ORIGINAL ARTICLE

Cognitive control of feature bindings: evidence from children with autistic spectrum disorder

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Received: 9 February 2011 / Accepted: 23 November 2011 © The Author(s) 2011. This article is published with open access at Springerlink.com

Abstract Understanding how the brain integrates features from different domains that are processed in distinct cortical regions calls for the examination of integration processes. Recent studies of feature-repetition effects demonstrated interactions across perceptual features and action-related features: repeating only some features of the perception–action episode hinders performance. These partial-repetition costs point to the existence of temporary memory traces (event files). However, the principles and the constraints that govern the management of such traces are still unclear. Here, we investigated whether children with autistic spectrum disorder (ASD) differ from typically developing children in managing episodic memory traces. The results show that both groups integrate stimulus features along with action features, but children with ASD exhibit larger partial-repetition costs, suggesting lesser control and flexibility in updating episodic memory traces. The findings are discussed in the light of evidence for a central role of the dopaminergic system in cognitive integration, ASD, and cognitive control.

Introduction

Given that the primate cortex processes the various features of perceptual events and actions in distinct brain regions (e.g., DeYoe & van Essen, 1988), it has been assumed that

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Published online: 06 December 2011

is easier after having paired A and X, or the unrelated object B and action Y, than after having responded differently to the same object $(A \rightarrow Y)$ or responded similarly to a different object $(B \rightarrow X)$. Apparently, then, a single pairing of a stimulus (feature) and a response is sufficient to create an episodic binding (an event file; Hommel, 1998) that interferes with partially, but not completely overlapping bindings. This suggests that repeating at least one (stimulus and/or response) feature leads to the retrieval of the just created binding, which interferes with current processing if that involves the reactivation of a previous (no longer valid)

repetitions or alternations).

the representations of these features need to be integrated into coherent episodic bindings (e.g., Hommel, 2004; Kahneman, Treisman, & Gibbs, 1992). Evidence for the spontaneous integration of perceptual features comes from analyses of (interactions between) repetition effects. For instance, people not only respond faster to letters that they just saw in a preview display (a standard priming or repetition effect), but they are particularly fast if the repeated letter also appears in the same location (Kahneman et al., 1992). This suggests that processing a perceptual event induces the binding of the codes of its features, so that repeating the particular conjunction of features allows for particularly efficient processing.

Comparable observations have been made for auditory

features (Mondor, Hurlburt, & Thorne, 2003; Zmigrod &

Hommel, 2009), perceptual features from different sensory

modalities (Zmigrod, Spapé, & Hommel, 2009), and for

perceptual and action features (Hommel, 1998). For

instance, responding to object A by carrying out response X

feature code (Hommel, 2004). This produces costs which we refer to as partial-repetition costs (i.e., the performance deficits with incomplete repetitions of stimulus-feature or stimulus-response combinations as compared to complete



Partial-repetition costs can be observed if two conditions are met: feature codes need to be integrated in the respective prime trial (a process reflecting binding efficiency); and this created binding needs to be retrieved in the present (probe) trial (a process reflecting the efficiency to control episodic bindings). Interestingly, attempts to dissociate these two processes provided evidence that the binding process proper is more or less automatic (Hommel, 2005). For instance, the degree and strength of binding two given features is independent from the frequency and probability of their co-occurrence (Colzato, Raffone, & Hommel, 2006; Hommel & Colzato, 2009) and the availability of attentional resources (Hommel, 2005; Takegata et al., 2005). Nevertheless, bindings involving task-relevant features have a stronger impact on behavior (Hommel, 1998, 2007) and can persist longer (Zmigrod & Hommel, 2010). Presumably, they are more likely to be retrieved (and/or created) as well as decay slower than irrelevant features. Furthermore, the retrieval process seems to be affected by task instructions and individual differences. For instance, partial-repetition costs have been found to be *more* pronounced in populations that are unlikely to possess particularly well-functioning binding processes, such as individuals with low fluid intelligence (Colzato, van Wouwe, Lavender, & Hommel, 2006) and in young children and elderly participants (Hommel, Kray, & Lindenberger, 2011). This suggests at least some degree of control, assumingly during episodic retrieval. Thus, Given that executivecontrol functions are related to fluid intelligence (Duncan et al., 2000), not fully developed in young age (Hongwanishkul, Happaney, Lee, & Zelazo, 2005), and impaired in old age (Fisk & Sharp, 2004), these observations suggest that people differ with respect to the efficiency of managing episodic representations. More specifically, it makes sense to assume that individuals with more efficient executive-control functions like people high in fluid intelligence and younger adults are better able to either inhibit no longer relevant bindings or update those bindings to represent current feature combinations (Colzato, Raffone, et al., 2006).

