsing alcohol use.

Pain intensity. Pain intensity was assessed on a numerical rating scale (NRS) using the PROMIS Pain Intensity scale²⁷. Respondents were asked to rate their average pain intensity over the previous 7 days on a scale of 0–10. Assessment of pain intensity using an NRS has been supported in prior studies²⁸.

Pain catastrophizing. The Pain Catastrophizing Scale (PCS) is a 13-item questionnaire that assesses distress regarding the cognition and emotion associated with actual or anticipated pain²⁹. The PCS comprises three different subscales: rumination, magnification, and helplessness. Patients rate each item on a 5-point Likert Scale (0 = Not at all to 4 = All the time). The PCS has a minimum score of 0 and a maximum score of 52 with higher scores indicative of more catastrophic thinking.

The patient-reported outcomes measurement information system (PROMIS). The PROMIS measures are well-validated and widely used to assess physical and psychosocial health status in patients with chronic illnesses, including chronic pain^{30–35}. Detailed information about the measure development and validation is available

at http://www.healthmeasures.net. All PROMIS measures in the current study were administered using computerized adaptive testing (CAT). T-scores were calculated for each patient (M=50, SD=10). PROMIS CAT instruments were administered for pain interference, physical function, fatigue, sleep impairment, depression, anxiety, anger, social isolation, emotional support, and satisfaction with roles and activities. Higher scores on each PROMIS measure generally indicate greater severity of each symptom domain. However, higher scores on PROMIS physical function, emotional support, and satisfaction with roles and activities indicate better health status in each domain.

Statistical analysis. Data were checked for completeness and nonsensical input (e.g., negative ages) before each demographic variable was quantified. We analyzed data of people who completed all PROMIS measures and the PCS. Subsequently, missing values existed only in demographic data. The proportion of patients belonging to each demographic variable were compared between pre- and post-COVID-19 pandemic onset cohorts using chi-squared tests to detect significant changes in cohort composition across timepoints. Additionally, age and the number of office visits, ER visits, painful sites, and weekly drinks were compared between the pre- and post-COVID-19 cohorts using a two-samples t-test.

SPSS 26 was used to conduct 2 (sex: female and male patients) by 2 (time: pre- and post-COVID-19 pandemic onset) by 7 (month: April to October) weighted-means MANOVAs for three sets of variables: (a) pain rating and PCS scores, (b) PROMIS physical health measures, and (c) PROMIS psychosocial health measures. Due to unbalanced samples, with small sample sizes for the post-COVID-19 cohorts, Type 1 (sequential) Sum of Squares was used for MANOVA and post-hoc univariate analyses³⁶. To reduce Type I error for multiple comparisons, *p* values from post-hoc analyses were false discovery rate (FDR) adjusted in R using the Benjamini and Hochberg (BH) procedure³⁷. Addressing our primary research aim, which was to investigate impacts of the COVID-19 pandemic on patients' health status, we examined the main effect of time and its interaction with month predicting patient health status. To address our secondary research aim, which was to examine sex differences in the effect of the pandemic, we focused on the main effect of time and its interaction with sex and/or month.

Results

Sample characteristics. Characteristics of both cohorts are presented in Table 1. The majority of the patients across post-COVID-19 pandemic onset and pre-COVID-19 cohorts were female (67.3 and 71.0%), White/Caucasian (58.4 and 60.2%), and middle-aged [M=51.4 (17.0) and M=50.2 (17.5)], respectively. Most patients were not on disability across both cohorts (78.8% and 79.9%), and over half of patients reported that they were not currently working (55.7% and 56.8%). In sum, no significant differences were noted across cohorts on demographic variables except for sex and age, with the post-COVID-19 cohort including slightly more female patients and slightly younger in age.

The proportion of patients drinking alcohol was similar between the two cohorts (41.6% and 40.2%, p = 0.544). Among patients drinking alcohol, the number of drinks per week were also not significantly different between the pre- and post-COVID-19 cohorts (3.7 and 3.5 drinks, p = 0.358). While no significant differences in healthcare utilization were found (doctor's office and ER visits, ps > 0.068), the post-COVID-19 cohort endorsed significantly more painful sites on the body map (M = 12.3) than the pre-COVID-19 cohort (M = 11.5, p = 0.030, Table 1).

