

Auditory Time Thresholds for Subsecond but not Suprasecond Stimuli are Impaired in ADHD

Giovanni Anobile

University of Florence

Mariaelisa Bartoli

IRCCS Fondazione Stella Maris

Chiara Pfanner

IRCCS Fondazione Stella Maris

Gabriele Masi

IRCCS Fondazione Stella Maris

Giovanni Cioni

IRCCS Fondazione Stella Maris

Francesca Tinelli (✉ francesca.tinelli@fsm.unipi.it)

IRCCS Fondazione Stella Maris

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Abstract

Background. The literature on time perception in individuals with ADHD is extensive but inconsistent, probably reflecting the use of different tasks and performances indexes.

Methods. A sample of 40 children/adolescents (20 with ADHD, 20 neurotypical) was engaged in two identical psychophysical tasks measuring auditory time thresholds for both subsecond (0.5-1s) and suprasecond (0.75-3s) stimuli.

Results. Results showed a severe impairment in ADHD for subsecond thresholds ($\text{Log10BF}=1.9$). The deficit remained strong even when non-verbal IQ was regressed out and correlation with age suggests a developmental delay. Suprasecond thresholds were indistinguishable between the two groups ($\text{Log10BF} = -0.5$) and not correlated with subsecond thresholds.

Conclusions. Since much evidence suggest that perception of subsecond stimuli does not load on cognitive control and working memory, the current results are consistent with a pure timing deficit in individuals with ADHD.

Introduction

According to current diagnostic systems (DSM-5), attention deficit hyperactivity disorder (ADHD) is defined by pervasive and severe symptoms of inattention, hyperactivity, and impulsivity that have a direct negative impact on social, academic, or occupational functioning. ADHD children often show deficits in planning, organization and in executive functions, such as response inhibition, interference control, reasoning, hindsight, anticipation and working memory set-shifting [1].

Along with these deficits, findings suggest perceptual dysfunctions related to time processing. Time perception in ADHD has been widely investigated but the results are mixed, probably reflecting the many different methods and performance parameters. Measures of time processing includes accuracy (how far from the target) and/or precision (response variability) measured by motor reproduction, verbal estimation, discrimination and odd-ball tasks. Moreover, time processing has been investigated across different timing ranges (from a few milliseconds to several seconds) and across sensory modalities [for a review see 2]. Even if the literature and the clinical practice indicate an impaired time processing in ADHD, the highly heterogeneous results make it difficult to draw firm conclusions.

A meta-analysis considering 27 studies on ADHD children and adolescents found a significant timing deficit in ADHD with impairments in both accuracy and precision and across visual and auditory stimuli [3]. Another recent meta-analysis considering 12 studies, despite generally confirming a time deficit in ADHD, suggested stronger effect size for tasks involving relatively long intervals (5s) and for specific tasks such as estimation and reproduction [4]. An example of how task and timing range could affect the results comes from the study conducted by Smith et al. [5]. The authors measured time discrimination thresholds in a sample of ADHD children by asking participants to indicate which one of two audio-visual

stimuli, ranging around 1s, lasted longer. Results revealed higher thresholds (lower precision) in ADHD. On the contrary, no group differences were found when time processing was measured with a reproduction task or with a verbal estimation task with longer (5s-10s) stimuli. Toplak & Tannock [6] also measured duration discrimination thresholds in a sample of ADHD adolescents for visual and auditory stimuli and for both short (around 200ms) and longer (around 1s) intervals. Results confirmed higher thresholds compared to controls, but across all the tasks. Similar impairments across vision and audition were later obtained by Rubia et al. [7] with younger ADHD children and by Dölek et al. [8] with adults. Plummer and Humphrey [9] measured ADHD children's timing performance with a motor reproduction task. Children were asked to reproduce, by key press, the duration of visual, auditory, or audio-visual stimuli ranging from 1 to 60s. Results showed higher errors in ADHD across all the conditions. Radonovich and Mostofsky [10] measured ADHD duration discrimination thresholds for short (around 0.5s) and longer (around 4s) auditory stimuli. At odds with some of the previous studies, the results showed no deficits for the short stimuli but poorer performance for longer intervals. Gooch et al. [11] measured auditory time discrimination thresholds in ADHD by asking children to detect which of three sounds was different in duration (odd-one-out task) with stimuli ranging from 400 ms to 1s. The results revealed higher thresholds (lower precision) in children with ADHD, compared to controls. The same results were obtained with a motor reproduction task for longer visual stimuli (from 2 to 10s). Barkley [12] measured time perception with a verbal estimation and a reproduction task testing the same visual stimuli, lasting from 2 to 60 s. The results revealed poorer time perception (accuracy) for the reproduction, but not for the estimation task. Two independent studies employing similar visual time intervals found the same pattern of results [13, 14].

