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Executive and attentional functions in chronic pain: Does performance decrease with increasing task load?

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BACKGROUND: Diminished executive function and attentional control has been reported in chronic pain patients. However, the precise pattern of impairment in these aspects of cognition in chronic pain remains unclear. Moreover, a decline in psychomotor speed could potentially influence executive and attentional control performance in pain patients.

OBJECTIVE: To examine different aspects of executive and attentional control in chronic pain together with the confounding role of psychomotor slowing.

METHODS: Neuropsychological tests of sustained attention, planning ability, inhibition and mental flexibility were administered to 34 participants with chronic pain and 32 control participants.

RESULTS: Compared with the controls, participants with chronic pain took longer to complete tests of sustained attention and mental flexibility, but did not perform worse on inhibition or planning tasks. The decreased performance on the mental flexibility task likely reflects a reduction in psychomotor speed. The pattern of performance on the sustained attention task reveals a specific decline in attention, indicated by a disproportionate decline in performance with an increase in task duration and by increased fluctuations in attention during task performance. No additional effect was noted of pain intensity, pain duration, pain catastrophizing, depressive symptoms, reduced sleep because of the pain or opioid use.

CONCLUSIONS: Executive and attention functions are not uniformly affected in chronic pain. At least part of the previously reported decline in executive function in this group may reflect psychomotor slowing. Overall, limited evidence was found that executive and attention performance is indeed lower in chronic pain. Therefore, it can be concluded that in chronic pain sustained attention performance is diminished while mental flexibility, planning and inhibition appear to be intact.

Key Words: Attention; Chronic pain; Executive function; Pain catastrophizing; Pain intensity; Psychomotor speed

The influence of pain on neuropsychological test performance has been widely acknowledged (1,2). Many studies have reported that executive and attentional functions are affected in chronic pain patients, presumably as a result of competing attentional demands of chronic pain (2-5). Due to this interruptive nature of pain, it is presumed that less cognitive resources are available for other processes, including cognitive function. Because multiple functions are commonly attributed to the concepts of executive and attentional control (6), it is one of the most heterogeneous and least well understood cognitive domains to date. Furthermore, it is difficult to segregate concepts of attention and executive functions at a conceptual level. For example, attention encompasses multiple functions including sustained, divided and selective (or focused) attention (6,7), but a strong overlap exists between these attentional functions and those that are commonly regarded as executive control. This is nicely illustrated by tests such as the Trail Making Test (TMT) and the Stroop Colour Word task. The TMT part B (8) requires participants to switch between different stimuli and to keep track of previously encountered items. Hence, this test requires divided attention for successful performance but is commonly regarded as a test of mental flexibility (9). The Stroop Colour Word test (10) requires the inhibition of one type of stimulus while responding to another stimulus. This test is, therefore, commonly regarded as a measure of cognitive inhibition, but it also requires the ability to focus attention on one aspect of the task while suppressing another aspect, ie, selective attention (6). For the sake of clarity, when we refer to a task in the present article, we will...
describe it in terms of the cognitive functions that are most commonly used in the research literature, such as inhibition for the Stroop task and mental flexibility for the TMT.

The studies to date reveal conflicting findings regarding the precise effect of chronic pain on these specific executive and attention functions. Some studies failed to find decreased performance of chronic pain patients compared with controls on tasks of inhibition (11,12), mental flexibility (12-14) and letter fluency (12). In contrast, other studies reported on significant associations between chronic pain and inhibition (4), mental flexibility (15,16) and sustained attention (5,17). The precise pattern of performance on tasks of executive and attention tasks in chronic pain patients thus remains unclear.