Comparing pain ratings and pain catastrophizing scores between pre- and post-COVID-19 onset cohorts. Two (sex: male and female) by two (time: pre- and post-COVID-19 pandemic onset) by seven (month: April to October) weighted-means MANOVAs were conducted for pain ratings and pain catastrophizing scores (Table 2). There was a significant main effect of sex (p<0.001) and a significant interaction between time and month (p=0.020). Post-hoc univariate analyses showed that the time-by-month interaction was significant for PCS scores [F(6, 7,507)=2.88, p=0.016] but not for pain ratings (p=0.143). Specifically, PCS scores significantly differed between cohorts in June [F(1, 938)=7.09, p=0.028] and July [F(1, 998)=7.98, p=0.028], but did not differ in the other months (ps>0.478). In June, PCS scores were significantly lower in the post-COVID-19 cohort (M=19.1, SD=12.6) compared to the pre-COVID-19 cohort (M=20.0, SD=12.8, η ²=0.007; Fig. 2A). In July, the opposite pattern was observed: PCS scores were significantly higher in the post-COVD-19 cohort (M=22.1, SD=12.6) compared to the pre-COVID cohort (M=19.5, SD=12.3, η ²=0.008; Fig. 2A).

Taken together, we found small but significant differences in pain catastrophizing in June and July 2020 compared to pre-pandemic months. PCS scores were significantly lower after the onset of the pandemic in June 2020 when the clinic was re-opened for in-person visits, and increased in July 2020 when new COVID-19 cases peaked. However, these changes in PCS scores in June and July 2020 were not paralleled by changes in pain ratings (Fig. 2B).

Comparing PROMIS-physical health measures between pre- and post-COVID-19 onset cohorts. Two (sex: male and female) by two (time: pre- and post-COVID-19 pandemic onset) by seven (month: April to October) weighted-means MANOVAs were conducted for PROMIS-pain interference, physical function, fatigue, and sleep impairment T-scores (Table 2). There were significant main effects of sex (p < 0.001) and time (p < 0.001), as well as a significant time-by-sex interaction (p = 0.027). Post-hoc univariate analyses revealed significant effects of time for the pain interference, physical function, fatigue, and sleep impairment scores, F s> 5.10, p s < 0.036, η^2 = 0.001, suggesting a slight improvement in these physical health measures (Fig. 3A–D). Post-hoc univariate analyses revealed a significant sex-by-time interaction only for fatigue, F (1, 7659) = 5.84, p = 0.027, η^2 = 0.001, but not for the other measures (p s > 0.338). PROMIS-fatigue T-scores were

	Pre- COVID-19 pandemic N=5737		Post- COVID-19 pandemic onset N=1798			
Characteristic	n	(%)	n	(%)	X^2	p
Gender					8.57	0.003
Female	3860	(67.3)	1276	(71.0)		
Male	1877	(32.7)	522	(29.0)		
Race/ethnicity					1.87	0.171
White	3348	(58.4)	1082	(60.2)		
Others [^]	2389	(41.6)	716	(39.8)		
(Asian)	569	(9.9)	187	(10.4)		
(Hispanic/Latino)	448	(7.8)	155	(8.6)		
(Black or African American)	278	(4.8)	55	(3.1)		
(bi-racial, Native Indians, etc.)	824	(14.4)	229	(12.7)		
(Missing/refused to disclose)	270	(4.7)	90	(5.0)		
Disability status					1.03	0.309
Not on disability	4521	(78.8)	1437	(79.9)		
On disability	1216	(21.2)	361	(20.1)		
Employment status					0.73	0.391
Currently working	2542	(44.3)	776	(43.2)		
Not currently working	3195	(55.7)	1022	(56.8)		
Marital status					2.77	0.598
Married	3137	(55.4)	972	(54.7)		
Never married	1144	(20.2)	380	(21.4)		
Divorced/separated	758	(13.4)	223	(12.6)		
Living together	345	(6.1)	119	(6.7)		
Widowed	277	(4.9)	82	(4.6)		
(missing) [^]	76	-	22	-		
Drinking alcohol					0.37	0.544
Yes	1864	(41.6)	648	(40.2)		
No	2621	(58.4)	963	(59.8)		
(missing) [^]	1252	-	187	-		
	М	(SD)	М	(SD)	t	p ^
Age	51.4	(17.0)	50.2	(17.5)	2.96	0.015
Doctor's office visits	7.0	(8.8)	6.6	(8.3)	2.04	0.068
ER visits	0.8	(2.4)	0.8	(2.5)	0.37	0.714
Painful sites (body map)	11.5	(12.3)	12.3	(12.3)	- 2.50	0.030
Number of drinks per week*	3.7	(4.5)	3.6	(6.8)	0.57	0.563

significantly lower in the post-COVID-19 cohort versus pre-COVID-19 cohort among female patients, F(1, 5134) = 16.66, p < 0.001, but not among male patients (p = 0.838, Supplementary Table).

