Time-on-Task in Children with ADHD: An ex-Gaussian Analysis

Vincenza Tarantino,¹ Simone Cutini,¹ Carla Mogentale,² AND Patrizia Silvia Bisiacchi¹ ¹Department of General Psychology, University of Padua, Padua, Italy ²Centro Phoenix di Psicologia, Neuropsicologia, Psicoterapia - Bassano del Grappa, Italy

Centro Filoenix di Fsicologia, Neuropsicologia, Fsicolerapia - Bassano dei Grappa, naiy

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Abstract

Although it is widely known that high intra-individual variability (IIV) is a key characteristic of attention deficit/ hyperactivity disorder (ADHD), a detailed exploration of the IIV pattern during the time course of a cognitive task has never been carried out. In this study, 30 children with ADHD and 30 controls, were administered the Conners' Continuous Performance Task (CPT-II). The across-block individual performance of the groups was analyzed using an ex-Gaussian approach, which enabled a clearer understanding of how individual response times (RTs) fluctuate during a task in comparison with conventional measures of central tendency. While the conventional measures showed a significant group effect on mean RTs but similar RT trends across blocks between the two groups, the ex-Gaussian results revealed no actual differences between the two groups in the normally distributed component of mean RTs (*mu*). In contrast to the control group, the children with ADHD showed a steep increase in the exponentially distributed component of RTs (*tau*) across blocks, thereby indicating that extremely long RTs progressively increased soon after the beginning of the task. Taken together, the results demonstrated that sustained attention deficit in ADHD can be detected by analyzing the IIV in the first few task blocks. (*JINS*, 2013, *19*, 820–828)

Keywords: Attention-deficit/hyperactivity disorder, Intra-individual variability, Sustained attention, Continuous performance task, Ex-Gaussian function, Developmental disorder

INTRODUCTION

Although most children with Attention Deficit/Hyperactivity Disorder (ADHD) are clinically described as impulsive and fast to respond, their performance in cognitive tasks is surprisingly slow, given that they generally report significantly higher mean response times (RTs) compared to a control group, regardless of the specific ability under examination (e.g., Klein, Wendling, Huettner, Ruder, & Peper, 2006; Nigg, Willcutt, Doyle, & Sonuga-Barke, 2005; Willcutt, Doyle, Nigg, Faraone, & Pennington, 2005). This apparent divergence has recently been accounted for through finer-grained analyses of RTs during the execution of computerized tasks requiring repeated responses. In fact, when examining the trial-by-trial distribution of RTs collected during a task, children with ADHD typically show both extremely fast and extremely slow responses (for a review, see Castellanos, Sonuga-Barke, Milham, & Tannock, 2006), thus presenting a high intra-individual variability (IIV).

Therefore, higher mean RTs have been hypothesized to be the result of a periodic fluctuation in response times rather than a general slowing of responses.

Remarkably, high IIV in ADHD can consistently be observed in a number of studies (Epstein, Langberg, et al., 2011), and it has been suggested that high IIV should not be considered as simply a reflection of error variance, but as one of the core features of ADHD, and one which can be regarded as a reliable clinical index because it correlates with ADHD diagnosis (Castellanos et al., 2005; Epstein et al., 2003; Leth-Steensen, Elbaz, & Douglas, 2000). Crucially, stimulant medication (e.g., methylphenidate) has been found to attenuate individual RT variability in children with ADHD (Castellanos et al., 2005; Epstein et al., 2006; Epstein, Brinkman, et al., 2011; Spencer et al., 2009; Tannock, Schachar, & Logan, 1995; Teicher, Lowen, Polcari, Foley, & McGreenery, 2004). Studies have analyzed the frequency of extremely long responses in time series RT data by means of Fast Fourier Transform (FFT), and have found that they have a characteristic periodicity in children with ADHD, namely they occur at a frequency of 0.05 Hz (i.e., approximately every 20 s; Castellanos et al., 2005). Later findings support

Correspondence and reprint requests to: Vincenza Tarantino, Department of General Psychology, Via Venezia, 8, 35131 Padua, Italy. E-mail: vincenza. tarantino@unipd.it

the existence of a low frequency periodicity for long RTs (in the range of 0.03–0.07 Hz) in a variety of tasks, such as the Sustained Attention to Response Task (SART; Johnson et al., 2007), the Ericksen Flanker Task (Di Martino et al., 2008), and the Go/NoGo Task (Vaurio, Simmonds, & Mostofsky, 2009).

