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DYNAMICS OF COGNITIVE ACTION CONTROL IN LATE-LIFE DEPRESSION DURING ACTION SELECTION

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List of Abbreviations:

- AES: Apathy evaluation scale
- C: Congruent Condition
- CAC: Cognitive action control
- cR2 : conditional R2
- CE: Congruency effect
- CPT-3: Continuous performance test, third version
- CSE: Congruency sequence effect
- DRS: Depressive retardation scale
- HC: Healthy controls
- HDRS: Hamilton depressive rating scale
- LLD: Late-life depression
- MoCA: Montreal cognitive assessment scale
- MCST: Modified Wisconsin card sorting test
- mR2 : marginal R2
- NC: Non-congruent Condition
- ST: Simon Task
- STAI: State and Trait Anxiety Inventory

1 **1. INTRODUCTION**

2

Late-Life Depression (LLD), (>60 years old), affects 7% of the population (World Health 3 4 Organization, 2017). Cognitive control deficits lie at the core of LLD being persistent during remission (Bhalla et al., 2006; Butters et al., 2004; Diniz et al., 2013), correlating with 5 disability and predicting depressive relapses (Alexopoulos et al., 2002, 2000). Cognitive 6 7 action control (CAC) refers to "a subset of cognitive control processes involved in the requirement to coordinate one's instantaneous urges vis-à-vis actions that concord with our 8 9 intentions or instructions" (Ridderinkhof et al., 2011). Such processes include cognitive inhibition, cognitive flexibility or inhibitory control. They can be altered during normal aging 10 (Maldonado et al., 2020; Rey-Mermet and Gade, 2018) but may be magnified in LLD. 11 12 Indeed, CAC deficits in LLD have been associated with higher suicidality (McGirr et al., 13 2012; Richard-Devantoy et al., 2015, 2011) possibly mediated by impulsivity (Dombrovski et al., 2011), and apathy (Feil et al., 2003; Funes et al., 2018). However, these studies lack an 14

15 integrative framework to explore in-depth CAC dysfunction.

The Simon task (ST) is a tool of choice to explore CAC in LLD as it measures electively the 16 17 integrity of the processes coding for perceptual and action monitoring (Hommel, 2011) and is negligibly influenced by other domains of cognition. During this task, subjects are instructed 18 19 to make a fast and accurate left or right hand response according to the color of a stimulus that is presented either to the left or right side of a central fixation point (Hedge and Marsh, 1975). 20 21 Conflict lies in the competition between the location (non-relevant information) and the color of the stimulus (relevant information). During non-congruent trials (i.e. when color and 22 location do not match) participants show longer reaction time (RT) and decreased accuracy 23 compared to congruent trials (i.e. when color and location match). These modifications 24

between non-congruent and congruent conditions, named the congruency effect (CE), are
believed to evaluate online CAC. Additionally, RT and error rates are greater when trials
congruency alternates (Gratton et al., 1992). This so-called congruency sequence effect (CSE;
Duthoo et al., 2014) is an adaptation of CAC to conflict (Botvinick et al., 2001). Hence, the
ST provides a unifying paradigm to evaluate online (CE) and adaptive CAC (CSE).

30 Using the ST, depressive symptoms have been associated with larger CE (Ng et al., 2012) 31 interpreted as a poorer performance. However, the association between depressive symptoms and adaptive CAC has yielded mixed results: decreased CSE in depression during negative 32 feedback (Holmes and Pizzagalli, 2007), no association between symptoms severity and CSE 33 34 (Ng et al., 2012) and increased CSE in depressed participants (van Steenbergen et al., 2012). This discrepancy might be due to the use of averaged RTs, thereby omitting the temporal 35 dynamics of CAC. To resolve this, we used the activation-suppression model (Ridderinkhof, 36 2002a). This model incorporates the temporal dynamics of top-down suppression of 37 38 impulsive action selection by distinguishing a fast and early impulsive action selection from a 39 late and slow selective response suppression. Impulsive action selection results in increased errors for the fastest responses of the participants in conflict situations while selective 40 response suppression of the non-relevant information takes time to buildup and is most 41 42 evident at the slowest RTs. Discretizing CAC by reaction times is particularly relevant in LLD because it is suggested that abnormal cognitive control is mediated by decreased 43 processing speed (Nebes et al., 2000; Sheline et al., 2006; Snyder, 2013). 44

The main purpose of the present study is to show that patients with LLD have a deficit in both online and adaptive CAC. Secondly, we aim to show that LLD patients have specific patterns of impulsive action selection and selective response suppression. We hypothesized an impulsive behavior in LLD reflected by increased fast errors and lower selective suppression. Finally, we explored how impulsive action selection and selective response suppression of both online and adaptive CAC were associated to apathy and impulsivity.

8 2. MATERIALS AND METHODS

9

7

10 2.1. Participants

11

12 Thirty-one elderly diagnosed with DSM-5 criteria of Major Depressive Disorder and 31 healthy elderly controls (HC) participated. LLD group individuals were recruited by their 13 psychiatrist, while HC were recruited from local retirees associations. Inclusion and exclusion 14 15 criteria were assessed during a psychiatric interview by a trained old-age psychiatrist. Inclusion criteria for the LLD group were: Hamilton Depressive Rating Scale (HDRS, 16 17 Hamilton, 1960) score > 10. Exclusion criteria were: age < 60 years-old, schizophrenia, bipolar and neurological disorders, history of addiction (tobacco excluded), uncorrected 18 19 vision or motor impairments, Montreal Cognitive Assessment Scale (MoCA, Nasreddine et 20 al., 2005) score < 20 (Blair et al., 2016; Dautzenberg et al., 2021), cerebral modulation during the last 6 months. Written informed consents were collected. The study was approved by the 21 22 local ethics committee of Rennes University Hospital in accordance with current French 23 legislation.

24

25 **2.2. Experimental design**

26

27 We used the Apathy Evaluation Scale (AES, Marin, 1991), the Depressive Retardation Scale (DRS, Widlöcher, 1983), the Urgency Premeditation Perseverance Sensation seeking 28 29 impulsive behavior scale (UPPS, Whiteside and Lynam, 2001) exploring four dimensions of 30 impulsivity, and the State and Trait Anxiety Inventory (STAI, Spielberger et al., 1983). 31 HDRS quantified depression severity (LLD patients only). General cognitive function was assessed with the MoCA. The Stroop (Golden, 1978), Trail Making Test A and B (Reitan, 32 1958), Modified Wisconsin Card Sorting (MCST, Nelson, 1976) and Continuous 33 34 Performance tests (CPT-3, Conners et al., 2018) assessed cognitive control, cognitive 35 flexibility, set shifting and sustained plus selective attention, respectively.

36

37 2.3. Task and Procedure

38

The ST (**Figure 1**) was programmed under E-prime Professional version 2.0. This task comprised a familiarization phase (72 trials) followed by an experimental one (five blocks of 72 trials, with a pause between each block). All stimuli were displayed on a 60 cm white computer screen, placed 90 cm away from the participant. Each trial began with a black fixation cross displayed at the center of the screen for 1000 ms. A blue or yellow circle (target) of 2.1 cm diameter appeared on the right or left side of the cross. The distance
between the center of the computer screen and the edge of the circle was of 1.1 cm.