Contrary to our primary hypothesis, we found small but significant improvements in pain interference, physical function, and sleep impairment in the post-COVID-19 cohort. Addressing our secondary hypothesis regarding sex differences, we found that female patients in the post-COVID-19 cohort showed significant improvements in fatigue symptoms (Fig. 3D).

Comparing PROMIS-psychosocial health measures between pre- and post-COVID-19 onset cohorts. Two (sex: male and female) by two (time: pre- and post-COVID-19 pandemic onset) by seven (month: April to October) weighted-means MANOVAs were conducted for PROMIS-depression, anxiety, anger, emotional support, social isolation, and satisfaction with social roles and activities (Table 2). There was a significant main effect of sex (p < 0.001) and time (p < 0.001). Post-hoc univariate analyses showed that the main effect of time was significant for emotional support F(1, 7,553) = 15.01, p < 0.001, $\eta^2 = 0.002$, but not for the other measures (ps > 0.084). PROMIS-emotional support T-scores were significantly higher in the post-COVID-19 cohort (M = 52.2, SD = 9.5) than the pre-COVID-19 cohort (M = 51.2, SD = 9.5, Fig. 4).

Measures		Λ	F	df	p
Average pain intensity PCS scores	Sex	0.994	22.62	(2, 7506)	< 0.000
	Time	1.000	0.07	(2, 7506)	0.932
	Month	0.998	0.95	(12, 15,012)	0.455
	Time × month	0.997	2.01	(12, 15,012)	0.020
	Sex × time	0.999	2.34	(2, 7506)	0.096
	Sex × month	0.999	0.70	(12, 15,012)	0.752
	$Sex \times time \times month$	0.999	0.79	(12, 15,012)	0.660
Pain interference Physical function Fatigue Sleep impairment	Sex	0.984	30.26	(4, 7504)	< 0.001
	Time	0.998	4.41	(4, 7504)	0.001
	Month	0.996	1.24	(24, 26,180)	0.196
	Time × month	0.996	1.19	(24, 26,180)	0.240
	Sex × time	0.999	2.74	(4, 7504)	0.027
	Sex × month	0.997	1.01	(24, 26,180)	0.443
	$Sex \times time \times month$	0.998	0.67	(24, 26,180)	0.887
Depression Anxiety Anger Social isolation Emotional support Satisfaction with social roles and activities	Sex	0.983	22.10	(6, 7502)	< 0.001
	Time	0.996	5.12	(6, 7502)	< 0.001
	Month	0.995	1.04	(36, 32,946)	0.396
	Time × month	0.997	0.69	(36, 32,946)	0.917
	Sex × time	0.999	0.71	(6, 7502)	0.643
	Sex × month	0.994	1.25	(36, 32,946)	0.917
	$Sex \times time \times month$	0.997	0.71	(36, 32,946)	0.906

Table 2. Three-way MANOVA for pain ratings, PCS scores, and PROMIS measures. Bold font indicates significance at p < 0.05.

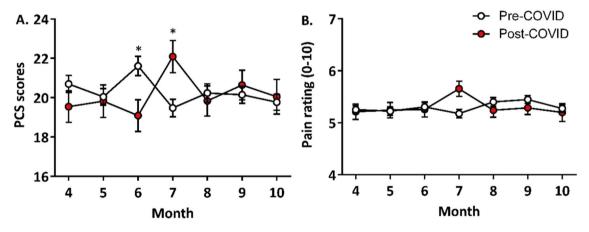


Figure 2. Comparing the changes in PCS scores (**A**) and pain ratings (**B**) by month between the pre- and post-COVID-19 cohorts.

These results suggest that patients in the post-COVID-19 cohort reported small but significant improvements in emotional support compared to the pre-COVID-19 cohort. Contrary to our hypothesis, we did not see a significant worsening of depression, anxiety, anger, social isolation, or satisfaction with social roles and activities in the post-COVID-19 patient cohort.

Discussion

We conducted a cross-sectional study matched by season to examine patient-reported health symptoms in patients with chronic pain pre- and post-COVID-19 pandemic onset (April-October 2020). To our knowledge, this is the first study to investigate the impacts of the COVID-19 pandemic and sex on the health status of patients with chronic pain seeking treatment.