Therefore, a considerable body of research has recently been developed to understand the nature of the IIV of RTs in ADHD. It has been thought to reflect moment-to-moment fluctuations in attention (Castellanos & Tannock, 2002; Douglas, 1999), lack of top-down control (Bellgrove, Hester, & Garavan, 2004; Castellanos et al., 2005), failure of response inhibition (Ridderinkhof, 2002), or state regulation (Geurts et al., 2008; Kuntsi, Oosterlaan, & Stevenson, 2001; Sergeant, 2005; Sergeant, Geurts, Huijbregts, Scheres, & Oosterlaan, 2003). The exact nature of such phenomena, however, is still a matter of debate.

When analyzing RTs one needs to take into account that individual RTs are not adequately described by conventional measures of central tendency, such as total mean and standard deviation (SD; Hockley & Corballis, 1982; Ratcliff & Murdock, 1976). In both typically developing controls and clinical populations, the RT distribution does not present a normal shape but is positively skewed (Luce, 1986). Specifically, the distribution of RTs has been demonstrated to fit optimally with an ex-Gaussian probability density function resulting from the convolution of a Gaussian and an exponential curve (Burbeck & Luce, 1982). Three parameters describe the ex-Gaussian fit: $mu(\mu)$, sigma (σ), and tau (7) (Heathcote, Popiel, & Mewhort, 1991; Ratcliff & Murdock, 1976). The *mu* and the *sigma* parameters represent the normally distributed components of the curve. The tau parameter represents the exponentially distributed component, which accounts for the positive skew of the RT distribution. The mean of the ex-Gaussian distribution corresponds to the sum of mu and tau. The standard deviation of the distribution corresponds to the sum of *tau* and *sigma*. Notably, according to the ex-Gaussian approach, extremely high RT values are not treated as outliers but are included in the analysis as they contribute both to the calculation of the mean response time and the size of the variance, and consequently lead to higher values of global mean and standard deviation. Therefore, a child showing higher mu values and low tau values likely has generalized slowing response times, whereas a child showing lower mu values and higher tau values likely has generally fast responses intermixed with extremely slow responses.

Decomposing response times recorded during computerized tasks into ex-Gaussian parameters has been shown to be an efficient approach for providing a more sensitive and specific measure of variability in patients with ADHD (e.g., Borella, de Ribaupierre, Cornoldi, & Chicherio, 2012; Buzy, Medoff, & Schweitzer, 2009; Epstein, Langberg, et al., 2011; Hervey et al., 2006; Leth-Steensen et al., 2000; Vaurio et al., 2009). In a foreperiod task, for example, Leth-Steensen et al. (2000) found that the RT distribution in children (9–13 years old) with ADHD was characterized by significantly higher values of the exponential tau parameter when compared to an age-matched control group, but had similar values for *mu* and *sigma* parameters. Furthermore, the authors showed that *mu* and *sigma* values were significantly higher in younger control children (7 years old) compared to the ADHD group. On the one hand, these results indicated that the RTs of children with ADHD differed from the non-clinical sample because of the presence of a larger number of excessively long RTs (i.e., beyond the individual mean), and on the other hand that ex-Gaussian parameters were differentially affected by individual variables, such as age. In a Continuous Performance Task (the Conners' CPT-II), Hervey et al. (2006) found that while traditional RT analyses showed significantly slower and more variable RTs in children with ADHD, ex-Gaussian analyses revealed lower mu (i.e., faster RTs) and higher sigma values. Notably, the largest group difference was reported in the *tau* values, which significantly increased as the inter-stimulus interval (ISI) duration increased, especially in the ADHD group. This result strongly suggested the presence of an inefficient phasic attention, which in turn affected response preparation. In two Go/NoGo tasks, Vaurio et al. (2009) found that children with ADHD presented significantly higher values of both the normal (sigma) and the exponential (tau) components of variability, regardless of the working memory load.