3 Participants were asked to press a blue or yellow button on a computer keyboard with their

4 right or left hand, as quickly and accurately as possible, according to the color of the target 5 while ignoring its position. The position of the target defined the congruency of the trial. The 6 target disappeared as soon as participants pressed one of the two buttons and a new target 7 appeared after a 1550 to 1650 ms randomly defined interval. The position of the color button 8 was pseudo-randomized across participants, and all conditions were pseudo-randomized with

- 9 an equal number of trials in each condition.
- 10

11 **2.4. Behavioral Analysis**

12

Only responses from the experimental phase with RT between 200 and 1500ms and less than
three standard deviations from the mean individual RT were analyzed (1.66 % of the trials

15 removed).

Main analyses focused on between-group differences in RT and accuracy rates between the 16 congruent (C) and non-congruent (NC) conditions to assess the CE, adding the interaction 17 18 with preceding congruency for evaluating the CSE. The dynamics of CAC (i.e., impulsive 19 action selection and selective response suppression) were assessed by distributional analyses (Ridderinkhof, 2002b). Impulsive action selection is represented by conditional accuracy 20 functions (CAF) which plot correct responses as a function of the RT distribution of all 21 22 *responses* discretized in 5 bins (62 trials per bin per participant) for each condition (C or NC) and each group (HC or LLD). An increased impulsive action selection corresponds to a lower 23 accuracy rate at the first bin of RT (i.e. fastest responses). Selective response suppression is 24 25 represented by delta plots which plot the congruency effect (difference in RT between NC and C conditions) as a function of the RT distribution of the correct responses, discretized in 5 26

bins. We then estimated the slopes between these bins (four slopes). According to the
activation-suppression model, the steeper the fourth slope is, the more effective the selective
suppression on slow responses. We performed the CAF and delta-plots considering

30 systematically the current trial as a function of the congruency of the preceding trial (CSE).

Three participants, 2 LLD and 1 HC, were excluded from the distribution analyses because20% of their completed trials were missing.

33

34 2.5. Statistical Analysis

35

Statistical analyses were performed using R software (version 3.0; R core team, 2017). A
significance threshold of two-sided p < 0.05 was used.

38

39 2.5.1. Patients characteristics

40 Demographic, clinical and psychological data were compared between the LLD and HC 41 groups using χ^2 tests for qualitative variables and independent t-tests (or Wilcoxon's signed-

- 42 rand test, depending on visual inspection of Q-Q plots).
- 43

1 2.5.2. Behavioral data

2 As RTs were not normally distributed, they were first converted using a logarithmic3 transformation.

- 4 The CSE on RT was estimated using a linear mixed model comparing the log RTs between 5 the four conditions and both groups. The fixed factors were the congruency of the preceding trial (Pre_Cong: Pre_congruent vs. Pre_non-congruent), the congruency of the current trial 6 7 (Congruency: congruent vs. non-congruent) and the group (LLD and HC). Participant was 8 added as a random intercept. Mixed models optimize statistical power as they use all available data, account for intra-subject repeated measures correlation and easily handle unbalanced 9 10 designs (Hox et al., 2017). As recommended by Nakagawa and Schielzeth (2013), we reported the conditional (cR2) and marginal R2 (mR2) using the {MuMIn} package (Bartoń, 2013). 11 The linear mixed model was executed using the lme function of the {nlme} package (Pinheiro 12
- 13 et al., 2013). Post-hoc Tukey-Kramer tests were performed using the {multcomp} package
- 14 (Hothorn et al., 2008).

15 The CSE on accuracy was estimated using a mixed logistic regression model comparing the

- 16 correct responses between the four conditions and both groups. The fixed factors were the
- 17 congruency of the preceding trial, the congruency of the current trial and the group.

18 Participant was added as a random intercept. The model was executed with the glmer function

19 of the {lme4} package (Bates et al., 2015). P-values from the logistic models were derived

- 20 from the Wald Chi-square tests using the {car} package (Fox and Weisberg, 2011).
- 21 The **CE on RT and accuracy** were estimated from the effect of the congruency of the current
- trial from the respective models.
- Impulsive action selection was tested using a mixed model logistic regression with accuracy
 at the first bin of the CAF as the independent variable. 'Group' and 'congruency of the
 current trial' were fixed factors while 'participant' was added as a random intercept.
- 26 Selective suppression was tested with a linear regression mixed model with steepness of the
- 27 fourth slope of the delta plots as independent variable and with the same factorial design as
- 28 for impulsive action selection.
- 29
- 30 Statistically different between-group sociodemographic variables were included as covariates.
- 31

32 **2.5.3.** Behavioral and Clinical Data Relationships

- Associations between the ST indices and the clinical scores of LLD were tested usingspearman correlations (detailed in supplemental methods).
- 35
- All codes are available in supplemental material. Clinical and behavioral data are available at
 the Mendeley data repository (10.17632/5644kyzncp.1).
- 38
- **39 3. RESULTS**
- 40

41 **3.1. Patient characteristics**

42

Between-group sociodemographic, clinical and neuropsychological variables are summarized in **Table 1**. Twenty-seven (87%) LLD patients were taking antidepressants. LLD patients had lower educational status (p<0.001) than HC but comparable age (p=0.11) and sex ratio (p=0.57). Because of the expected effect of education status on cognitive performance (Seblova et al., 2020), all following between-group analyses were adjusted on education status.

7

8 **3.2.** Between group difference on the CE and CSE

9

10 **3.2.1. Congruency Effect**

RTs were similar between the LLD patients and HC (Group: log(RT): $F_{(1, 59)} = 3.17$, p = 0.07). 11 RTs were longer and accuracy rates lower for non-congruent than for congruent trials across 12 all participants (Congruency: $\log(RT)$, F(1, 18362) = 2350.39, p < 0.0001, $mR^2 = 0.24$, $cR^2 =$ 13 0.33; accuracy, $\chi^2_{(1, 19305)} = 246.34$, p < 0.0001, $mR^2 = 0.13$, $cR^2 = 0.49$). This CE was 14 significantly different between the groups on accuracy rates (Group × Congruency: accuracy, 15 γ^2 (1. 19305) = 15.11, p < 0.001) but not for RT (Group × Congruency: log(RT), F(1. 18362) = 16 3.45, p = 0.06). Post-hoc analysis showed that LLD patients made more errors on congruent 17 18 trials (Group: z = -3.91, p < 0.001) which was not found in non-congruent trials (Group: z = -1.24, p = 0.57). 19

20

21 **3.2.2.** Congruency sequential effect

Across groups, accuracy rates were higher for trials that followed congruent trials in 22 comparison to trials that followed non-congruent trials, as opposed to RTs (Pre_Cong: 23 accuracy, $\chi^2(1, 19296) = 51.58$, p < 0.0001; log(RT), $\chi^2(1, 16888) = 0.014$, p = 0.91; Figure 2). CE 24 was associated with the congruency of the preceding trial (RT: Congruency × Pre_Cong: 25 log(RT), χ^2 (1, 16888) = 2308.51, p < 0.0001; accuracy, χ^2 (1, 19296) = 15.11, p = 0.0001). Post-hoc 26 27 comparisons indicated that the CE was larger when the preceding trial was congruent for both RT (Pre_Cong: log(RT), z = -9.06, p < 0.001) and accuracy (Pre_Cong: accuracy, z = -9.76, p 28 < 0.001). No group differences were found on the CSE (RT: Group x Congruency x 29 Pre_Cong: log(RT), χ^2 (1, 16888) = 0.48, p = 0.49; accuracy, χ^2 (1, 19296) = 0.58, p = 0.44). 30