A crucial factor that should be considered when examining executive and attentional task performance in chronic pain patients is whether this reflects a specific deficit or a decline in more basic cognitive abilities. More specifically, executive function and attention performance may be particularly vulnerable to a reduction in psychomotor speed (6), with psychomotor slowing being a consistent finding in chronic pain patients (18). This is an important consideration because frequently used tests to assess executive function and attention, such as the Stroop Colour Word test and the TMT, are strongly dependent on psychomotor speed performance (19-22). A decline in executive or attentional performance can therefore only be revealed when an increase in task demands substantially affects performance (3), with an effect that extends beyond a potential reduction in basic cognitive abilities such as psychomotor speed. Some studies attempted to adjust for reductions in these abilities by calculating difference scores, for example, by subtracting the completion time of different parts of the TMT. However, it was recently demonstrated that psychomotor speed still significantly contributes to performance on the TMT difference score (21). Interestingly, a study that adequately controlled for reductions in basic cognitive abilities by calculating the interference score of the Stroop Colour Word test did not find an effect of chronic pain on task performance (12). These findings imply that a decrease in executive and attentional task performance may at least be partly attributable to a decline in psychomotor speed. In addition, factors such as pain catastrophizing and a reduction in sleep quality and opioid use may all influence cognitive functioning (1), and could therefore potentially account for the heterogeneous findings regarding attention and executive function in chronic pain.

The aim of the present study was to examine the extent to which chronic pain affects psychomotor speed, and whether the effects of chronic pain on executive and attention functions reflect a decline that extends beyond a reduction in this psychomotor slowing. In the present study, a repeated measures ANOVA was employed to allow for examining disproportional diminished task performance in participants with chronic pain with increasing task demands or duration (3). As such, it could be determined whether group differences on tasks represented specific executive and attention impairments. In line with previous findings (18), chronic pain participants were expected to perform more slowly on the neuropsychological tests. In addition, specific lower performance on the attention and executive function tests was anticipated (4,5,16,17).

METHODS

Participants
A total of 34 participants with chronic pain were involved in the study. Participants with chronic pain were included in cooperation with the Pain Clinic at the University Medical Centre Utrecht, the Netherlands (n=7), or were participants of a previous (unrelated) study (n=9) or were recruited through advertisements (ie, pain internet sites and oral advertising n=18). All participants reported persistent pain for >6 months (mean duration 11.7 years, median 6.5 years, range one to 50 years). Chronic pain conditions could be categorized as visceral pain (n=8), musculoskeletal pain (n=21), neuropathic pain (n=1) and other (n=5). Three participants suffered from more than one chronic pain condition. The majority of participants with chronic pain used pain medication: 17 participants used acetylsalicylic acid (paracetamol, 13 used a nonsteroidal anti-inflammatory drug (ie, diclofenac, diclofenac plus misoprostol, naproxen, ibuprofen), 11 were receiving opioids (ie, tramadol, fentanyl, acetaminophen plus codeine, acetaminophen plus tramadol), one used a tricyclic antidepressant (amitriptyline) and one used acetaminophen combined with caffeine. Four participants did not use any analgesic medication. Current pain intensity was rated using a 10 cm visual analogue scale of the Dutch translation of the McGill Pain Questionnaire (23). Thirty-two age- and IQ-matched control participants were also recruited. All participants reported to be free of a history of neurodegenerative disorders (eg, dementia), stroke, major depressive disorder, and alcohol or other substance abuse.

Premorbid IQ was estimated with the Dutch version of the National Adult Reading Test (24). Additionally, the Mini Mental State Examination (25) was administered to exclude participants with possible severe cognitive decline (defined as a score <24). No participants were excluded on this basis. The characteristics of the participants are presented in Table 1. Approval for the study was obtained from the Medical Ethics Committee of the University Medical Centre Utrecht. All participants signed an informed consent form.

Neuropsychological tests
All participants completed four tests to assess executive function and attention. First, the Stroop Colour Word test (10) was assessed to measure inhibition. This test consists of three cards, namely the Word (W) card, the Colour (C) card, and Colour/Word (C/W) card. The W card consists of 10 rows with 10 colour names in each row which are all printed in black ink. The participant is required to read aloud these colour names as fast as possible. For the C card, 10 rows are presented containing 10 coloured blocks per row, which the participant is required to name as fast as possible. The C/W card, colour names are printed in an incongruent colour, and the participant is required to name the colours in which the words are printed as fast as possible. The time needed to finish each of the cards was measured. Performance on the first two cards reflects basic cognitive abilities such as psychomotor speed and reading ability, whereas the third card specifically addresses inhibition. To establish diminished inhibition performance in chronic pain, a disproportional decline in performance should be present on the C/W card, ie, group differences should be more pronounced for the C/W card than for the W and C cards.