The possibility that individual differences relate mainly to control over memory traces is also consistent with recent findings of Keizer, Verment, and Hommel (2010), whose participants received neurofeedback to increase cortical gamma synchronization. Such training improved memory retrieval of relevant information in a standard recollection task and reduced partial-repetition costs due to irrelevant bindings. Convergent evidence comes from a study of Hommel, Fischer, Colzato, van den Wildenberg, and Cellini (2011, in press). Consistent with Kofman, Meiran, Greenberg, Balas, and Cohen (2006), Hommel et al. observed that inducing mild stress improved task-switching performance—an indicator of executive-control functioning. Interestingly, the same manipulation also reduced partial repetition costs, suggesting more effective control of

episodic retrieval. Thus, although the relationship between binding of stimulus and response features (as in the *event-file* task) and control processes is not entirely clear, studies using this task demonstrated that the partial-repetition costs are affected in populations that are impaired with respect to cognitive control or in various situations that associated with lack of cognitive control (Colzato, van Wouwe, et al., 2006; Colzato, Kool, & Hommel, 2008; Hommel et al., 2011; 2011, in press). Given that performing the event-file task does require, or benefit from, the retrieval of previous episodes, efficient control would result in the prevention of retrieval in the first place or, if that is not possible, in the inhibition of retrieved bindings.

In view of these hints to a link between executive control functions and the management of episodic event files, we considered binding-task performance in individuals suffering from autistic spectrum disorder (ASD)—a disorder that has been related to deficits in both episodic binding (Frith, 2003) and cognitive control (Solomon et al., 2009)—particularly interesting. ASD is one of the most common childhood disorders and characterized by social communication impairment, deficits in language skills, and repetitive behaviors. Various authors have advocated various factors that might account for the disorder, but it is fair to say that there is a rather general agreement that impairments related to executive control functions play a major role (Corbett, Constantine, Hendren, Rocke, & Ozonoff, 2009; Hill, 2004; Kenworthy, Black, Harrison, Della Rosa, & Wallace, 2009). Among other things, these impairments are assumed to render ASD patients cognitively less flexible, which would account for both impaired performance in clinical tests like the Wisconsin card sorting task (WCST: Willcutt, Sonuga-Barke, Nigg, & Sergeant, 2008) and behavioral rigidity in everyday life behavior.

Unfortunately, however, experimental evidence supporting the link between ASD and cognitive flexibility is still scarce and equivocal (Geurts, Corbett, & Solomon, 2009; Poljac et al., 2010). As suggested by Geurts et al., this might be due to the fact that most clinical tests are rather complex and unlikely to provide process-pure measures of the cognitive processes of interest. For instance, the WCST relies on a good understanding of the task, working memory, learning from feedback, the availability of multiple strategies, and so on. Not all of these abilities and skills are related to the processes targeted by executive-control and flexibility accounts of ASD, so that the outcomes on these kinds of tests are open to various interpretations. Clearly, there is a need for more diagnostic experimental tasks that provide more process-pure measures of cognitive control and flexibility.

In the present study, we investigated whether ASD is associated with impairments in managing episodic event files. More specifically, we were interested to see whether ASD patients are impaired in controlling and flexibly switching between stimulus–response bindings. This would



fit with the observation of specific deficits in children suffering from ASD in tasks requiring cognitive flexibility, such as Intradimensional/Extradimensional (ID/ED) shift task of the Cambridge Neuropsychological Test Automated Battery (CANTAB) (Ozonoff et al., 2004), or the WCST (for a review, see Hill, 2004). If ASD patients have a deficit in integrating stimulus and/or response features—as some authors have claimed (e.g., Frith, 2003), one would expect ASD patients to show small or no partial-repetition costs. However, if ASD would indeed be associated with poorer event-file management abilities, one would expect ASD patients to show more pronounced partial-repetition costs than control participants do. We tested our hypothesis by comparing the performance of a group of ASD-diagnosed children and a group of normally developing children in a standard event-file task (e.g., Hommel, 1998), which was only slightly adapted for use with children.

Method

Participants

The participants in this study included 40 children of age between 11 and 18 years : 20 children (13 males) with autism spectrum disorder (ASD) (age 11–18, mean 15, SD 2.02), and 20 (14 males) typically developing children (the control group; age 11–18, mean 15, SD 2.01).