31

32 **3.3. Impulsive action selection**

33

34 **3.3.1.** Between group differences on online impulsive action selection

Figure 3 shows the dynamics of accuracy as a function of RT for congruent and noncongruent conditions. Across groups, we found a reduced accuracy on the fastest RTs during non-congruent trials (Congruency: χ^2 (1, 3927) = 273.57, p < 0.0001; mR^2 = 0.43, cR^2 = 0.53), showing greater impulsive action selection in the non-congruent condition. The RT of the first bins were equivalent between the groups (Group: log(RT): F(1, 59) = 0.65, p = 0.42), even when only considering the congruent trials (Group: log(RT): F(1, 59) = 0.04, p = 0.27).

- 41 While no main group differences across congruency was found (Group: accuracy, χ^2 (1, 59) =
- 42 0.99, p = 0.32), there was a significant interaction between group and congruency (Group x
- 43 Congruency: accuracy, χ^2 (1, 3927) = 10.77, p = 0.01). Post-hoc analysis indicates that LLD

- 1 patients made more errors than HC during congruent trials (Group: z = -3.44, p = 0.003) and
- 2 not for non-congruent trials (Group: z = -0.86, p = 0.81).
- We found no effect of the group on the reaction times at the first quantile (Pre_Cong x Congruency x Group: $\log(RT)$: F_(1, 59) = 0.46, p =0.50) particularly when congruent conditions
- 5 were preceded by non-congruent conditions (Group: z = 0.036, p = 0.73).
- 6

7 **3.3.2.** Between group differences on adaptation of impulsive action selection

8 Figure 4 reports the CAFs of each group as a function of the congruency of the preceding 9 trials. Across groups, the accuracy of the fastest responses was dependent of the congruency 10 of the preceding trial (Pre_Cong: χ^2 (1, 59) = 116.95, p < 0.0001) as well as the CSE 11 (Pre_Cong x Congruency: χ^2 (1, 59) = 40.28, p <0.0001). Hence, an adaptation effect was 12 present on the fastest responses.

- 13 There was no significant interaction between the group and CSE (Group x Pre_Cong x 14 Congruency: χ^2 (1, 59) = 2.46, p =0.12). As the graphical reading of the CAFs suggests
- between-group differences in impulsive action selection when congruencies in trials change (grey plots in **Figure 4**), we explored an association between 'Group' and impulsive action selection for each pair of congruency sequences. Impulsive responses in LLD patients were
- greater when congruent trials occur after non-congruent trials (Group, z = -3.13, p = 0.03)
- 19 compared to HC (Figure 4D), not found in other sequences of adaptation (all p = 0.99).
- 20

21 **3.4. Selective response suppression**

22

23 **3.4.1.** Between group differences on online selective response suppression

Figure 5 displays the delta plots for each group. As predicted by the activation–suppression model, the fourth slope was steeper for the slowest RT, without between group differences (Group: slope, $F_{(1, 56)} = 9.55$, p = 0.008; $mR^2 = 0.11$, $cR^2 = 0.89$).

27

28 **3.4.2.** Between group differences on adaptation of selective response suppression

Delta plots as a function of trial sequence are represented in supplementary materials (**Figure S1**). We did not find any significant differences in selective suppression according to the congruency of the preceding trial (Pre_Cong: slope, χ^2 (1,56) = 0.08, p = 0.78) and no significant interaction between group and preceding congruency (Group x Pre_Cong: slope, χ^2 (1,56) = 0.98, p = 0.32).

34

35 **3.5.** Associations between CAC metrics and symptoms severity in LLD

36

We did not find any significant associations between CAC metrics and apathy, impulsivity,depression or anxiety scores in LLD. These results are detailed in supplemental material.

39 40

41 **4. DISCUSSION**

42

1 We combined measures of online CAC (CE) and adaptive CAC (CSE) from the ST to the 2 activation-suppression model (Ridderinkhof, 2002a) to assess impulsive action selection and 3 response suppression. We found mixed results on the integrity of CAC in LLD: while LLD 4 participants showed reduced cognitive control during impulsive action, they had a more 5 effective selective response suppression compared to HC.

6

4.1. Influence of Depression on Global Cognitive Action Control

7 8

9 Regarding online CAC, we found increased errors in LLD patients on congruent conditions,
10 unlike Ng and colleagues (2012) who found increased error rates in non-congruent trials
11 among sub-depressive adults. However, the task they used required working memory, which
12 may affect CAC by increasing cognitive load (Engström et al., 2017). In contrast, our simpler
13 task was specifically designed to assess the CAC.

14 Regarding overall adaptive control abilities, we found the expected CSE on both accuracy and

15 RTs, suggesting the task was adequately performed. We did not find any group differences,

16 suggesting preserved overall adaptive control abilities in LLD, as shown in adult depression

17 (Dillon et al., 2015; van Steenbergen et al., 2012).

18 The increased errors during congruent trials among LLD participants compared to HC suggest19 a subtle decreased facilitation of action selection. Indeed, congruent trials facilitate the

20 transformation of a spatial stimulus location into a motor response (Hasbroucq and Guiard,

21 1991; Kornblum and Lee, 1995). Our results suggest this facilitation of the performance by

the task-irrelevant dimension of the stimulus is reduced in LLD. In the ST, the CE specifically informs on response selection, as the conflict operates between the response set and the taskirrelevant dimension of the stimulus. This explains why this CE was different from the

interference effect at the Stroop test, where the conflict occurs at the stage of stimulus identification (Li et al., 2014).

27

4.2. Influence of depression on the dynamics of CAC

29

30 4.2.1. Influence of depression on impulsive action selection

CEs associated to LLD are underpinned by an increased impulsive online action selection in congruent condition. All participants had increased errors in non-congruent conditions for the fastest RT in accordance with the activation-suppression hypothesis. LLD participants showed a greater impulsive action selection on congruent trials suggesting a decreased facilitation by the congruent situation, hence a reduced sensitivity to the irrelevant dimension of the stimulus (Dillon et al., 2015).