To date, the IIV of RTs in ADHD has been investigated by taking into account only the overall performance or the effect of ISI. In the present study, we overcome such limitation by exploring how IIV fluctuates over the course of a CPT in children with ADHD and a control group. To this end, we compared error rates and IIV in RTs across the six blocks of the CPT; RTs were analyzed separately for each block using both a conventional and an ex-Gaussian approach. We expected to find an increasing trend of *tau* values (reflecting abnormally long RTs) across blocks in children with ADHD, significantly higher than in the control group. A different ex-Gaussian pattern emerging from the two groups would demonstrate that managing sustained attention resources over the course of a task is one of the core abilities impaired in ADHD.

METHOD

Participants

A total of 30 children with a diagnosis of ADHD, aged 8–13 years (M = 11.48; SD = 1.73; male = 26), were enrolled at a local Clinical Service ("Centro Phoenix di Psicologia, Neuropsicologia, Psicoterapia", Bassano del Grappa, Italy). A diagnosis of ADHD was established by qualified neuropsychologists on the basis of DSM-IV-TR criteria (APA, 2000), using a clinical interview and by asking parents to complete two behavioral checklists, standardized for the Italian population, for assessing hyperactivity and inattentive behaviors (SDAG; Cornoldi, Gardinale, Masi, & Pettenò, 1996), and the Child Behavior Checklist (CBCL; Achenbach, 1991). The clinical interviews with parents assessed whether

the child met DSM-IV-TR criteria for ADHD, both for inattention and hyperactivity/impulsivity, namely the presence of six symptoms displayed at home and at school. The SDAG scale contains 18 items investigating all DSM-IV-TR symptoms, which are divided into two subscales: one for inattention and one for hyperactivity-impulsivity behaviors. Parents are required to evaluate the frequency of each behavior on a fourpoint Likert scale, ranging from "almost never" (0) to "very often" (3). The cutoff for considering a child for a possible diagnosis of ADHD was a total score of 14 in either one of the two subscales (Marzocchi, Re, & Cornoldi, 2010). In the ADHD group, the mean SDAG score on the inattention scale was 18.5 (SD = 2.70), and on the hyperactivity-impulsivity scale was 15.8 (SD = 3.50). A total of three (10.00%) children obtained scores above the clinical cutoff on the internalizing scale of the CBCL (Frigerio et al., 2004), and a total of four (13.33%) children obtained scores higher than the clinical cutoff on the externalizing scale. On the basis of both the clinical evaluation and the scores derived from the questionnaires, the following percentages relating to ADHD subtypes were identified: 8 (26.66%) predominantly inattentive, 3 (10.00%) predominantly hyperactive/impulsive, and 16 (63.33%) combined subtype. All the children included in the study obtained full-scale IQ scores above 85 (M = 102.73; SD = 11.53), assessed by the Wechsler Intelligence Scale for Children (WISC-III; Orsini & Picone, 2006; Wechsler, 1991). Based on reading, writing, and calculation tests, none of them presented learning disabilities. Reading abilities were assessed by means of words and non-words lists (Sartori, Job, & Tressoldi, 2009) and text reading (Cornoldi & Colpo, 1998); writing skills were assessed by means of a sentence writing test (Sartori et al., 2009) and a praxis (Tressoldi & Cornoldi, 1991) test; mathematical abilities were assessed by means of a battery of tests (Biancardi & Nicoletti, 2004). Learning disabilities were defined if children scored below the 10th percentile in at least two measures for each domain.

A total of 30 age and sex-matched children were included in the control group (age range = 8-13 years; M = 11.30; SD = 1.26; male = 26). The two groups did not differ in age (p = .642). Control participants were recruited at schools in the same geographical area (a region in northeast Italy) as the ADHD group, and came from the same socioeconomic background. Children were included in the control group if their IQ scores were above 85, if teachers reported neither cognitive or behavioral deficits nor learning disabilities, and if their parents scored lower than the cutoff in both the two subscales included in the SDAG questionnaire. The mean full IQ in the control group (M = 107.56; SD = 10.43) did not differ from the mean IQ in the ADHD group (p = .104). The mean SDAG scores of the control group on the inattention scale was 9.80 (SD = 3.70); the mean SDAG score on the hyperactivity-impulsivity scale was 7.20 (SD = 3.20). The SDAG scores of the ADHD group were significantly lower than the control group (p = .020, and p < .001, respectively).