Second, the Trail Making Test (8) is a visual search task, measuring mental flexibility. The TMT part A consists of 25 encircled numbers that are randomly distributed on a sheet of paper. The participant is required to connect these numbers sequentially. On the TMT part B, both numbers and letters are distributed. This time, the participant is instructed to alternate between the numbers and letters (ie, 1, A, 2, B, etc). Completion times of part A and B were measured. Part A is considered to measure basic abilities such as motor speed and visual search (19), whereas part B examines mental flexibility. Again, to establish whether performance on part B extends beyond a mere reduction in basic cognitive abilities in chronic pain, a disproportional decline in TMT part B performance in the chronic pain group should be present, which extends beyond group differences in TMT part A performance.

Third, the Bourdon-Vos test (26) was administered to measure sustained attention. This test consists of 33 lines containing groups of three, four or five dots presented on a sheet of paper. Participants are instructed to count the groups with four dots as fast and accurately as possible. Completion time and the number of omissions (27) are recorded for each line. When only examining the total score (eg, total completion time), no information about the ability to sustain attention can be obtained. Here, a longer completion time might only indicate basic cognitive slowing per se, without an impaired ability to sustain attention over time. Therefore, the study calculated the average time to complete a line, and the average number of omissions for each line separately for the first one-half of the test (lines 1-17) and
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the second one-half of the test (lines 18-33). A specific decline in sustained attention should be reflected by increased slowing, as well as an increased number of omissions, on the second half compared with the first half of this test in the chronic pain group. In addition, the variability in performance on this task was calculated by examining the reaction time and omission scores per row for all 33 rows (28). Lower sustained attention performance was indicated by an increased variability, reflecting an increase in fluctuations in attention, in the completion time and the number of omissions over the different lines of the task.

Finally, the Zoo Map test of the Behavioural Assessment of the Dysexecutive Syndrome (29) was used to assess planning ability. This test consists of an unstructured first part and a structured second part. In the first part, participants are instructed to plan their route through a map of the zoo, visiting a selection of places while bypassing others. While planning the route, participants also need to obey to some rules. The second part consists of the same map with the same places that have to be visited, only this time instructions are provided about the precise order of visits. This second part, therefore, strongly reduces the involvement of planning functions. For both cards, the maximum score ranged between 0 and 8. Points were deducted if an error was made (ie, using a path more than once, deviations from the path, failure to make a continuous line and inappropriate places visited). Both planning and completion time were measured. A specific planning problem in chronic pain means that performance (raw score, planning or completion time) on the first unstructured card should be disproportionately affected compared to the second, more structured, card.

Potential confounders

Pain catastrophizing was measured with the Pain Catastrophizing Scale (PCS) (30-32), because this may be an important mediator of the cognitive impairments observed in chronic pain patients (33,34). The Geriatric Depression Scale (GDS) (35) was administered to measure depressive symptoms, which is a reliable indicator of depressive symptoms in both the elderly population and in younger adults (36,37). In the chronic pain group, the number of hours lying awake because of the pain was also measured with the McGill Pain Questionnaire. Finally, the influence of pain duration on performance was examined, because an increase in the duration of pain has been associated with an increase in brain grey and white matter changes (38,39), which likely affect cognitive function.

Procedure

Participants were contacted through the pain clinic, internet sites, oral advertising or were participants of a previous study. After informed consent was obtained, participants completed a single test session. Tests and questionnaires were always administered in the same order: participants first completed the questionnaires and then the neuropsychological tests (all in a fixed order). A total session lasted approximately 1 h to 1.5 h.