Analysis of impulsive action selection adaptation suggests that the impulsive tendency of LLD may be due to a reduced disengagement of attention from conflict. Similar to HC, LLD improved their responses when a NC trial preceded a NC trial, but showed increased impulsivity when a NC trial preceded a C trial. Thus, LLD subjects were less "helped" by the stimulus location to promote fast and accurate responses after conflict. This is consistent with altered adaptation to 'aversive' stimulus like cognitive conflict (Botvinick et al., 2001;

43 Dreisbach and Fischer, 2015) in depression (Disner et al., 2011; Eshel and Roiser, 2010;

- 1 Meiran et al., 2011). This has been related to impaired attention (Nicoletti and Umiltà, 1994)
- 2 which LLD participants confirmed by lower CPT scores than HC.
- 3

4 4.2.3. Influence of depression on selective response suppression

5 LLD patients displayed a stronger selective response suppression than HC. This result aligns 6 with studies showing increased electrophysiological markers of inhibition during Go/No-Go 7 task in depression (Ruchsow et al., 2008; Zhang et al., 2016). The authors suggested that LLD 8 participants require additional resources to activate inhibition. In our study, mild to moderate 9 depressive subjects may have over-recruited their inhibition resources to improve their 10 performance in the slowest responses. Indeed, HC's last quantile of RT were approximately 11 100ms faster than LLD. A couple of studies found similar results with depressed individuals 12 taking more time to execute a more effective inhibitory control of pre-potent responses than

- 13 HC (Dillon et al., 2015; Vallesi et al., 2015).
- The absence of adaptive selective suppression across participants reproduces Ridderinkhof
 (2002a) as Wylie and colleagues (2010). This result is sound as the suppression of impulsive
- 16 responses, maximal for the slowest RT, should be less affected by the congruency of the
- 17 preceding trial.
- 18

19 **4.3.** No association between clinical features and CAC metrics

20

No significant correlation between CAC measures and clinical scores were found in LLD, similarly than previous results (Austin et al., 1999; Quinn et al., 2012). This might be explained by our sample wherein 80% of the patients had mild to moderate depressive symptoms (Zimmerman et al., 2013). As the performance at ST would not have been specific for too severely depressed elderly (Staub et al., 2013), we may have failed to find correlations

- 26 between clinical severity and CAC measures.
- LLD patients had higher scores in impulsive urgency and perseverance traits consistent with
 depressive disorder (d'Acremont and Van der Linden, 2007). The absence of association
 between impulsivity-trait and CAC has been found previously, reflecting poor consistency
 between self-reported temperament and cognitive measures in laboratory settings (Cyders and
- 31 Coskunpinar, 2011; Stahl et al., 2014).
- Explorative results found no association between anxiety and impulsive action selection nor response suppression. They also suggest a non-linear relationship between anxiety and online CE which might explain the mixed evidence for an association between cognitive inhibition and anxiety (Beaudreau and O'Hara, 2009; Delphin-Combe et al., 2016; König et al., 2021). We observed that anxiety and premeditation as perseverance impulsivities were correlated in LLD, which are in turn associated with shortest CE on RT, consistent with the meta-analysis from Berg and colleagues (2015). However, these analyses have to be interpreted with caution
- 39 as they are insignificant after p-value correction.
- 40

41 **4.4. Limitations**

- 42
- 43 To better consider our findings, several issues might be raised.

- The magnitude of the reduced facilitation after conflict found in LLD suggests that it might be
 a cognitive marker of the LLD. It should be outlined however that while statistically
- 3 significant, the group difference is marginal and questions the translational potential of this
- 4 finding. This is likely to be due to the criteria selection in terms of cognitive impairment we
- 5 used in order to homogenize the population sample.
- 6 By selecting participants on an estimation of their ability to complete the ST via the MoCA,
- 7 we may have excluded participants with poor executive functions, thereby reducing the effect
- 8 size of CAC deficit among LLD. Yet, the ST has shown to be sensitive to modifications in
- 9 CAC between groups of subjects with preserved executive functioning (Wylie et al., 2010).10 We cannot rule out the possibility that some members of our sample may have an underlying
- 11 neurodegenerative condition that is contributing to these low scores. However, a reduced
- 12 MoCA scores has been found in adult depression without mild cognitive impairment (Blair et
- 13 al., 2016; Dautzenberg et al., 2021), a score of 20 being the lowest bound of MoCA scores in
- 14 this population. Hence, we chose a lower threshold of 20 for inclusion and the absence of
- 15 DSM-5 criteria of mild neurocognitive impairment.
- 16 Most of LLD patients were under antidepressants and anxiolytic drugs. While being difficult
- to disentangle effects of drugs from effects of depression, tapering medication was notfeasible for ethical reasons.
- 19 Other parameters of conflict adaptation, as incentives or emotion, were not considered while
- 20 important (Shenhav et al., 2013) notably in depression (Grahek et al., 2019). Further studies
- evaluating them may shed light on complementary mechanisms involved in CAC in LLD.
- 22
- 23

24 CONCLUSION

Our study suggests that LLD subjects have a modified CAC, originating from increased impulsive errors during congruent trials, which suggest that LLD individuals have marginal difficulties to disengage their attention from conflict. Also, LLD was characterized by an increased selective suppression suggesting an over-recruitment of inhibition resources. Further research should investigate how motivation and emotion might influence these alterations.

- 31
- 32

33 AUTHORS' CONTRIBUTIONS

- 34 JCR: Conceptualization; Data curation; Formal analysis; Funding acquisition; Investigation;
- 35 Resources; Software; Validation; Visualization; Writing original draft.
- 36 JFH: Conceptualization ; Methodology; Software; Validation; Writing review & editing.
- 37 JD: Conceptualization ; Methodology; Validation; Writing review & editing.
- 38 MG: Conceptualization ; Investigation; Validation; Writing review & editing.
- 39 DD: Funding acquisition; Supervision
- 40 GR: Conceptualization; Funding acquisition; Investigation; Methodology; Project
- 41 Administration; Supervision; Validation; Writing review & editing.
- 42 All authors read and approved the final manuscript.
- 43

1

2 CONFLICTS OF INTEREST

- 3 None
- 4
- 5

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8 financial support to the study. We are thankful to all the volunteers who participated in this
9 study.

10

11 **TABLES**

12

13 <u>Table 1</u>

14 Demographic and Clinical Data for LLD and HC Groups

15

	LLD (n = 31)	HC (n=30)	Statistic	p-value
Age (years)	72 ± 14 $60 - 88$	69 ± 10 60 - 83	t = -1.65	0.11
Gender (M:F)	10:20	8:22	$\chi^2 = 0.32$	0.57
Education (years)	12 ± 5 $8 - 16$	14 ± 3 $9 - 21$	t = 3.81	<0.001
HDRS	17 ± 9 $11 - 32$	NA		
AES	43 ± 14 $23 - 58$	19.50 ± 2.25 18 - 24	W = -14.22	<0.001
DRS	$6 \pm 11 \\ 0 - 12$	2 ± 2 $0 - 2$	W = -11.68	<0.001
Anxiety				
STAI-A	57 ± 27 22 - 98	27 ± 10.75 20 - 45	W = -8.36	<0.001
STAI-B	52 ± 6 $33 - 68$	35 ±17.25 22 - 49	t = -6.69	<0.001