None of participants had a history of seizures, brain injury or other neurological damage, uncorrected visual, auditory or speech deficits, pervasive developmental disorders, or medical conditions that may mimic ADHD. All children with ADHD were medication-naïve. The study was conducted according to the principles stated in the Declaration of Helsinki and parents were informed about the general experimental procedures and provided written consent.

Procedure and Task

All children performed the Conners' Continuous Performance Test (CPT-II; Conners, 2000) within the standard neuropsychological assessment session. The task takes approximately 14 min to be completed. With respect to other previously investigated tasks (5.5 min, Johnson et al., 2007; 8 min, Vaurio et al., 2009), this is an optimal duration for observing potential deficits related to sustained attention. The task was presented individually to each child using a laptop computer. Children were instructed to press the spacebar whenever a letter appeared on the screen (Go trials), except for the letter "X" (NoGo trials). The task included a total of 360 letters consecutively presented at the center of the screen, each for 250 ms. The 360 trials were divided into six blocks, each consisting of 60 trials. The blocks were divided into three randomly presented sub-blocks, one for each ISI (1, 2, or 4s): the three levels of ISI were randomized across trials. The appearance of the NoGo stimulus ("X") was rare (10%: 36 times) and RTs were measured from the onset of the letter appearing on the screen.

Data Analysis

Two types of error were recorded: omissions (no response to the target letters) and commissions (response to the Xs). The ex-Gaussian parameters (mu, sigma, and tau) of the probability distribution were calculated using the *egfit* MATLAB function (Lacouture & Cousineau, 2008). This function computes an iterative search process to fit the ex-Gaussian probability density function to the frequency distribution, and generates the three parameters from which the observed RTs are most likely to be sampled. Anticipated responses (RTs < 100 ms) were excluded from the analysis. A total of 1.62 (SD = 2.08) anticipations in the ADHD group and 0.73 (SD = 1.34) in the control group was found. On average, the RT distribution did not fit the ex-Gaussian function but instead approximated to an exponential function (i.e., sigma values were lower than 1) in 7.3% of the blocks in the ADHD group and in 4% of the blocks in the control group. In these cases, estimated values of mu, sigma, and tau were not included in the analysis but were replaced by the average value of the sample. Notably, the Kolmogorov-Smirnov normality test conducted on each individual block did not yield statistically significant results.

To examine the effect of group and block on these measures, separate 2×6 mixed model analyses of covariance (ANCOVAs) were conducted, in which Group (ADHD *vs.* control) was entered as the between-subject factor, Block number as the within-subject factor, and Age as covariate. Since the focus of the study was to examine whether the time-on-task effect and the number of trials at each ISI is equally

distributed across blocks, we collapsed different ISIs; by doing so, we obtained several trials for a reliable estimate of ex-Gaussian parameters.

An alpha level of .050 was considered for statistical significance. To control for multiple testing (i.e., seven different outcomes for the same task), a false discovery rate correction (FDR) was applied to *p* values (Benjamini & Hochberg 1995). Within each ANCOVA model, the Bonferroni correction was applied in *post hoc* analyses. Effect sizes were calculated in terms of partial eta squares (η_p^2). The relationship between age and error rates or RT parameters, as well as the association between error rates and RT parameters, was further examined by calculating Spearman's correlation coefficients (rho) between errors, Gaussian and ex-Gaussian measures.

RESULTS

Errors

The ANCOVA revealed that the overall number of omissions was significantly higher in the ADHD group in comparison with the control group (F(1,57) = 30.112; p < .001; $\eta_p^2 = .346$). A main effect of Block was found (F(5,53) = 2.864; p = .035; $\eta_p^2 = .048$). Importantly, the Group × Block interaction was also significant (F(5,53) = 4.313; p = .003; $\eta_p^2 = .070$), showing that the block number (order) only had a significant effect on the number of omissions in children with ADHD. The *post hoc* analysis of the interaction revealed that while omissions did not differ across blocks in the control group, children with ADHD made significantly more omission errors in blocks 4, 5, and 6 compared to block 1 (all *ps* < .003), and in blocks 5 and 6 compared to block 2 (both *ps* = .004). Figure 1a shows the distribution of the mean omission errors across the six blocks in the two groups, corrected for age.