Statistical analyses

Logarithmic transformation was applied to the TMT part A and B completion time measures, and inverse normalizing transformation to the Bourdon-Vos omissions (both first half and second half), to the attention fluctuations reflected by the variability in completion time and omission score of each line of the Bourdon-Vos; and to the Zoo Map raw score, the planning and the completion time measures.

A 2×3 mixed-design ANOVA was used for the Stroop Colour Word test with group (control versus chronic pain patients) as the between-subject factor and completion time of the cards (W, C and C/W) as the within-subject factor. A 2×2 mixed-design ANOVA was performed for the TMT test, with test version (TMT-A, TMT-B) as the within-subject factor and group as the between-subject factor. A similar design was employed twice for the Bourdon-Vos test with the completion time (first half, second half) and the number of omissions (first half, second half) as separate within-subject variables and group as a between-subject factor. The variability in completion time and omissions of the different lines was analysed using a multivariate ANOVA (MANOVA) with group (control versus chronic pain) as the independent variable and the completion time and omission variance as dependent variables. A 2×2 mixed-design was also used for the Zoo Map test, where the analyses were run separately for the raw score, the planning time and the completion time of both cards. For all analyses the study focused on the main effect of group and on the interaction between group and the test variables. For all these analyses the sphericity assumption (Mauchly’s test), the homoscedasticity assumption (Box’s M test) and the homogeneity assumption (Levene’s test) were met.

T tests were employed to test for group differences in IQ, age and pain catastrophizing. Group differences in depressive symptoms were tested with the Mann-Whitney U test because these scores were not normally distributed. Sex distribution was examined with a Pearson x² test. For these and the above-mentioned tests, the level of significance was set at P<0.05.

Pearson correlations were calculated in the chronic pain group between current pain intensity, pain duration, pain catastrophizing, depressive symptoms and the hours lying awake on the previous night because of the pain on the one hand and the cognitive tasks on the other. This analysis was performed to examine whether an increase in pain severity, duration, depressive symptoms, catastrophizing or a reduction in sleep quality was related to a more pronounced decline in task performance. Mann-Whitney U tests were performed to examine potential group differences in cognitive task performance between chronic pain participants who did and did not use opioids. To reduce the risk of type I error due to multiple testing, the level of significance for the correlational tests was set at P<0.01.

Finally, insufficient effort has been reported in chronic pain patients, particularly in those seeking litigation (41,42). These factors may result in decreased neuropsychological test performance (43,44). To address this possibility, for each patient the present study compared their performance on each test to normative data and specifically examined the number of participants who performed in the impaired range (2 SD or more below average).

RESULTS

The chronic pain and healthy control groups did not differ significantly in age (t(64)=0.75, P=0.46), sex (2 [1.2]=2.30, P=0.13) and IQ (t(64)=1.88 [P=0.065]) (Table 1). Participants with chronic pain had in general higher PCS scores compared with the controls (t(64)=1.80 [P<0.05]) (36). Depressive symptoms were significantly increased in the chronic pain group (Z=-4.38 [P<0.001]). This variable (normalized using Blom transformation) was therefore used as a covariate in the subsequent repeated measures analyses. The performance of both groups on all tasks is presented in Table 2.

Inhibition

No significant main effect for group (F(1, 63)=0.25, P=0.62, η²=0.00) or a significant interaction between group and task performance (F(2, 126)=0.71, P=0.49, η²=0.00) was found on the completion time of the Stroop Colour Word test.

Mental flexibility

A significant main effect of group was observed for the TMT (F(1, 63)=7.74, P<0.01, η²=0.11). Overall, participants with chronic pain took longer to complete the TMT part A and part B than the controls. However, the interaction between group and task performance was not significant (F(1,63)=0.30, P=0.59, η²=0.00), implying that an increase in task demands (part B compared with part A) affected performance in the chronic pain and the control group to a similar extent.