Impulsivity

UPPS-Urgency	31.00 ± 7.00 17.00 - 38.00	24.50 ±13.25 16.00 - 38.00	t = -2.64	0.011
UPPS-Premeditation	21.00 ± 7.00 12.00 - 31.00	19.00 ± 5.00 12.00 - 28.00	t = -0.49	0.626
UPPS-Perseverance	19.00 ± 9.00 12.00 - 30.00	16.50 ± 5.25 10.00 - 22.00	W = 308	0.010
UPPS-Sensation	19.00 ± 9.00 12.00 - 41.00	23.50 ± 10.75 12.00 - 39.00	t = 1.40	0.168
МоСА				
Raw score	25 ± 4 $20 - 30$	28 ± 2 $23 - 30$	W = 4.99	<0.001
Visuospatial/Executive	4 ± 2 $1 - 5$	5 ± 1 $3 - 5$		
Naming	3 ± 0 $2 - 3$	3 ± 0 $3 - 3$		
Attention	6 ± 1 $3 - 6$	6 ± 0 $4 - 6$		
Language	2.5 ± 1 $2 - 3$	3 ± 1 $2 - 3$		
Abstraction	$1 \pm 1 \\ 0 - 2$	$\begin{array}{c} 2 \pm 0 \\ 0 - 2 \end{array}$		
Delayed Recall	3 ± 2 $0 - 5$	5 ± 1 $0 - 5$		
Orientation	6 ± 1 $4 - 6$	6 ± 0 5 - 6		
Stroop test				
Interference score	-1.31 ± 13.49 -18.10 - 27.00	1.20 ± 13.78 -20.53 - 22.87	t = 1.43	0.159

Correct responses in Interference	23.00 ± 22.25 0.00 - 54.00	39.00 ± 13.00 19.00 - 67.00	t = 3.89	<0.001
TMT - A	46 ± 15.75 27.00 – 196.00	36.50 ± 18 16.00 - 78.00	W = 543	P = 0.07
Cognitive flexibility				
TMT B-A	$-0.73 \pm 14.43 \\ -22.87 - 42.00$	0.00 ± 12.11 -27.00 - 54.00	W = 284	0.503
MCST perseverative errors	3.00 ± 3.75 0.00 - 9.00	1.50 ± 2.00 0.00 - 6.00	t = -1.89	0.064
MCST Complete categories	6.00 ± 1.75 2.00 - 6.00	6.00 ± 0.00 4.00 - 6.00	W = 402.0	0.066
CPT 3				
Detectability	55.00 ± 19.50 $41.00 - 83.00$	47.00 ± 15.25 30.00 - 69.00	t = -2.85	0.060
HRT	58.00 ± 9.50 50.00 - 90.00	51.00 ± 10.00 39.00 - 68.00	t = -3.72	<0.001
HRT SD	52.00 ± 12.5 33.00 - 85.00	47.00 ± 7.00 41.00 - 61.00	W = 188.5	0.018
Simon task				
Reaction times (ms)	_			
Mean	519 ± 190 201 - 1282	472 ± 145 237 - 1145	F=3.17*	0.07
Congruent	484 ± 174 201 - 1282	439 ± 134 237 - 1145		
Non-Congruent	556 ± 184 244 - 1280	507 ± 142 363 - 1053		

Accuracy rates (%)

Mean	94.87 ± 3.23 85.62 - 99.37	96.05 ± 3.05 83.92 – 99.68	$\chi^2 = 3.33^*$	0.07
Congruent	97.91 ± 2.19 91.82 – 99.75	99.14 ± 0.97 96.79 – 99.75		
Non-Congruent	91.83 ± 5.00 79.20 - 98.74	92.96 ± 6.2 69.68 - 100		

Summary descriptive results (except for Simon task results):

Median ± Interquartile Range

Min - Max

Acronyms: AES : Apathy Evaluation Scale, χ^2 : Chi-squared test, CPT-3 : Conners Continuous Performance Test third Edition, DRS : Depression Retardation Scale, F: Ftest, HC: Healthy Controls, HDRS : Hamilton Depressive Rating Scale, HRT: mean Hit Reaction Times, HRT SD: standard deviation of Hit Reaction Times , LLD: Late-Life Depression participants, MCST: Modified Card Sorting Test; MoCA: Montreal Cognitive Assesment, STAI: State-Trait Anxiety Inventory, t: Student's t-test , TMT B-A: difference in scores between versions B and A of the Trail Making Test, UPPS: UPPS impulsive scale, W: Wilcoxon's signed-rank test.

* Group Effects from mixed-models

1 2

23

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4 FIGURES

5

6 <u>Figure 1</u>

7 In the Simon task, subjects are instructed to make a left- or right-hand response as fast and as 8 accurately as possible according to the color of a stimulus that is presented either to the left or 9 right of a fixation cross. A. In congruent trials both dimensions of the stimulus (location and 10 color) point to the desired response, facilitating the response of the subject. B. In noncongruent trials, the automatic processing of the spatial location of the stimulus (plain arrow) 11 conflicts with the response requested by the controlled processing of color (dotted arrow). The 12 13 spatial location of the stimulus has to be ignored to produce a correct response, thus leading to increased RT and errors. 14

15

16 <u>Figure 2</u>

17 Mean RT and accuracy rates (% correct) for congruent (C) and non-congruent (NC) trials

18 with respect to the congruency of the preceding trial (preceding trial congruent (PreC) and

- 19 non-congruent (PreNC)) in each group. Situations in which the congruency of sequential trials
- 20 changes are shaded in grey. Error bars represent standard error of the mean.

1

6

2 <u>Figure 3</u>

- 3 Conditional accuracy functions for congruent and non-congruent trial LLD and HC. Error
- 4 bars represent the standard error of the mean.
- 5 *: Significant between-group differences.

7 Figure 4

- 8 Conditional Accuracy Functions in individuals with LLD and HC for congruent (C) and non-
- 9 congruent (NC) trials preceded by congruent trials (PreC) or by non-congruent trials (PreNC).
- 10 Accuracy rates in the context of shift in congruency are reported in grey-shaded cells. Error
- 11 bars represent the standard error of the means.
- 12 *: Significant between-group differences.
- 13

14 **<u>Figure 5</u>**

- 15 Delta-plot of reaction times of LLD and HC participants. Error bars represent the standard
- 16 error of the mean.
- 17 *: Significant between-group differences between the last slopes.
- 18
- 19

20 **REFERENCES**

- 21 Alexopoulos, G.S., Buckwalter, K., Olin, J., Martinez, R., Wainscott, C., Krishnan, K.R.R.,
- 22 2002. Comorbidity of late life depression: an opportunity for research on mechanisms and
- treatment. Biol. Psychiatry 52, 543–558.
- 24 Alexopoulos, G.S., Meyers, B.S., Young, R.C., Kalayam, B., Kakuma, T., Gabrielle, M.,
- 25 Sirey, J.A., Hull, J., 2000. Executive dysfunction and long-term outcomes of geriatric
- 26 depression. Arch. Gen. Psychiatry 57, 285–290.
- 27 Austin, M.P., Mitchell, P., Wilhelm, K., Parker, G., Hickie, I., Brodaty, H., Chan, J., Eyers,
- 28 K., Milic, M., Hadzi-Pavlovic, D., 1999. Cognitive function in depression: a distinct pattern
- 29 of frontal impairment in melancholia? Psychol Med 29, 73–85.
- 30 https://doi.org/10.1017/s0033291798007788
- 31 Bartoń, K., 2013. MuMIn: Multi-model inference, R package version 1.10.0.
- 32 Bates, D., Mächler, M., Bolker, B., Walker, S., 2015. Fitting Linear Mixed-Effects Models
- 33 Using lme4. Journal of Statistical Software 67, 1–48. https://doi.org/10.18637/jss.v067.i01
- 34 Beaudreau, S.A., O'Hara, R., 2009. The Association of Anxiety and Depressive Symptoms
- 35 with Cognitive Performance in Community-Dwelling Older Adults. Psychol Aging 24, 507–
- 36 512. https://doi.org/10.1037/a0016035
- 37 Berg, J.M., Latzman, R.D., Bliwise, N.G., Lilienfeld, S.O., 2015. Parsing the heterogeneity of
- 38 impulsivity: A meta-analytic review of the behavioral implications of the UPPS for
- 39 psychopathology. Psychol Assess 27, 1129–1146. https://doi.org/10.1037/pas0000111
- 40 Bhalla, R.K., Butters, M.A., Mulsant, B.H., Begley, A.E., Zmuda, M.D., Schoderbek, B.,
- 41 Pollock, B.G., Reynolds, C.F., Becker, J.T., 2006. Persistence of neuropsychologic deficits in
- 42 the remitted state of late-life depression. Am J Geriatr Psychiatry 14, 419–427.
- 43 https://doi.org/10.1097/01.JGP.0000203130.45421.69
- 44 Blair, M., Coleman, K., Jesso, S., Desbeaumes Jodoin, V., Smolewska, K., Warriner, E.,
- 45 Finger, E., Pasternak, S.H., 2016. Depressive Symptoms Negatively Impact Montreal
- 46 Cognitive Assessment Performance: A Memory Clinic Experience. Can J Neurol Sci 43, 513–
- 47 517. https://doi.org/10.1017/cjn.2015.399