No differences were found in the mean number of commissions between the two groups (F(1,57) = 0.172; p = .680; $\eta_p^2 = .003$). As illustrated in Figure 1b, the total number of commission errors was high in the ADHD group as well as in the control group (the maximum number of possible commission errors is six). Age did not influence commission but significantly interacted with block number on the number of omissions $(F(1,53) = 2.760; p = .040; \eta_p^2 = .046)$. In particular, age negatively correlated with the mean number of omissions in block 3 (rho = -.361; p = .005).

Gaussian RT Parameters

As expected, overall mean RTs were significantly higher in the ADHD group (F(1,57) = 17.977; p < .001; $\eta_p^2 = .243$). The ADHD group showed significantly higher mean RTs compared to the control group in all blocks (all ps < .016). Both groups showed a steep increase in mean RTs after the first block of trials; subsequently this remained relatively stable throughout the entire task (see Figure 2a). The ANCOVA did not yield a significant Block (F(5,53) = 0.706; p = .619; $\eta_p^2 = .012$) or Group × Block interaction (F(5,53) = 1.511; p = .187; $\eta_p^2 = .026$).

Overall, children with ADHD showed significantly more variability than children in the control group, as revealed by analyzing the mean RT standard deviation (F(1,57) = 21.221; p < .001; $\eta_p^2 = .275$). As shown in Figure 2b, the mean RT standard deviation of children with ADHD rose in the last three blocks of the task, whereas it remained stable in the control group. The ADHD group showed significantly higher SD than the control group in all blocks except the first and last ones. The Block (F(5,53) = 1.416; p = .219; $\eta_p^2 = .025$) or interaction effect (F(5,53) = 2.263; p = .064; $\eta_p^2 = .039$) did not reach statistical significance.

In all children, age significantly affected mean RTs $(F(1,57) = 14.48; p = .001; \eta_p^2 = .243)$ and mean RT standard deviation $(F(1,57) = 9.94; p = .009, \eta_p^2 = .151)$. The correlation analysis between mean RTs and age confirmed that, regardless of block, as age increased mean RTs and mean RT standard deviation significantly diminished (rho = -.371; p = .004 and rho = -.278; p < .033, respectively).

Ex-Gaussian RT Parameters

As shown in Figure 2c, mean *mu* values increased from the first to the second block in both groups, a pattern similar to that observed in mean RTs calculated according to the

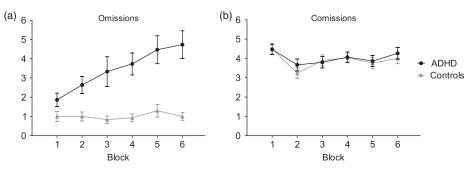


Fig. 1. Mean number of (a) Omission and (b) Commission errors plotted according to six blocks of the Conners' Continuous Performance Task (CPT) and to group (black line: attention deficit/hyperactivity disorder [ADHD], gray line: control). Data are age-corrected. Bars represent standard errors.

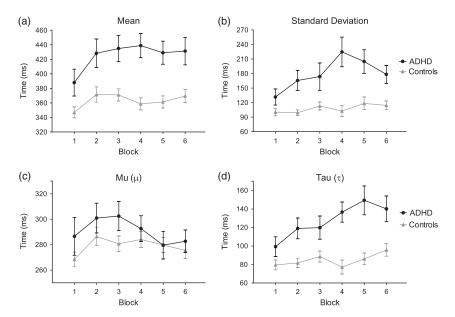


Fig. 2. Response times expressed in terms of Gaussian parameters: (a) mean, (b) standard deviation; and ex-Gaussian parameters: (c) $mu(\mu)$ and (d) $tau(\tau)$. Data are plotted according to the six blocks of the Conners' Continuous Performance Task (CPT) and to group (black line: attention deficit/hyperactivity disorder [ADHD]; gray line: control). Data are age-corrected. Bars represent standard errors.