Drastic changes in social interactions and medical practice imposed by the COVID-19 pandemic have created unique challenges to chronic pain management. Already, there is emerging evidence in the literature of the negative impacts of the pandemic on patients with chronic pain. Community-dwelling adults with chronic pain in one study reported worse pain and health status compared to their recall of pre-pandemic health status²³, and pain physicians have reported observations of increased pain levels in patients visiting their clinics after the

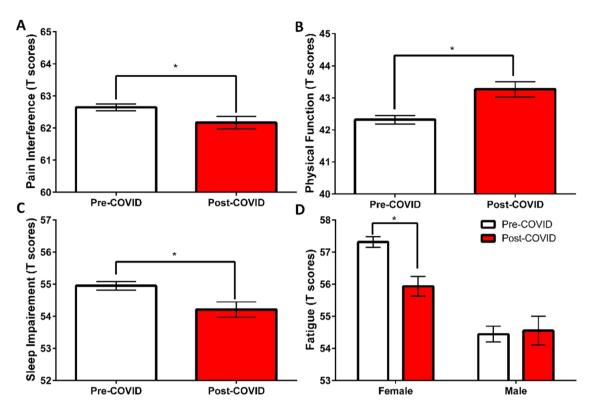


Figure 3. Comparing PROMIS-pain interference (**A**), physical function (**B**), sleep impairment (**C**), and fatigue (**D**) T-scores between pre- and post-COVID-19 pandemic onset cohorts.

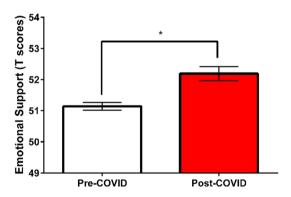


Figure 4. Comparing the PROMIS-emotional support T-scores between the pre- and post-COVID-19 cohorts.

pandemic onset compared to their recall of typical patients prior to the pandemic ³⁸. Given these findings, in the present study we hypothesized that the onset of the COVID-19 pandemic would be associated with worse pain and physical and psychosocial health among patients with chronic pain visiting our clinic. Consistent with our hypothesis, we found that patients' pain catastrophizing was significantly lower after the onset of the pandemic in June 2020 (when the clinic was re-opened for in-person visits), and higher in July 2020 (when new COVID-19 cases peaked in Santa Clara and San Mateo County) compared to pre-pandemic months. Counter to our hypothesis, however, we did not observe a similar change in pain ratings, and in fact observed small but significant improvements in patients' pain interference, physical function, and sleep impairment in the post-COVID-19 sample. These findings were not consistent with those reported among individuals living with chronic pain in two Pan-Canadian studies^{13,14}. Similarly, addressing our secondary research aim of investigating sex differences in the impact of the pandemic, we found that female patients in the post-COVID-19 cohort showed significant improvements in fatigue symptoms.

Overall, we found that the pandemic did not adversely influence markers of physical and mental health among patients seeking treatment for pain in our clinic. Our post-COVID-19 cohort also reported small but significant increases in emotional support and no significant changes in social isolation and satisfaction with social roles. Although this is contrary to published findings to date^{13,14}, an important distinction is that the current study sample were in treatment at the Stanford Pain Center and continued to receive virtual treatment during

the pandemic. Treatment was continuously available via virtual-platforms throughout the study timeframe, and engagement was very high. In addition, patients reported access to increases in emotional support in the post COVID group. These factors likely buffered the impacts of COVID-19 and were key contributors to the observed improvements. In addition, study findings are consistent with prior findings of resilience in response to the pandemic²⁻⁴, which may have buffered the effects of the pandemic on stress and emotional functioning. Furthermore, small but significant changes in pain interference, sleep impairment, and physical function were all in the direction of improvement after the pandemic. These results should also be considered in the context of our patients' social situations. Most of our patients were married or living together, of which proportions were not significantly different between the pre- (68.8%) and post-COVID-19 cohorts (69.3%). Additionally, employment status, disability status, and the numbers of doctor's office visits and ER visits for the past six months were not significantly different between the pre- and post-pandemic cohorts.

Past research has suggested worsening outcomes following epidemics and past natural disasters^{39–41}. Alcohol use, PTSD, anxiety, anger, fear of contagion, perceived risk, and distrust emerge as potential effects from the COVID-19 pandemic⁴⁰. As such, future work should focus on identifying risk and protective factors that may mitigate or exacerbate negative impacts, including mental health status, substance use, and related demographic variables. Importantly, current study findings support the notion that having access to treatment in the context of a pandemic was likely protective and allowed patients to benefit from the care in spite of the pandemic. Additionally, research on resilience among individuals with chronic pain is growing⁴², and studies show that certain positive characteristics, such as optimism, pain acceptance, and purpose in life have been associated with increased resilience to chronic pain⁴³. It is possible that the development of coping strategies required to manage chronic pain may be protective when facing a widespread disruptive event such as a pandemic.