- 1 Boone, K.B., Lesser, I.M., Miller, B.L., Wohl, M., et al, 1995. Cognitive functioning in older
- 2 depressed outpatients: Relationship of presence and severity of depression to
- neuropsychological test scores. Neuropsychology 9, 390–398. https://doi.org/10.1037/08944105.9.3.390
- 5 Botvinick, M.M., Braver, T.S., Barch, D.M., Carter, C.S., Cohen, J.D., 2001. Conflict
- 6 monitoring and cognitive control. Psychol Rev 108, 624–652.
- 7 Butters, M.A., Whyte, E.M., Nebes, R.D., Begley, A.E., Dew, M.A., Mulsant, B.H., Zmuda,
- 8 M.D., Bhalla, R., Meltzer, C.C., Pollock, B.G., Reynolds, C.F., Becker, J.T., 2004. The nature
- 9 and determinants of neuropsychological functioning in late-life depression. Arch. Gen.
- 10 Psychiatry 61, 587–595. https://doi.org/10.1001/archpsyc.61.6.587
- 11 Cyders, M.A., Coskunpinar, A., 2011. Measurement of constructs using self-report and
- 12 behavioral lab tasks: is there overlap in nomothetic span and construct representation for
- 13 impulsivity? Clin Psychol Rev 31, 965–982. https://doi.org/10.1016/j.cpr.2011.06.001
- 14 d'Acremont, M., Van der Linden, M., 2007. How is impulsivity related to depression in
- 15 adolescence? Evidence from a French validation of the cognitive emotion regulation
- 16 questionnaire. J Adolesc 30, 271–282. https://doi.org/10.1016/j.adolescence.2006.02.007
- 17 D Spielberger, C., Gorsuch, R., E Lushene, R., Vagg, P., A Jacobs, G., 1983. Manual for the
- **18** State-Trait Anxiety Inventory (Form Y1 Y2).
- 19 Dautzenberg, G., Lijmer, J., Beekman, A., 2021. Clinical value of the Montreal Cognitive
- 20 Assessment (MoCA) in patients suspected of cognitive impairment in old age psychiatry.
- 21 Using the MoCA for triaging to a memory clinic. Cognitive Neuropsychiatry 26, 1–17.
- 22 https://doi.org/10.1080/13546805.2020.1850434
- 23 Delphin-Combe, F., Bathsavanis, A., Rouch, I., Liles, T., Vannier-Nitenberg, C., Fantino, B.,
- 24 Dauphinot, V., Krolak-Salmon, P., 2016. Relationship between anxiety and cognitive
- performance in an elderly population with a cognitive complaint. European Journal of
 Neurology 23, 1210, 1217, https://doi.org/10.1111/ene.13004
- 26 Neurology 23, 1210–1217. https://doi.org/10.1111/ene.13004
- 27 Dillon, D.G., Wiecki, T., Pechtel, P., Webb, C., Goer, F., Murray, L., Trivedi, M., Fava, M.,
- 28 McGrath, P.J., Weissman, M., Parsey, R., Kurian, B., Adams, P., Carmody, T., Weyandt, S.,
- 29 Shores-Wilson, K., Toups, M., McInnis, M., Oquendo, M.A., Cusin, C., Deldin, P., Bruder,
- 30 G., Pizzagalli, D.A., 2015. A Computational Analysis of Flanker Interference in Depression.
- 31 Psychol Med 45, 2333–2344. https://doi.org/10.1017/S0033291715000276
- 32 Diniz, B.S., Butters, M.A., Albert, S.M., Dew, M.A., Reynolds, C.F., 2013. Late-life
- 33 depression and risk of vascular dementia and Alzheimer's disease: systematic review and
- 34 meta-analysis of community-based cohort studies. Br J Psychiatry 202, 329–335.
- 35 https://doi.org/10.1192/bjp.bp.112.118307
- 36 Disner, S.G., Beevers, C.G., Haigh, E.A.P., Beck, A.T., 2011. Neural mechanisms of the
- 37 cognitive model of depression. Nat. Rev. Neurosci. 12, 467–477.
- 38 https://doi.org/10.1038/nrn3027
- 39 Dombrovski, A.Y., Szanto, K., Siegle, G.J., Wallace, M.L., Forman, S.D., Sahakian, B.,
- 40 Reynolds, C.F., Clark, L., 2011. Lethal Forethought: Delayed Reward Discounting
- 41 Differentiates High- and Low-Lethality Suicide Attempts in Old Age. Biol Psychiatry 70,
- 42 138–144. https://doi.org/10.1016/j.biopsych.2010.12.025
- 43 Dreisbach, G., Fischer, R., 2015. Conflicts as Aversive Signals for Control Adaptation. Curr
- 44 Dir Psychol Sci 24, 255–260. https://doi.org/10.1177/0963721415569569
- 45 Duthoo, W., Abrahamse, E.L., Braem, S., Boehler, C.N., Notebaert, W., 2014. The
- 46 heterogeneous world of congruency sequence effects: an update. Front Psychol 5.
- 47 https://doi.org/10.3389/fpsyg.2014.01001
- 48 Engström, J., Markkula, G., Victor, T., Merat, N., 2017. Effects of Cognitive Load on Driving
- 49 Performance: The Cognitive Control Hypothesis. Hum Factors 59, 734–764.
- 50 https://doi.org/10.1177/0018720817690639