Gaussian distribution (cf., Figure 2a). After the second block, however, mean *mu* values demonstrated a different pattern, in that they decreased in the ADHD group while they remained approximately stable in the control group. A main effect of Block was found (F(5,53) = 2.606; p = .049; $\eta_p^2 = .044$). The *post hoc* comparisons showed that mean *mu* values in block 2 were significantly higher than in block 5 and 6 (both ps = .026). Notably, *mu* values did not significantly differ between the two groups in any block (F(1,57) = 2.012; p = .162; $\eta_p^2 = .035$).

The ex-Gaussian parameter *sigma* was significantly higher in the ADHD group in all task blocks (F(1,57) = 18.420; p < .001; $\eta_p^2 = .248$). There was no significant main effect for Block (F(5,53) = 1.299; p = .264; $\eta_p^2 = .023$) or Group × Block interaction effect (F(5,53) = 1.668; p = .142; $\eta_p^2 = .029$).

The most interesting results were related to the *tau* parameter, which presented an evident increase from block 1 to block 6 in the ADHD group, as shown in Figure 2d. Overall, mean *tau* values were higher in the ADHD group $(F(1,57) = 28.268; p < .001; \eta_p^2 = .335)$. Moreover, the ANCOVA yielded a significant main effect of Block $(F(5,53) = 3.562; p = .012; \eta_p^2 = .060)$ and Group × Block interaction $(F(5,53) = 2.638; p = .048; \eta_p^2 = .045)$. Bonferroni *post hoc* analyses showed that *tau* values in the ADHD group were significantly higher in blocks 2, 4, 5, and 6 compared to block 1 (all *ps* < .033), in blocks 5 and 6 compared to block 2 (all *ps* < .047), and in block 5 compared to block 3 (*p* = .025). Mean *tau* values were significantly higher in the ADHD group compared to the control group in all blocks except the first one (all *ps* < .014).

The across-block pattern of the *tau* was explored in detail by fitting a linear regression equation, separately for

the ADHD and the control group. The mean slope significantly differed from zero only in the ADHD group ($\beta = 8.926 \pm 1.914$; 95% confidence interval [CI] = 3.613–14.240), not in the control group ($\beta = 2.358 \pm 1.378$; 95% CI = -1.477-6.186). Furthermore, the two linear regression were significantly different (*F*(1,8) = 7.749; *p* = .023).

Remarkably, the fact that in both groups the across-block trend of *tau* values mirrored the pattern observed for the normally distributed RT standard deviation (Figure 2b) suggests that intra-individual RT variability in ADHD children is mostly related to extremely slow responses.

Age significantly affected mean *mu* (*F*(1,57) = 7.719; p = .021; $\eta_p^2 = .121$), *sigma* (*F*(1,57) = 21.51; p < .001; $\eta_p^2 = .248$), and *tau* (*F*(1,57) = 15.762; p = .001; $\eta_p^2 = .220$) values, namely they significantly decreased with age. Furthermore, age significantly interacted with Block in influencing *mu* (*F*(5,53) = 2.935; p = .035; $\eta_p^2 = .050$) and *tau* (*F*(5,53) = 2.908; p < .035; $\eta_p^2 = .049$). Correlation analyses showed that age negatively correlated with *mu* values in blocks 1, 4, and 6 (rho< -.273; p < .037); whereas it negatively correlated with *tau* in all but the first block (rho< -.270; p < .038).

Correlations between Measures

Table 1 shows the results of correlation analyses between omission and commission errors with Gaussian and ex-Gaussian parameters. As illustrated in the table, in the ADHD group omission errors were significantly and positively correlated with the *sigma* parameter, while commission errors were negatively correlated with the *mu* parameter. Importantly, in the ADHD group, RT standard deviation was significantly correlated only

Spearman's rho	Omissions	Commissions	RT mean	RT standard deviation	mu	sigma	tau
Omissions	_	.512**	.017	.553**	330	.158	.372*
Commissions	.030	_	359	.289	562**	.217	.075
RT mean	.279	590**	_	.566**	.820***	.371*	.700***
RT standard deviation	.396*	125	.631***	-	.139	.567**	.885***
Mu	.047	708***	.722***	.073	_	.247	.260
Sigma	.522**	.056	.427*	.359	.281	_	.416*
tau	.396*	294	.850***	.889***	.351	.406*	-

Table 1. Non parametric correlation coefficients (rho) between errors, Gaussian and ex-Gaussian measures. Lower diagonal values refer to ADHD group, upper diagonal values refer to control group.

p < .05, p < .01, p < .01, p < .001.

with the *tau* parameter, while in the control group, it correlated with the *tau* and the *sigma* parameters. This confirms that the RT variability in ADHD is principally due to extremely long RTs.