However, it is possible that it is still too early to assess the full impacts of the pandemic on limiting access to care, medications, or other critical services. With a pattern of cyclic surges of COVID-19 cases and new strains emerging, patients with chronic pain will continue for the foreseeable future to experience social and emotional challenges arising from the pandemic. In our clinic, when new COVID-19 cases peaked in July 2020, our patients' pain catastrophizing scores increased compared to the same month of the previous three years. The extended social and emotional toll of the pandemic, with multiple surges of new COVID-19 cases and associated lockdowns, may be increasing patients' catastrophic thinking about their pain. Pain catastrophizing is a negative cognitive and emotional response pattern to actual or anticipated pain 44,45 that has been previously linked to increased pain report, opioid use, and pain-related disability⁴⁶⁻⁴⁸. More study is warranted to understand the longterm physical and mental health impacts of the pandemic. Alternatively, the Substance Abuse and Mental Health Services Administration (SAMHSA) acknowledges that lack of health insurance and limited access to mental and behavioral health services are a major health disparity issue during the COVID-19 pandemic⁴⁹. Yet, this is not the case for the majority of the patients seeking treatment at our pain clinic. Future study in more vulnerable patient populations, who may lack health insurance or have substantial barriers to care, is needed. Even though the confounding impact of alcohol use was examined in this study, future work should assess the confounding effect of other and recreational substance use, in addition to mental health status, on patient-outcomes.

Several limitations of the current study are worth noting. First, the cross-sectional design of our study precludes making causal inferences. The negative effects of the pandemic may increase over time with persistent virus spread, worsening economic hardship, social isolation, and inability to obtain medications. Hence, future longitudinal studies are required to assess changes in pain and functioning over time. Next, selection bias is another potential limitation, as it is possible that the patients most severely impacted by the COVID-19 pandemic were not sampled in our study because they were not able to successfully obtain care at our clinic. Finally, our study cohorts were recruited from a pain management clinic located in the San Francisco Bay Area. Regional variability in COVID-19 caseloads, mortality, and government-mandated lockdowns may influence the negative impacts of the pandemic on patients with pain. In San Mateo County, where the Stanford Pain Management Center is located, there have been considerably fewer deaths due to COVID-19 than in many other parts of the United States⁵⁰. Increased severity of an outbreak in a locality may result in increased detrimental physical and psychosocial impacts in patients with pain and other chronic illnesses. Notably, the current study was not designed to specifically query COVID-19 impacts. Future research is needed to probe patient experiences specific to the COVID-19 pandemic, such as testing positive for COVID-19, experiencing long-term effects of infection, loss of employment and related financial impacts, barriers to accessing healthcare, and childcare burden, and whether these new stressors correlate with pain-related outcomes.

Study findings are encouraging and could be bolstered by considering adjunctive digital health solutions to further address COVID-19-related misinformation, provide support, and maintain continuity of care⁵¹. For instance, we showed that a zoom-delivered behavioral health intervention "Empowered Relief" imparts enduring benefits⁵²; this and other online-delivered interventions, as well as on-demand digital treatments such as virtual reality⁵³, may be integrated into pain care pathways broadly. In light of the urgent clinical need and patient preference for digital services due to the COVID-19 pandemic, future studies could focus on assessing the impact of online-health communities, using phone calls, and AI-based approaches for providing counseling and promoting self-management skills among patients with chronic pain⁵¹.

Conclusion

Despite these limitations, this is the first study to report data on patients with chronic pain, controlling for seasonal effects, and examining sex differences in potential adverse health impacts of the pandemic. A notable strength of this study is the multidimensional nature of the data, which included both psychosocial and pain-related measures. Finally, we used a learning health system to examine changes in physical and psychosocial health among a large number of real-world patients seeking treatment for pain before and after the pandemic

onset. Subsequently, our results were free of recall bias and ecologically valid in understanding the health status of patients with whom treating physicians would interact in their daily practice. In conclusion, our findings overall provide evidence of encouraging resilience among patients seeking treatment for pain conditions in the face of the COVID-19 pandemic. However, our finding that patients' pain catastrophizing increased when COVID-19 cases peaked points to the need for future study and monitoring of patients as the effects of the pandemic continue.

Data availability

The datasets generated during and/or analyzed during the current study are available from the corresponding author on reasonable request.