- 1 Eshel, N., Roiser, J.P., 2010. Reward and Punishment Processing in Depression. Biological
- 2 Psychiatry, Vascular Function in Depression in Older Adults 68, 118–124.
- 3 https://doi.org/10.1016/j.biopsych.2010.01.027
- 4 Feil, D., Razani, J., Boone, K., Lesser, I., 2003. Apathy and cognitive performance in older
- 5 adults with depression. Int J Geriatr Psychiatry 18, 479–485. https://doi.org/10.1002/gps.869
- 6 Fox, J., Weisberg, S., 2011. An R Companion to Applied Regression. SAGE Publications.
- 7 Funes, C.M., Lavretsky, H., Ercoli, L., St Cyr, N., Siddarth, P., 2018. Apathy Mediates
- 8 Cognitive Difficulties in Geriatric Depression. Am J Geriatr Psychiatry 26, 100–106.
- 9 https://doi.org/10.1016/j.jagp.2017.06.012
- 10 Golden, C., 1978. A Manual for the Clinical and Experimental Use of the Stroop Color and
- 11 Word Test. Faculty Books and Book Chapters.
- 12 Grahek, I., Shenhav, A., Musslick, S., Krebs, R.M., Koster, E.H.W., 2019. Motivation and
- 13 cognitive control in depression. Neurosci Biobehav Rev 102, 371–381.
- 14 https://doi.org/10.1016/j.neubiorev.2019.04.011
- 15 Gratton, G., Coles, M.G., Donchin, E., 1992. Optimizing the use of information: strategic
- 16 control of activation of responses. J Exp Psychol Gen 121, 480–506.
- Hamilton, M., 1960. A rating scale for depression. J. Neurol. Neurosurg. Psychiatry 23, 56–
 62.
- 19 Hasbroucq, T., Guiard, Y., 1991. Stimulus-response compatibility and the Simon effect:
- 20 Toward a conceptual clarification. Journal of Experimental Psychology: Human Perception
- 21 and Performance 17, 246–266. https://doi.org/10.1037/0096-1523.17.1.246
- Hedge, A., Marsh, N.W., 1975. The effect of irrelevant spatial correspondences on two-choice
- 23 response-time. Acta Psychol (Amst) 39, 427–439. https://doi.org/10.1016/0001-
- 24 6918(75)90041-4
- 25 Holmes, A.J., Pizzagalli, D.A., 2007. Task feedback effects on conflict monitoring and
- 26 executive control: Relationship to subclinical measures of depression. Emotion 7, 68–76.
- 27 https://doi.org/10.1037/1528-3542.7.1.68
- Hommel, B., 2011. The Simon effect as tool and heuristic. Acta Psychol (Amst) 136, 189–
- 29 202. https://doi.org/10.1016/j.actpsy.2010.04.011
- 30 Hothorn, T., Bretz, F., Westfall, P., 2008. Simultaneous inference in general parametric
- 31 models. Biom J 50, 346–363. https://doi.org/10.1002/bimj.200810425
- 32 Hox, J.J., Moerbeek, M., Schoot, R. van de, Moerbeek, M., Schoot, R. van de, 2017.
- 33 Multilevel Analysis : Techniques and Applications, Third Edition. Routledge.
- 34 https://doi.org/10.4324/9781315650982
- 35 Katz, R., De Sanctis, P., Mahoney, J.R., Sehatpour, P., Murphy, C.F., Gomez-Ramirez, M.,
- 36 Alexopoulos, G.S., Foxe, J.J., 2010. Cognitive control in late-life depression: response
- 37 inhibition deficits and dysfunction of the anterior cingulate cortex. Am J Geriatr Psychiatry
- 38 18, 1017–1025. https://doi.org/10.1097/JGP.0b013e3181d695f2
- 39 Keith Conners, C., Sitarenios, G., Ayearst, L.E., 2018. Conners' Continuous Performance
- 40 Test Third Edition, in: Kreutzer, J.S., DeLuca, J., Caplan, B. (Eds.), Encyclopedia of Clinical
- 41 Neuropsychology. Springer International Publishing, Cham, pp. 929–933.
- 42 https://doi.org/10.1007/978-3-319-57111-9_1535
- 43 König, N., Steber, S., Borowski, A., Bliem, H.R., Rossi, S., 2021. Neural Processing of
- 44 Cognitive Control in an Emotionally Neutral Context in Anxiety Patients. Brain Sci 11, 543.
- 45 https://doi.org/10.3390/brainsci11050543
- 46 Kornblum, S., Lee, J.-W., 1995. Stimulus-response compatibility with relevant and irrelevant
- 47 stimulus dimensions that do and do not overlap with the response. Journal of Experimental
- 48 Psychology: Human Perception and Performance 21, 855–875. https://doi.org/10.1037/0096-1523 21 4 855
- 49 1523.21.4.855
- 50 Li, Q., Nan, W., Wang, K., Liu, X., 2014. Independent Processing of Stimulus-Stimulus and

- 1 Stimulus-Response Conflicts. PLOS ONE 9, e89249.
- 2 https://doi.org/10.1371/journal.pone.0089249
- 3 Maldonado, T., Orr, J.M., Goen, J.R.M., Bernard, J.A., 2020. Age Differences in the
- 4 Subcomponents of Executive Functioning. J Gerontol B Psychol Sci Soc Sci 75, e31–e55.
- 5 https://doi.org/10.1093/geronb/gbaa005
- 6 Marin, R.S., 1991. Apathy: a neuropsychiatric syndrome. J Neuropsychiatry Clin Neurosci 3,
- 7 243–254. https://doi.org/10.1176/jnp.3.3.243
- 8 McGirr, A., Dombrovski, A.Y., Butters, M.A., Clark, L., Szanto, K., 2012. Deterministic
- 9 learning and attempted suicide among older depressed individuals: Cognitive assessment
- 10 using the Wisconsin Card Sorting Task. Journal of Psychiatric Research 46, 226–232.
- 11 https://doi.org/10.1016/j.jpsychires.2011.10.001
- 12 Meiran, N., Diamond, G.M., Toder, D., Nemets, B., 2011. Cognitive rigidity in unipolar
- 13 depression and obsessive compulsive disorder: examination of task switching, Stroop,
- 14 working memory updating and post-conflict adaptation. Psychiatry Res 185, 149–156.
- 15 https://doi.org/10.1016/j.psychres.2010.04.044
- 16 Nakagawa, S., Schielzeth, H., 2013. A general and simple method for obtaining R2 from
- 17 generalized linear mixed-effects models. Methods in Ecology and Evolution 4, 133–142.
- 18 https://doi.org/10.1111/j.2041-210x.2012.00261.x@10.1111/(ISSN)2041-
- 19 210X.TOPMETHODS
- 20 Nasreddine, Z.S., Phillips, N.A., Bédirian, V., Charbonneau, S., Whitehead, V., Collin, I.,
- 21 Cummings, J.L., Chertkow, H., 2005. The Montreal Cognitive Assessment, MoCA: a brief
- screening tool for mild cognitive impairment. J Am Geriatr Soc 53, 695–699.
- 23 https://doi.org/10.1111/j.1532-5415.2005.53221.x
- 24 Nebes, R.D., Butters, M.A., Mulsant, B.H., Pollock, B.G., Zmuda, M.D., Houck, P.R.,
- 25 Reynolds, C.F., 2000. Decreased working memory and processing speed mediate cognitive
- 26 impairment in geriatric depression. Psychol Med 30, 679–691.
- 27 https://doi.org/10.1017/s0033291799001968
- 28 Nelson, H.E., 1976. A modified card sorting test sensitive to frontal lobe defects. Cortex 12,
- 29 313–324. https://doi.org/10.1016/s0010-9452(76)80035-4
- Ng, J., Chan, H.Y., Schlaghecken, F., 2012. Dissociating effects of subclinical anxiety and
- depression on cognitive control. Adv Cogn Psychol 8, 38–49. https://doi.org/10.2478/v10053008-0100-6
- 33 Nicoletti, R., Umiltà, C., 1994. Attention shifts produce spatial stimulus codes. Psychol. Res
- 34 56, 144–150. https://doi.org/10.1007/BF00419701
- 35 Pinheiro, J., Bates, D., DebRoy, S.S., Sarkar, D., 2013. Nlme: Linear and Nonlinear Mixed
- 36 Effects Models. R package version 31-110 3, 1–113.
- 37 Quinn, C.R., Harris, A., Kemp, A.H., 2012. The impact of depression heterogeneity on
- 38 inhibitory control. Aust N Z J Psychiatry 46, 374–383.
- 39 https://doi.org/10.1177/0004867411432073
- 40 Reitan, R.M., 1958. Validity of the Trail Making Test as an Indicator of Organic Brain
- 41 Damage. Percept Mot Skills 8, 271–276. https://doi.org/10.2466/pms.1958.8.3.271
- 42 Rey-Mermet, A., Gade, M., 2018. Inhibition in aging: What is preserved? What declines? A
- 43 meta-analysis. Psychon Bull Rev 25, 1695–1716. https://doi.org/10.3758/s13423-017-1384-7
- 44 Richard Ridderinkhof, K., Forstmann, B.U., Wylie, S.A., Burle, B., van den Wildenberg,
- 45 W.P.M., 2011. Neurocognitive mechanisms of action control: resisting the call of the Sirens.
- 46 Wiley Interdiscip Rev Cogn Sci 2, 174–192. https://doi.org/10.1002/wcs.99
- 47 Richard-Devantoy, S., Annweiler, C., Le Gall, D., Garré, J.-B., Olié, J.-P., Beauchet, O.,
- 48 2011. Cognitive inhibition in suicidal depressed elderly: a case-control pilot study. J Clin
- 49 Psychiatry 72, 871–872. https://doi.org/10.4088/JCP.10106797
- 50 Richard-Devantoy, S., Szanto, K., Butters, M.A., Kalkus, J., Dombrovski, A.Y., 2015.