DISCUSSION

The aim of the study was to examine intra-individual variability during a sustained attention task in children with ADHD by investigating their block-by-block fluctuation in responses, in terms of variations of errors (omission, commission), conventional RT measures (mean RT and standard deviation), and ex-Gaussian RT parameters (*mu*, *sigma*, and *tau*). To this end, these measures were compared across the six blocks of a CPT lasting 14 min (CPT-II; Conners, 2000), based on a sample of children with ADHD and a control group.

The results revealed that omission errors progressively increased after the first block of trials (i.e., at approximately 2.5 min from the beginning of the task) in the ADHD group but not in the control group. As expected, the conventional Gaussian analysis showed a significant main effect of group on mean RTs. In addition, it revealed the presence of a similar across-block trend between the two groups, namely RTs significantly increased after the first block and remained steady afterward. In contrast, the ex-Gaussian analysis revealed no actual differences between the two groups in terms of the normally distributed RT parameter mu. On the other hand, mean mu values showed a completely different across-block pattern from that of the Gaussian values (Figure 2a and 2c): they significantly decreased in the last two blocks. Importantly, the tau trend across blocks differed between the two groups: the ADHD group showed a significant increase, namely tau values steeply and progressively increased from the first block to block 6, whereas the control group they remained steady until the end of the task. These findings clearly suggest that the higher omission rates and mean RTs observed in children with ADHD were mostly due to the exponential component of RTs (i.e., to extremely long RTs), which significantly increased as the task proceeded. In addition, the fact that tau values did not differ between groups in the first block but differed in all following blocks suggests that intra-individual variability increases with time-on-task demands.

The across-block variation in intra-individual variability was consistent with that reported by Klein et al. (2006). Indeed, in a similar Go/NoGo task, these authors found a significant increase of RT standard deviation across the 15 blocks of a CPT in a group of children with ADHD. Our study further extended these findings by demonstrating that intra-individual variability of RTs in children with ADHD was mostly caused by abnormally long responses, as suggested by the tau parameter distribution. The present findings are in agreement with previous studies showing the tau parameter to be a sensitive marker of ADHD-related between-groups differences, based on CPT (Hervey et al., 2006), choice discrimination (Leth-Steensen et al., 2000), and Go/NoGo (Epstein, Langberg, et al., 2011; Vaurio et al., 2009) tasks. Furthermore, our results support previous evidence which shows significant low frequency fluctuations throughout a task in individuals with ADHD (Castellanos et al., 2005; Di Martino et al., 2008; Helps, Broyd, Bitsakou, & Sonuga-Barke, 2011; Johnson et al., 2007; Vaurio et al., 2009). Indeed, it is possible that the prolonged RTs are related to abnormally slow spontaneous oscillations in brain activity.

Particularly important is the finding that children with ADHD showed a steep increase of intra-individual variability after few minutes from the commencement of the task (i.e., after the first block), while the control group did not show any relevant across-block variation in errors, Gaussian, and ex-Gaussian measures.

Unlike the Gaussian mean RT, the fact that overall the *mu* parameter (i.e., the normally distributed component of mean RTs) did not significantly differ between the two groups confirmed previous findings that children with ADHD are generally no slower than children in control groups (Hervey et al., 2006; Leth-Steensen et al., 2000). Furthermore, the fact that *mu* values in the ADHD group decreased after the third block while RT standard deviation simultaneously increased corroborates the idea the time-on-task demands affect intra-individual variability not the overall response time speed.

The normally distributed *sigma* parameter was overall higher in the ADHD group, which is consistent with previous reports (Hervey et al., 2006; Vaurio et al., 2009). In addition, overall mean *tau* values were positively correlated with *sigma* values only in the control group, suggesting that phasic and tonic attention are subject to covariation.

Commission errors did not differ between groups, given that the rate of NoGo trials was very low (10%); not surprisingly, the children in the control group also found it very difficult to inhibit their motor response. Similarly, previous investigations using the Conners CPT observed no differences in the total number of commission errors between the ADHD and control group (Epstein et al., 2010; Hervey et al., 2006).