The children with ASD were selected from consecutive referrals to the outpatient and inpatient Department of Child and Adolescent Psychiatry at the University Medical Centre of Utrecht, the Netherlands. In order to qualify for the current study, the ASD participants had to meet the following three criteria: First, a diagnosis of ASD was required. Two certified experienced child psychiatrists diagnosed these participants and needed to reach a consensus on diagnosis. They used DSM-IV criteria for autistic disorder (Diagnostic and Statistical Manual of Mental Disorder, Fourth Edition; American Psychiatric Association, 1994) to evaluate the developmental history of the child as well as current social, emotional, and adaptive functioning. Furthermore, medical records and structured questionnaires completed by the child's parents and teacher(s) were included in the diagnostic process (Child Behavior Checklist: CBCL; and Teacher's Report Form: TRF; Achenbach & Edelbrock, 1986). Both child psychiatrists observed and interviewed the child in a semi-structured playroom session according to a DSM checklist made available for the Dutch situation. Second, Full-Scale IQs (FSIQ) were required to be 70 or above (Mean IQ 101; SD 9.71), as measured with the Dutch adaptation of the Wechsler Intelligence Scale for Children (Wechsler, 1997). Third, all participants were to have normal or corrected-to-normal vision.

In the healthy controls we included children with the same age range and IQ (there was no significant difference IQ between the groups as measured by t-test, t(38) = 2.011; p = .051), but without any known neurological or developmental disorders. Their IQ was above 70 (Mean IQ 107; SD 8.81) as measured by the Dutch adaptation of the Wechsler Intelligence Scale for Children (Wechsler, 1997), with normal or corrected-to-normal vision.

Finally, the study was conducted in accordance with the declaration of Helsinki and guidelines of the local ethics committee. All parents signed a written consent before participating in the study.

Procedure and design

The event file task measures binding-related effects, which can be quantified by calculating partial-repetition costs related to combinations of stimulus features (shape and color in our case) and combinations of stimulus features and the response. One of the perceptual features was task-relevant feature (shape) and the other (color) was the irrelevant feature for the task. To manipulate the repetition versus alternation of stimulus features and responses, the task comprises of pairs of trials with a prime trial (S1 \rightarrow R1) followed by a probe trial (S2 \rightarrow R2), see Fig. 1. The probe trial required a manual binary-choice response (R2) to the shape of the second stimulus S2 (an apple or a banana). The prime trial required a manual response (R1) to the mere onset of the first stimulus (S1). The correct R1 was signaled in advance of S1 (through a left- or right-pointing arrowhead), so that S1 and R1 could be varied independently, which was necessary to create orthogonal repetitions and alternations of stimulus shape and response. As an additional stimulus feature, color was also varied by presenting the apple or banana in green or yellow (see Colzato, van Wouwe, et al., 2006). Stimulus color could repeat or alternate independently of stimulus shape and responses, thus creating a $2 \times 2 \times 2$ -factorial design.

Each trial consisted of a prime-probe pair. The experiment was composed of a practice block with ten practice

¹ Partial-repetition costs for a given interaction between factors X and Y were calculated as the difference between the RTs/PEs for partial repetitions (feature X repeated and feature Y alternated, or vice versa) and the RTs/PEs for complete repetitions and "complete" alternations. E.g., the partial repetition costs in RTs for the shape X response interaction at a given group would be PRCshapeXresponse = (RT shape repeated/response alternated + RT shape alternated/response repeated)/2 - (RT shape repeated/response repeated + RT shape alternated/response alternated)/2. Partial repetition costs thus correspond to the 2-way interaction term of the respective features (and are thus immune to possible, but theoretically less relevant, main effects of feature repetition); a value close to zero means that the repetition effects of the two given features do not interact; a value greater than zero indicates a "binding-type" interaction of the sort described in the text.