Sustained attention

A significant main effect of group (F(1,63)=5.28, P<0.05, η²=0.07) was found for the completion time of the Bourdon-Vos test, indicating that, overall, the chronic pain group took longer to complete this test. The interaction between group and task performance was not
significant (F[1,63]=3.11, P=0.08, n²=0.05). Analyses of the omissions on this test revealed no significant main effect of group (F[1,63]=2.51, P=0.12, n²=0.04), but it did reveal a significant interaction effect between group and the number of omissions (F[1,63]=5.51, P<0.05, n²=0.08). While the control group performed slightly better with regard to the number of omissions in the second half compared with the first half of this test (t[31]=1.79, P=0.08), a marginally significant increase in the number of omissions in the second part of this test was found for participants with chronic pain (t[33]=-1.93, P=0.06). Finally, a significant effect was found for fluctuations in attention as indicated by the variability in the completion time and omission scores (F[2,62]=3.15, P=0.05, n²=0.09). Chronic pain patients were more variable with regard to the completion time (F[1,63]=5.67, P<0.05, n²=0.08), and a marginally significant effect appeared with regard to the omissions made on the different lines of this test (F[1,63]=3.11, P=0.08, n²=0.05).

Planning

Nearly all participants obtained the maximum raw score on the second card (with the exception of two controls and two participants in the chronic pain group). The study, therefore, did not analyze the raw score of the second card but simply analyzed the raw score of the first card with an ANOVA. No significant effect for group was found (F[1,62]=0.18, P=0.67, n²=0.00). For the planning time, neither the effect of group (F[1,61]=0.54, P=0.47, n²=0.01) nor the interaction between group and planning time (F[1,61]=0.01, P=0.91, n²=0.00) was significant. The same pattern of results was also obtained for the completion time measures, where the main effect of group (F[1,61]=0.03, P=0.86, n²=0.00) and the interaction between group and completion time (F[1,61]=0.99, P=0.32, n²=0.02) did not reach significance.

Current pain intensity, pain duration and confounders

Current pain intensity ranged between 0 mm and 79 mm on a 0 cm to 10 cm scale. The correlations between the current pain intensities and the cognitive task scores did not correlate significantly with pain duration, number of hours lying awake, depressive symptoms or the PCS scores. Finally, use of opioids was not associated with performance on the attention and executive function tests.

Effort to complete tasks

For the Stroop and TMT, age- and education-corrected normative data were available. For the Zoo Map test, no normative data were available but performance on this test could be converted to a profile score. This profile score ranged between 0 and 4. For the current analysis, a score of 0 or 1 was defined as impaired, a score of 2 as...
flexibility may be due to reductions in basic cognitive functions, present study is the first to indicate that slowing on a task of mental flexibility was affected in older chronic pain patients, compared with the first part, whereas the opposite pattern was observed in the control group. Also, the chronic pain group revealed increased fluctuations in attention, as indicated by an increased variability in completion time and omissions across the different lines of the task. To summarize, several executive and attentional functions (e.g., mental flexibility, planning, and inhibition) appear to be largely intact in chronic pain except for a specific decline in the ability to sustain attention.

In the present study, we found intact inhibition, which is in agreement with previous studies on selective attention or inhibition in this patient group (11,12). For the planning test, little is known about the potential disruptive effects of chronic pain on task performance. To our knowledge, we are the first to examine this function in chronic pain and show that planning is still intact in this group. Our study findings are further in line with previous studies that demonstrated diminished performance on sustained attention and mental flexibility tasks in chronic pain patients. Sjögren et al (5) examined neurological functioning of chronic pain patients on a sustained attentional performance reaction time test in which patients had to respond as fast as possible to auditory signals delivered over a 10 min period. The authors found longer reaction times for patients compared with controls for this test. Interestingly, they also found lower performance of the chronic pain patients on a finger tapping task, measuring motor speed. It thus remains unclear from these findings whether sustained attention was affected independently of diminished speed or whether the slower reaction times on the auditory reaction time test were due to impairments in speed in this patient group. The same holds for the study by Wiener et al (45). These authors demonstrated decreased performance on part B of the TMT in older chronic pain patients, but performance on a test for fine motor performance (the Grooved Pegboard) was also diminished in this patient group. Also in that study it remained unclear whether mental flexibility was affected or whether psychomotor slowing could explain the findings. The present study is the first to indicate that slowing on a task of mental flexibility may be due to reductions in basic cognitive functions, whereas difficulties in sustaining attention may be independent of these processes.