Of interest, overall omission rates were positively correlated with the exponential component of mean RTs (tau) in both groups, meaning that both phenomena reflect a common process. On the other hand, the negative correlation between commission errors and the Gaussian component of mean RTs (mu) reveals that children who failed more often were those with overall faster overall RTs. The positive correlation between omission rates and tau, together with the negative correlation between commission rates and mu, may also reflect the effect of a speed-accuracy trade-off in both groups. Furthermore, the positive correlation between omissions and tau, together with the negative correlation between omissions and mu, in children with ADHD, may partially be explained by the presence of pre- and post-error slowing (Epstein et al., 2010). These results suggest that tau and mu parameters are likely linked to two independent processes, namely, lapses of attention (Castellanos & Tannock, 2002; Douglas, 1999) and failures of response inhibition (Ridderinkhof, 2002).

The findings presented here support the hypotheses of the presence of either a deficit of top-down control (Barkley, 1997; Bellgrove et al., 2004) or sub-cortical state regulation (Geurts et al., 2008; Kuntsi et al., 2001; Sergeant et al., 2003) in ADHD. From a neural perspective, the increase in excessively long RTs and in error rates over the course of the task might reflect an inefficient suppression of activity in those brain areas involved in the default network, such as the medial prefrontal cortex (Fassbender et al., 2009; Kelly et al., 2004; Sonuga-Barke & Castellanos, 2007; Weissman, Roberts, Visscher, & Woldorff, 2006). Furthermore, the activation of frontal brain areas might be involved as the task demands sustained attention (e.g., Manly et al., 2003; Stuss, Murphy, Binns, & Alexander, 2003; Wilkins, Shallice, & McCarthy, 1987). Specifically, dorsolateral prefrontal regions have been found to be associated with increased over-time variability and inter-trial fluctuations, even in easy choice RT tasks (Bellgrove et al., 2004; Stuss et al., 2003); these regions are also considered to be dysfunctional in individuals with ADHD (for reviews, see Bush, 2010, 2011).

Strength of this study is that, in addition to having confirmed the IIV potential as a biomarker of ADHD, we demonstrated that an increased time-to-task demand is closely associated with increasing IIV and error rates in ADHD. Based on these observations, IIV in cognitive tests can no longer be considered "statistical noise", and theories that stem from such a perspective should be dismissed (e.g., Castellanos et al., 2006).

Moreover, our study extended previous investigations on sustained attention deficit in ADHD, by showing that variations in individual performance across a task in children with ADHD are associated with difficulties in sustaining attention. We interpreted such results as a consequence of an inability to distribute attentional resources strategically over time, which in turns leads to off-task behaviors. Importantly, the analysis of the ex-Gaussian *tau* parameter allows the detection of an ADHD-related deficit in sustained attention even in the first few blocks of the task. These findings are relevant given that although the existence of sustained attention deficit in ADHD is a well-known phenomenon, it has been surprisingly poorly investigated and still remains controversial (Johnson et al., 2007; Klein et al., 2006; van der Meere, Shalev, Borger, & Gross-Tsur, 1995; van der Meere, Wekking, & Sergeant, 1991).

Some limitations of the study should be considered when interpreting the findings. First, the diagnostic interview with parents and teachers that aimed to identify behavioral and emotional disorders was not standardized; therefore, the inclusion of some children with such comorbid disorders in the ADHD group as well as in the control group could not be completely ruled out. Moreover, the control group was not systematically assessed for learning disabilities. Although the control group was recruited from schools in the same geographical area and is assumed to have similar socioeconomic background as the ADHD group, no data were available to confirm this. Given the relatively small sample size, the present results should be replicated by future studies; nevertheless, it is worth noting that the results are consistent with previous findings.

In conclusion, the block-by-block analysis of variations in errors and RTs unveiled the presence of a marked increase in errors and abnormally slow responses in children with ADHD as the task progressed. Of interest, there was no difference in the distribution of RTs across blocks between the two groups in terms of normal mean, a difference was found in extremely long responses. These results suggested that intra-individual variability of children with ADHD over the course of a sustained attention task should be regarded as a crucial aspect when evaluating these patients.

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