Received: 20 April 2021; Accepted: 6 April 2022

Published online: 19 April 2022

References

- 1. Eccleston, C. et al. Managing patients with chronic pain during the COVID-19 outbreak: Considerations for the rapid introduction of remotely supported (eHealth) pain management services. Pain 161, 889 (2020).
- 2. Wang, C. et al. A longitudinal study on the mental health of general population during the COVID-19 epidemic in China. Brain Behav. Immun. 87, 40–48 (2020).
- 3. Luchetti, M. et al. The trajectory of loneliness in response to COVID-19. Am. Psychol. 75, 897-908 (2020).
- 4. Hamm, M. E. et al. Experiences of American older adults with pre-existing depression during the beginnings of the COVID-19 pandemic: A multicity, mixed-methods study. Am. J. Geriatr. Psychiatry 28, 924–932 (2020).
- Yamada, M. et al. Effect of the COVID-19 Epidemic on physical activity in community-dwelling older adults in Japan: A crosssectional online survey. J. Nutr. Health Aging 24, 1–3 (2020).
- Twenge, J. M. & Joiner, T. E. US Census Bureau-assessed prevalence of anxiety and depressive symptoms in 2019 and during the 2020 COVID-19 pandemic. *Depress. Anxiety* 37, 954–956 (2020).
- Ettman, C. K. et al. Prevalence of depression symptoms in US adults before and during the COVID-19 pandemic. JAMA Netw. Open 3, e2019686–e2019686 (2020).
- 8. Pollard, M. S., Tucker, J. S. & Green, H. D. Changes in adult alcohol use and consequences during the COVID-19 pandemic in the US. JAMA Netw. Open 3, e2022942–e2022942 (2020).
- Ciążyńska, M. et al. Quality of life of cancer patients during coronavirus disease (COVID-19) pandemic. Psycho-oncology. https://doi.org/10.1002/pon.5434 (2020).
- 10. Crettaz, B. et al. Stress-induced allodynia-evidence of increased pain sensitivity in healthy humans and patients with chronic pain after experimentally induced psychosocial stress. PLoS One 8, e69460 (2013).
- Generaal, E. et al. Biological stress systems, adverse life events and the onset of chronic multisite musculoskeletal pain: A 6-year cohort study. Ann. Rheum. Dis. 75, 847–854 (2016).
- Clauw, D. J., Häuser, W., Cohen, S. P. & Fitzcharles, M.-A. Considering the potential for an increase in chronic pain after the COVID-19 pandemic. *Pain* 161, 1694–1697 (2020).
- Pagé, M. G. et al. A cross-sectional study of pain status and psychological distress among individuals living with chronic pain: the Chronic Pain & COVID-19 Pan-Canadian Study. Health Promot. Chronic Dis. Prev. Can. Res. Policy Pract. 41, 141–152 (2021).
- 14. Lacasse, A. *et al.* Impact of the COVID-19 pandemic on the pharmacological, physical, and psychological treatments of pain: Findings from the Chronic Pain & COVID-19 Pan-Canadian Study. *Pain Rep.* **6**, e891 (2021).
- Hruschak, V. et al. Cross-sectional study of psychosocial and pain-related variables among patients with chronic pain during a time of social distancing imposed by the coronavirus disease 2019 pandemic. Pain 162, 619 (2021).
- 16. Karos, K. et al. The social threats of COVID-19 for people with chronic pain. Pain 161, 2229-2235 (2020).
- 17. Karayannis, N. V., Baumann, I., Sturgeon, J. A., Melloh, M. & Mackey, S. C. The impact of social isolation on pain interference: A longitudinal study. *Ann. Behav. Med.* **53**, 65–74 (2019).
- 18. Ziadni, M. S., You, D. S., Sturgeon, J. A., Mackey, S. C. & Darnall, B. D. Perceived injustice mediates the relationship between perceived childhood neglect and current function in patients with chronic pain: A preliminary pilot study. *J. Clin. Psychol. Med. Settings* 28, 1–12 (2020).
- 19. Wolf, L. D. & Davis, M. C. Loneliness, daily pain, and perceptions of interpersonal events in adults with fibromyalgia. *Health Psychol.* 33, 929 (2014).
- Sturgeon, J. A. et al. Social disruption mediates the relationship between perceived injustice and anger in chronic pain: A collaborative health outcomes information registry study. Ann. Behav. Med. 50, 802–812 (2016).
- Sturgeon, J. A., Dixon, E. A., Darnall, B. D. & Mackey, S. C. Contributions of physical function and satisfaction with social roles to emotional distress in chronic pain: A Collaborative Health Outcomes Information Registry (CHOIR) study. *Pain* 156, 2627 (2015).
- 22. Parrish, B. P., Zautra, A. J. & Davis, M. C. The role of positive and negative interpersonal events on daily fatigue in women with fibromyalgia, rheumatoid arthritis, and osteoarthritis. *Health Psychol.* 27, 694 (2008).
- 23. Nieto, R., Pardo, R., Sora, B., Feliu-Soler, A. & Luciano, J. V. Impact of COVID-19 lockdown measures on Spanish people with chronic pain: An online study survey. *J. Clin. Med.* **9**, 3558 (2020).
- 24. Stubbs, D. et al. Sex differences in pain and pain-related disability among primary care patients with chronic musculoskeletal pain. Pain Med. 11, 232–239 (2010).
- Lamé, I. E., Peters, M. L., Vlaeyen, J. W., Kleef, M. V. & Patijn, J. Quality of life in chronic pain is more associated with beliefs about pain, than with pain intensity. Eur. J. Pain 9, 15–24 (2005).
- 26. Scherrer, K. H. et al. Development and validation of the Collaborative Health Outcomes Information Registry body map. Pain reports 6, e880 (2021).
- 27. Cook, K. F. et al. Pain assessment using the NIH Toolbox. Neurology **80**, S49–S53 (2013).
- 28. Farrar, J. T., Young, J. P. Jr., LaMoreaux, L., Werth, J. L. & Poole, R. M. Clinical importance of changes in chronic pain intensity measured on an 11-point numerical pain rating scale. *Pain* 94, 149–158 (2001).
- 29. Sullivan, M. J., Bishop, S. R. & Pivik, J. The pain catastrophizing scale: Development and validation. Psychol. Assess 7, 524 (1995).
- 30. Askew, R. L., Cook, K. F., Revicki, D. A., Cella, D. & Amtmann, D. Evidence from diverse clinical populations supported clinical validity of PROMIS pain interference and pain behavior. *J. Clin. Epidemiol* 73, 103–111 (2016).
- 31. Cella, D. et al. The Patient-Reported Outcomes Measurement Information System (PROMIS) developed and tested its first wave of adult self-reported health outcome item banks: 2005–2008. J. Clin. Epidemiol. 63, 1179–1194 (2010).
- 32. Cella, D. et al. PROMIS fatigue item bank had clinical validity across diverse chronic conditions. J. Clin. Epidemiol. 73, 128–134 (2016).
- 33. Cook, K. F. et al. PROMIS measures of pain, fatigue, negative affect, physical function, and social function demonstrated clinical validity across a range of chronic conditions. J. Clin. Epidemiol. 73, 89–102 (2016).