Fig. 1 Sequence of events in the event file task. A visual response cue signaled a left or right response (R1) that was to be delayed until presentation of the first stimulus S1 (S1 is used as a detection signal for R1). The second stimulus S2 appeared 1,000 ms after S1. S2 signaled R2, a speeded left or right response according to the shape

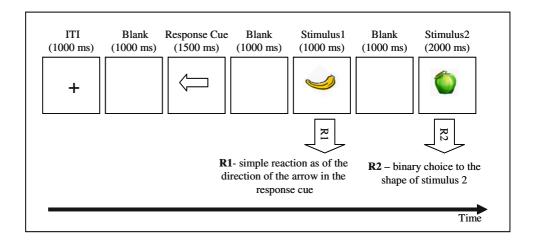


Table 1 Means of mean reaction times for responses to stimulus 2 (RT_{R2} in ms) and SE in parenthesis, as a function of group (ASD children vs. control—typically developing children), the relationship be-

tween the responses (R1 and R2), and the relationship between the stimuli features (S1 and S2) for shape and color

Group	Response repeated		Response alternated		Partial repetition
	Shape repeated	Shape alternated	Shape repeated	Shape alternated	costs
RTs (ms)					
ASD	518 (20)	649 (24)	591 (24)	549 (24)	86
Control	576 (20)	635 (24)	616 (24)	568 (24)	53
Errors (%)					
ASD	5.7 (0.8)	18.1 (1.9)	14.5 (1.6)	4.5 (0.8)	11.2
Control	1.3 (0.8)	6.1 (1.9)	9.6 (1.6)	1.6 (0.8)	6.4
Group	Response repeated		Response alternated		Partial repetition
	Color repeated	Color alternated	Color repeated	Color alternated	costs
RTs (ms)					
ASD	580 (22)	587 (21)	575 (24)	566 (23)	8
Control	600 (22)	611 (21)	592 (24)	592 (23)	5
Errors (%)					
ASD	11 4 (1 2)	12.4 (1.4)	10.2 (1.2)	8.7 (1.2)	1.2
ASD	11.4 (1.3)	12.4 (1.4)	10.2 (1.2)	0.7 (1.2)	1.2
Control	3.6 (1.3)	3.8 (1.4)	7.3 (1.2)	4.0 (1.2)	1.8

The rightmost column gives the partial repetition costs (see footnote 1), which differed significantly in response-shape between the two groups, p < .005, both in reaction times and error rates

trials, which were not further analyzed, and an experimental block with 196 experimental trials. The order of the trials was randomized but all eight conditions appeared equally often. Half of the participants responded to the apple and the banana by pressing on the left and right key press, respectively, while the other half received the opposite mapping. The participants were asked to respond as quickly and accurately as possible.

Results

Trials with incorrect R1 responses (0.75%), as well as missing (RT > 1,500 ms) or anticipatory (RT < 100 ms) R2

responses (0.02%) were excluded from analysis. From the remaining data, mean reaction time (RTs) and percentage of errors (PEs) for R2 (see Table 1) were analyzed as a function of the four variables: the relationship between S1 and S2 (repetition vs. alternation) with regard to shape and to color, and the relationship between responses R1 and R2 (repetition vs. alternation), which all varied within participants, and group (ASD vs. control). Mixed-design ANO-VAs were performed with repeated measures on three variables and with group as between-participant variable.

ASD patients tended to be faster than control participants (RTs: 576 vs. 611 ms), F(1,38) = .852, p = .362, but were less accurate (PEs: 9.57 vs. 4.3%), F(1,38) = 14.42, p < .001, suggesting a somewhat different speed-accuracy



strategy. There was a significant main effect of shape repetition in RTs, F(1,38) = 13.54, p < .001, due to faster responses to repeated (581 ms) than alternated shapes (605 ms). This effect was modified by group, F(1,38) = 5.20, p < .05; separate ANOVAs, split by group, revealed that it was statistically significant only in the ASD group, F(1,19) = 13.207, p < .005, where responses were slower to shape alternation (596 ms) than to shape repetition (556 ms), indicating slower adaptation to alternations. These results are in line with studies showing that ASD children demonstrate deficits in adaptive skills (e.g., Kanne et al., 2011; Paul et al., 2004) and repetitive behaviors (South, Ozonoff, & Mcmahon, 2007).

There were significant interactions between shape repetition and response repetition in RTs, F(1,38) = 113.77, p < .0001; and PEs, F(1,38) = 61.08, p < .0001. These findings followed the common pattern with worse performance if only one of the features (shape or response) is repeated while the other is not, as compared with complete repetitions or alternations (see Hommel, 1998). In addition, there was a significant interaction between color, the irrelevant feature, and the response in PEs, F(1,38) = 9.87, p < .005, showing that the response was more accurate for total repetition or total alternation of the response and the color than partial repetition (see Table 1).

More importantly for our study, the response–shape interaction was further modified by group in both RTs, F(1,38) = 4.478, p < .05; and PEs, F(1,38) = 6.82, p < .05 (see Fig. 2). This was due to more pronounced interactions in children with ASD than in typically developing children. In contrast, group was not involved in either the three-way interactions with shape and color or with color and response, all Fs(1,38) < 1, or the four-way interaction, F(1,38) = 0.51, p = .48, and F(1,38) = 2.65, p = .11, for RTs and PEs, respectively, indicating that the impact of group is restricted to effects of features from relevant dimensions.