The mechanisms for the decline in attention and executive function in chronic pain patients remain largely unclear. Several different hypotheses have been postulated to explain reduced cognitive, and in particular attentional, functions in chronic pain patients. It has been argued that patients have an attentional bias toward painful sensations which may cause distraction from other stimuli. In a recent study, rats with monarthiritis caused by complete Freund’s adjuvant showed a marked decline in a sustained attention task (five-choice serial reaction time task) (46). After pain was induced, rats were less accurate and made more omissions on this task. Interestingly, this decline in performance was not altered by transient analgesic drug administration, indicating that distraction due to pain is probably not the main cause of attentional deficits in chronic pain.

Another hypothesis is that attention and pain share common resources (47). The experience of pain may reduce the capacity to attend to another stimulus because it captures attentional resources itself. In this perspective, it is expected that with increasing task demands cognitive task performance decreases disproportionately in chronic pain patients. Our results are not completely in line with this view because such a specific pattern of results was not observed for three different executive function tests. An alternative perspective proposes that perhaps attentional capacity is not impaired in chronic pain, but chronic pain patients have the tendency to allocate attention differently (48,49). An attentional bias toward stimuli, especially pain, has been described by the hypervigilance perspective. Recent data suggest that allocation of attention is different in chronic pain patients, and not necessarily only when pain-related information is involved (49). It seems reasonable to assume that the allocation of attention is particularly sensitive to long durations over time. Our present finding that participants with chronic pain have a specific difficulty in sustaining attention might therefore potentially reflect an underlying problem in the allocation of attention.

Increasing pain severity has been previously found to be associated with decreased cognitive task performance (3,45,50). For example, Grisart and Flachiyi (4) found that selective attention or inhibition, as assessed with the Stroop Colour Word task, was affected in chronic pain patients, but only in patients with pain of high intensity with a visual analogue scale score of 60 or more. In the present study, no significant association was observed between pain severity and performance on the neuropsychological tasks. Potentially, the level of pain intensity in our chronic pain sample was not severe enough to replicate the previously observed associations between pain intensity and cognitive task performance. Future studies on the pattern of attentional task performance in chronic pain patients could consider including patients with higher pain intensity levels. It is possible that the specific decline in task performance with increasing task demands is present in chronic pain patients with high pain intensity levels.

One potential limitation of our study is that we included a heterogeneous sample of participants with chronic pain. The extent to which this explains the absence of a specific executive deficit in our sample is unclear. However, in a previous study with the same heterogeneous patient sample, we were able to find significantly lower performance on multiple memory tests (50). Nevertheless, the current results should be replicated in more homogeneous patient samples.

Also, the attention and executive function domains cover multiple aspects of cognition, some of which were not addressed in the current study. For example, working memory and set shifting have also been denoted as aspects of executive function (6), while others have tried to differentiate between alerting, orienting and executive attention (51). Only several indices of attention and executive function were examined in the present study; future studies are therefore needed to obtain a more complete assessment of the executive and attention domains.

A final point of consideration concerns the high variability in performance on some of the tests (e.g., the TMT-B and the Zoo Map test) in the chronic pain group. This could possibly reflect low validity of these tests, but an increase in variances in test performance has often been reported in different clinical groups (52,53).
Executive and attention functions appear to be largely intact in chronic pain, with the exception of a significant decline in attentional performance on sustained attentional task performance. The lower performance on mental flexibility is presumably due to more basic cognitive processes such as psychomotor slowing. No impairment in inhibition or planning ability was found. This study provides new insights into executive function and attention deficits in chronic pain patients.

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SUMMARY

Executive and attention functions appear to be largely intact in chronic pain, with the exception of a significant decline in attentional resources on sustained attentional task performance. The lower performance on mental flexibility is presumably due to more basic cognitive processes such as psychomotor slowing. No impairment in inhibition or planning ability was found. This study provides new insights into executive function and attention deficits in chronic pain patients.

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