- 34. Schalet, B. D. et al. Validity of PROMIS physical function measured in diverse clinical samples. J. Clin. Epidemiol. 73, 112–118 (2016).
- 35. Buysse, D. J. et al. Development and validation of patient-reported outcome measures for sleep disturbance and sleep-related impairments. Sleep 33, 781–792 (2010).
- 36. Tabachnick, B. G. & Fidell, L. S. Experimental Designs Using ANOVA. (Thomson/Brooks/Cole Belmont, 2007).
- 37. Thissen, D., Steinberg, L. & Kuang, D. Quick and easy implementation of the Benjamini–Hochberg procedure for controlling the false positive rate in multiple comparisons. *J. Educ. Behav. Stat.* 27, 77–83 (2002).
- 38. Lynch, M. E., Williamson, O. D. & Banfield, J. C. COVID-19 impact and response by Canadian pain clinics: A national survey of adult pain clinics. *Can. J. Pain* 4, 204–209 (2020).
- Lee, A. M. et al. Stress and psychological distress among SARS survivors 1 year after the outbreak. Can. J. Psychiatry 52, 233–240 (2007).
- 40. Esterwood, E. & Saeed, S. A. Past epidemics, natural disasters, COVID19, and mental health: Learning from history as we deal with the present and prepare for the future. *Psychiatr. Q.* **91**, 1–13 (2020).
- 41. La Greca, A. M., Silverman, W. K., Vernberg, E. M. & Prinstein, M. J. Symptoms of posttraumatic stress in children after Hurricane Andrew: A prospective study. J. Consult. Clin. Psychol. 64, 712 (1996).
- 42. Sturgeon, J. A. & Zautra, A. J. Resilience: A new paradigm for adaptation to chronic pain. Curr. Pain Headache Rep. 14, 105–112 (2010).
- Smith, B. W. & Zautra, A. J. Vulnerability and resilience in women with arthritis: Test of a two-factor model. J. Consult. Clin. Psychol. 76, 799 (2008).
- 44. Rosenstiel, A. K. & Keefe, F. J. The use of coping strategies in chronic low back pain patients: Relationship to patient characteristics and current adjustment. *Pain* 17, 33–44 (1983).
- 45. Sullivan, M. J. et al. Theoretical perspectives on the relation between catastrophizing and pain. Clin. J. Pain 17, 52-64 (2001).
- 46. Schütze, R. et al. How can we best reduce pain catastrophizing in adults with chronic noncancer pain? A systematic review and meta-analysis. J. Pain 19, 233–256 (2018).
- 47. Lazaridou, A. et al. Influence of catastrophizing on pain intensity, disability, side effects, and opioid misuse among pain patients in primary care. J. Appl. Biobehav. Res. 22, e12081 (2017).
- 48. Picavet, H. S. J., Vlaeyen, J. W. & Schouten, J. S. Pain catastrophizing and kinesiophobia: Predictors of chronic low back pain. *Am. J. Epidemiol.* **156**, 1028–1034 (2002).
- Jeopardy, D. COVID-19 and Behavioral Health Disparities for Black and Latino Communities in the US Substance Abuse and Mental Health Services Administration. https://www.samhsa.gov/sites/default/files/covid19-behavioral-health-disparities-black-latino-communities.pdf (accessed 07 Aug 2020) (2020).
- 50. JHU. Coronavirus Resource Center: California State Overview. https://coronavirus.jhu.edu/region/us/california (2021).
- 51. Li, L. W., Chew, A. M. & Gunasekeran, D. V. Digital health for patients with chronic pain during the COVID-19 pandemic. *BJA Br. J. Anaesth.* **125**, 657 (2020).
- 52. Ziadni, M. S., Gonzalez-Castro, L., Anderson, S., Krishnamurthy, P. & Darnall, B. D. Efficacy of a single-session "empowered relief" zoom-delivered group intervention for chronic pain: Randomized controlled trial conducted during the COVID-19 pandemic. *J. Med. Internet. Res.* 23, e29672 (2021).
- 53. Garcia, L. M. et al. An 8-week self-administered at-home behavioral skills-based virtual reality program for chronic low back pain: Double-blind, randomized, placebo-controlled trial conducted during COVID-19. J. Med. Internet. Res. 23, e26292 (2021).