To further analyze the response–shape interaction, we calculated partial repetition costs (see footnote 1) for RTs and PEs and ran an ANOVA with group type. There were reliable effects of group in RTs: F(1,38) = 4.47 p < .05, and PEs: F(1,38) = 6.83, p < .05, indicating higher costs associated with shape–response integration for ASD than the control group (see Fig. 3).

As the experiment was relatively long, participants could develop particular strategies to deal with the task, especially with its relevant aspects (i.e., shape and response features) which might have changed the respective effects over time. To test this possibility and to see whether it might have emerged in the two groups differently, we split the data into a first and second half, calculated (response-shape) partial repetition costs for the two practice levels for each group in RTs and PEs, and ran an ANOVA with practice level and group type as factors. However, there were no

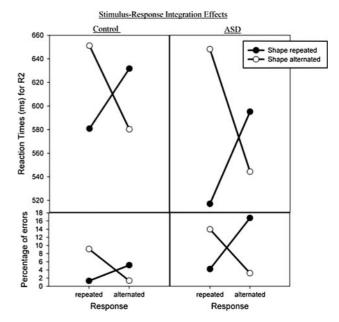


Fig. 2 Mean reaction times and error percentages (on R2) for repetition versus alternation of the shape as a function of response repetition versus alternation and the group

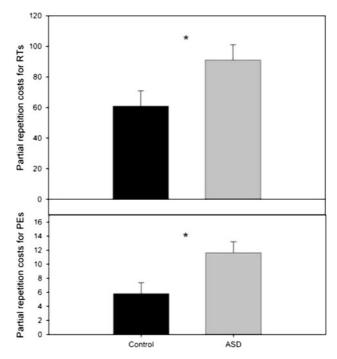


Fig. 3 Effects indicating shape-response partial repetition costs (footnote 1) in reaction times and error rates (on R2), for the ASD and control groups. *p < .005

significant main effects of practice level in RTs and PEs (Fs < 1) or interactions between group and practice level (Fs < 1). This does not provide evidence that all or some of the participants developed particular strategies to deal with the task over time, at least not in a systematic fashion.

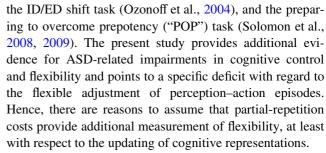


Discussion

The aim of this study was to investigate whether children suffering from ASD would show a specific deficit in a task tapping into the handling of episodic event files, that is, of bindings between codes that represent the features of experienced objects and stimulus-response episodes (Hommel, 1998, 2004). Both normally developing controls and the ASD group showed partial-repetition costs for combinations of the two task-relevant stimulus and response features: stimulus shape and response location. It is known that task relevance modulates feature-integration effects (e.g., Hommel, 1998; Hommel & Colzato, 2004) so that it is not surprising that the reliable effects were mainly restricted to the features that mattered for the task. More interesting, however, is that these effects were observed in both groups. Given that partial-repetition costs in S2–R2 performance can only occur if the respective features are integrated when processing S1 and R1, this observation implies that binding as such does not seem to be impaired in ASD. This does not seem to fit with the claim that ASD is associated with difficulties in feature integration (e.g., Frith, 2003).

Importantly, the aftereffects of binding were larger but not smaller in the ASD group (see Fig. 3), suggesting that ASD children are impaired in managing of episodic representations. That is, both healthy controls and children suffering from ASD seem to spontaneously integrate stimulus-response episodes and automatically retrieve traces of these episodes when facing a similar, that is, feature-overlapping episode thereafter (see Fig. 2). However, healthy controls seem to be more efficient in preventing these traces from affecting ongoing processes and/or updating the traces if they do not fit with the current feature combinations, reflecting a specific deficit in cognitive control in ASD. We found no evidence that the after-effects of binding the relevant task features change over time or that such changes differed between the two groups, suggesting that the group differences are relatively stable. These findings are in line with the recent observations that partial-repetition costs are more pronounced in individuals with low fluid intelligence (Colzato, van Wouwe, et al., 2006) and in young children and elderly participants, as compared with young adults (Hommel et al., 2011). Taken together, these findings suggest that less efficient control functions increase the impact of previously created feature bindings.