- 1 Cognitive inhibition in older high-lethality suicide attempters. Int J Geriatr Psychiatry 30,
- 2 274–283. https://doi.org/10.1002/gps.4138
- 3 Ridderinkhof, K. Richard, 2002. Micro- and macro-adjustments of task set: activation and
- 4 suppression in conflict tasks. Psychol Res 66, 312–323. https://doi.org/10.1007/s00426-002 5 0104-7
- 6 Ridderinkhof, K. R., 2002. Activation and suppression in conflict tasks: Empirical
- 7 clarification through distributional analyses.
- 8 Ruchsow, M., Groen, G., Kiefer, M., Beschoner, P., Hermle, L., Ebert, D., Falkenstein, M.,
- 9 2008. Electrophysiological evidence for reduced inhibitory control in depressed patients in
- 10 partial remission: a Go/Nogo study. Int J Psychophysiol 68, 209–218.
- 11 https://doi.org/10.1016/j.ijpsycho.2008.01.010
- 12 Seblova, D., Berggren, R., Lövdén, M., 2020. Education and age-related decline in cognitive
- 13 performance: Systematic review and meta-analysis of longitudinal cohort studies. Ageing
- 14 Res. Rev. 58, 101005. https://doi.org/10.1016/j.arr.2019.101005
- 15 Sheline, Y.I., Barch, D.M., Garcia, K., Gersing, K., Pieper, C., Welsh-Bohmer, K., Steffens,
- 16 D.C., Doraiswamy, P.M., 2006. Cognitive function in late life depression: relationships to
- 17 depression severity, cerebrovascular risk factors and processing speed. Biol. Psychiatry 60,
- 18 58–65. https://doi.org/10.1016/j.biopsych.2005.09.019
- 19 Shenhav, A., Botvinick, M.M., Cohen, J.D., 2013. The expected value of control: an
- 20 integrative theory of anterior cingulate cortex function. Neuron 79, 217–240.
- 21 https://doi.org/10.1016/j.neuron.2013.07.007
- 22 Snyder, H.R., 2013. Major Depressive Disorder is Associated with Broad Impairments on
- 23 Neuropsychological Measures of Executive Function: A Meta-Analysis and Review. Psychol
- 24 Bull 139, 81–132. https://doi.org/10.1037/a0028727
- 25 Stahl, C., Voss, A., Schmitz, F., Nuszbaum, M., Tüscher, O., Lieb, K., Klauer, K.C., 2014.
- 26 Behavioral components of impulsivity. J Exp Psychol Gen 143, 850–886.
- 27 https://doi.org/10.1037/a0033981
- 28 Vallesi, A., Canalaz, F., Balestrieri, M., Brambilla, P., 2015. Modulating speed-accuracy
- strategies in major depression. Journal of Psychiatric Research 60, 103–108.
- 30 https://doi.org/10.1016/j.jpsychires.2014.09.017
- 31 van Steenbergen, H., Booij, L., Band, G.P.H., Hommel, B., van der Does, A.J.W., 2012.
- 32 Affective regulation of cognitive-control adjustments in remitted depressive patients after
- acute tryptophan depletion. Cogn Affect Behav Neurosci 12, 280–286.
- 34 https://doi.org/10.3758/s13415-011-0078-2
- 35 Whiteside, S.P., Lynam, D.R., 2001. The five factor model and impulsivity: Using a structural
- 36 model of personality to understand impulsivity. Personality and Individual Differences 30,
- 37 669–689. https://doi.org/10.1016/S0191-8869(00)00064-7
- 38 Widlöcher, D.J., 1983. Psychomotor Retardation: Clinical, Theoretical, and Psychometric
- 39 Aspects. Psychiatric Clinics 6, 27–40. https://doi.org/10.1016/S0193-953X(18)30838-4
- 40 World Health Organization, Department of Ageing and Life Course, 2017. Integrated care for
- 41 older people: guidelines on community-level interventions to manage declines in intrinsic
- 42 capacity.
- 43 Wylie, S.A., Ridderinkhof, K.R., Bashore, T.R., van den Wildenberg, W.P.M., 2010. The
- 44 Effect of Parkinson's Disease on the Dynamics of On-line and Proactive Cognitive Control
- 45 during Action Selection. J Cogn Neurosci 22, 2058–2073.
- 46 https://doi.org/10.1162/jocn.2009.21326
- 47 Zhang, B.-W., Xu, J., Chang, Y., 2016. The Effect of Aging in Inhibitory Control of Major
- 48 Depressive Disorder Revealed by Event-Related Potentials. Front. Hum. Neurosci. 10.
- 49 https://doi.org/10.3389/fnhum.2016.00116
- 50 Zimmerman, M., Martinez, J.H., Young, D., Chelminski, I., Dalrymple, K., 2013. Severity

- classification on the Hamilton Depression Rating Scale. J Affect Disord 150, 384–388. https://doi.org/10.1016/j.jad.2013.04.028
- 1 2 3