Acknowledgements

We acknowledge funding from the National Institutes of Health (NIH) National Institute on Drug Abuse (NIDA) which supported this work: K23DA047473 (MSZ), K23DA048972 (DSY), K24DA029262 and K24NS126781 (SCM). All authors report no conflicts of interest.

Author contributions

M.S.Z. was the lead author on the conception and design of the study, acquisition of the data, and drafting of the manuscript. D.S.Y. has made significant contribution to the analysis and interpretation of the data and drafting the manuscript. E.C. has contributed to the data analysis and drafting the manuscript. S.R.A. and B.H. have contributed to drafting the manuscript. S.C.M. and B.D.D. have contributed to substantially revising the manuscript. G.H. has contributed to drafting of the manuscript. All authors have approved the enclosed version and agreed both to be personally accountable for the author's own contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature.

Competing interests

Dr. Darnall discloses that she receives royalties for four pain treatment books she has authored. She is the principal investigator for pain research grants and awards from the National Institutes of Health (NIH) and the Patient-Centered Research Outcomes Research Institute (none specific to the current work). Dr. Darnall is chief science advisor at AppliedVR. She serves on the Board of Directors for the American Academy of Pain Medicine and is on the Board of Directors for the Institute for Brain Potential. She is a scientific member of the NIH Interagency Pain Research Coordinating Committee, a member of the Centers for Disease Control and Prevention (CDC) Opioid Workgroup (2020–2021) and is a member of the Pain Advisory Group of the American Psychological Association. Dr. Mackey receives research funding from the NIH, FDA, and PCORI (administered through Stanford University). He is an unpaid advisor to the following organizations: American Chronic Pain Association (Scientific Advisory Board); Analgesic, Anesthetic, and Addiction Clinical Trial Translations, Innovations, Opportunities, and Networks (ACTTION), a public–private partnership with the FDA (Oversight committee); National Pain Advocacy Center (Science and Policy Advisory Council). There are no additional competing interests to declare.

Additional information

Supplementary Information The online version contains supplementary material available at https://doi.org/10.1038/s41598-022-10431-5.

Correspondence and requests for materials should be addressed to M.S.Z.

Reprints and permissions information is available at www.nature.com/reprints.

Publisher's note Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Open Access This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/.

© The Author(s) 2022