Studies have demonstrated ASD-related impairments in various executive functions (Hill, 2004; Ozonoff et al., 2004; Solomon, Ozonoff, Cummings, & Carter, 2008). Specifically, deficits in cognitive control and flexibility have long been associated with ASD (Geurts et al., 2009). Numerous studies have shown such deficits in various neuropsychological tests, such as the WCST task (Hill, 2004),



Converging evidence for an interesting theoretical and empirical link between ASD, control functions, and partialrepetition costs comes from research on the neuromodulation of the underlying cognitive processes and in particular from the fact that they all seem to rely on prefrontal dopaminergic pathways and their interaction with striatal dopaminergic pathways. Atypical dopaminergic activity is associated with ASD along with ADHD and schizophrenia (Previc, 1999, 2007). Considering the comorbidity between ASD and ADHD, it seems plausible that a dopaminerelated abnormality is the common source for the similar symptoms in these two disorders (Gillberg & Billstedt, 2000). In addition, some of the behavioral symptoms in ASD children were reduced by administrating risperidone (a dopamine-receptor antagonist; McCracken et al., 2002). Moreover, ASD is considered as one of the most highly heritable developmental disorder, and a number of genes linked to it (see: Yonan et al., 2003) are associated with dopamine, such as DBH (Robinson, Schutz, Macciardi, White, & Holden, 2001).

The dopaminergic system is also involved in a number of executive control functions, such as planning, working memory, or temporal sequencing (for a review, see Previc, 1999). More relevant to our study, there is compelling evidence that the dopaminergic system is important for mental flexibility and cognitive shifting operations. For instance, older adults show declines in dopaminergic transmission related to D1 (Rinne, Lonnberg, & Marjamaki, 1990; Suhara et al. 1991) and D2 receptors (Rinne et al., 1990; Volkow et al., 1996), and these declines are associated with poor performance in many neuropsychological controlrelated tests, such as the Stroop task, the WCST, and others (Volkow et al., 1998). This decline in cognitive ability can be corrected by administrating dopaminergic agonist such as Piribedil (Ollat, 1992). Furthermore, flexibility is improved by inducing positive affect (Dreisbach, 2006; van Wouwe, Band, & Ridderinkhof, 2011), which is assumed to induce temporary increases of the dopamine level (Ashby, Isen, & Turken, 1999; Dreisbach & Goschke, 2004). Thus, cognitive control and flexibility in particular appear to be modulated by dopamine.

Given that both ASD and cognitive flexibility seem to depend on dopaminergic pathways, it is interesting that the same seems to be true for partial-repetition costs. For



instance, these costs are systematically modulated by affect (Colzato, Wouwe, & Hommel, 2007a) and related to individual differences in the spontaneous eyeblink rate (Colzato, Wouwe, & Hommel 2007b), a clinical marker for the level of dopaminergic functioning (Blin, Masson, Azulay, Fondarai, & Serratrice, 1990; Kleven & Koek, 1996). Moreover, aftereffects of stimulus-response bindings are affected by stress (Colzato et al., 2008) and the use of cannabis, but not cocaine (Colzato & Hommel, 2008), suggesting that it is mainly dopaminergic D1 receptors that are involved, but not D2 receptors. Given that D1, but not D2, receptors are dominant in the mesocortical dopaminergic pathways, which are also assumed to drive executive control functions including working memory (e.g., Arnsten & Goldman-Rakic, 1998), these observations provide converging evidence for a link between ASD, executive control, and the management of episodic feature bindings.

To conclude, meeting the challenges a dynamic environment poses requires the human brain to find a balance between stability—as in integrating and keeping integrated the features of perceived event along with the actions they afford-and flexibility-as in updating, managing, and adapting integrated representations. Distortions of this balance can cause inflexibility and maladaptive behavior as can be seen in developmental disorders (Previc, 2007), substance abuse (Colzato & Hommel, 2008; Previc, 2007), and stress-related behaviors (Arnsten & Goldman-Rakic, 1998; Colzato et al., 2008). The present study provides evidence that ASD is associated with a specific deficit in updating episodic stimulus-response representations. From a general neuroscientific perspective, this implies that ASD is targeting processes and functions that are crucial not so much for the creation of episodic bindings but for their cognitive control and updating. Accordingly, ASD can serve as a useful "model" for the breakdown of the control of retrieval from episodic memory. From a neuropathological perspective, it is interesting that even the relatively simple task we used to assess aftereffects of feature binding captures the essence of processes that also impair performance in more complex experimental tasks and neuropsychological tests.

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