Interestingly, psychophysical and pharmacological studies indicates that the perception of relatively long stimuli requires and loads on cognitive control as well as working memory, while relatively shorter intervals (< 1s) are automatically processed and targeting a "pure" sense of time [15–17].

Overall, these and many other investigations suggest a time processing deficit in ADHD, with potentially relevant clinical and practical implications [18]. However, given the heterogeneity of results, further investigation is needed. To this aim, 20 children/adolescents with ADHD were engaged in a testing protocol in which we psychophysically measured auditory time thresholds for both subsecond (0.5s) and suprasecond (1.5s) durations. To avoid methodological confounds, the same psychophysical technique (categorization task, see methods) was used for both timing ranges. The performance on these tasks was compared to those obtained from 20 age matched neurotypical controls. As many studies found time perception deficits in children and adolescents with ADHD, we expected higher thresholds, at least on one of the timing range tested here. As subsecond but not suprasecond stimuli depends on cognitive control and working memory, a specific deficit for subsecond stimuli would suggest a pure time deficit.

Materials And Methods

Participants

Forty children/adolescents participated in this study: 20 with ADHD (6 female, 14 males, mean age = 11.2 year old, age range 8–16) and 20 neurotypical (11 female, 9 males, mean = 11.2 year old, range 8.1–16.2). Individuals with ADHD were enrolled from the Stella Maris Foundation Institute in Pisa, a main centers for ADHD care in Italy. ADHD inclusion criteria were: clinical diagnosis of ADHD based on DSM-5; a total intelligence quotient (TIQ), evaluated with the Wechsler Intelligence Scale for Children-IV above 75 [19], no neurological or sensory deficits, no psychiatric comorbidities, no current or past pharmacological treatment. Three children with ADHD met the criteria for a diagnosis of developmental dyslexia. Non-verbal reasoning skills were computed by a combined index of WISC-IV measuring Visual Perceptual Reasoning (IRP). The IQ of four ADHD participants was measured by an external independent institute and, for those participants, we did not have the possibility to calculate IRP. ADHD symptoms were measured by Conners Rating Scale (parent version). General clinical symptoms were measured by the Clinical Global Impression – Severity scale (CGI) and the Children Global Assessment Scale (CGAS). Detailed information about the group with ADHD is reported in Table 1.

Table 1
Descriptive characteristics of the ADHD group.

Participants	Sex	Age	TIQ	IRP	Subtype	CGI-S	CGAS	z-score			
								C1	C2	C3	C4
1	M	15y 6m	103	1.13	1	3	60 - 51	n.a.	n.a.	n.a.	n.a.
2	F	9y 6m	90	-0.73	1	2	70 - 61	45	60	52	65
3	F	12y 3m	88	-1.2	3	2	70 - 61	42	68	51	67
4	M	9y 3m	94	-0.4	3	4	50 - 41	77	73	63	80
5	M	14y 11m	97	-1.2	3 + dd	4	50 - 41	77	75	61	75
6	M	13y 3m	125	n.a.	3	2	70 - 61	55	49	35	56
7	M	10y 5m	114	n.a.	1	2	60 - 51	74	70	70	72
8	M	8y 4m	91	-0.13	3	4	60 - 51	78	75	70	80
9	M	11 y	110	n.a.	3	5	50 - 41	71	75	80	80
10	M	9y 1m	114	1.26	3	4	60 - 51	80	75	63	80
11	M	12y 6m	88	n.a.	3	2	70 - 61	60	53	57	65
12	F	11y 1m	98	-1.2	3	4	50 - 41	n.a.	n.a.	n.a.	n.a.
13	M	11y 1m	100	0	3	3	60 - 51	55	50	47	61
14	F	9y 8m	105	-0.13	3	3	60 - 51	45	56	52	63
15	F	12y 1m	92	0.13	3 + dd	4	60 - 51	n.a.	n.a.	n.a.	n.a.
16	M	11y 9m	77	-1	1	3	70 - 61	70	100	56	100
17	M	12y 6m	101	-1.2	3	6	60 - 51	n.a.	n.a.	n.a.	n.a.

Participants	Sex	Age	TIQ	IRP	Subtype	CGI-S	CGAS	z-score			
								C1	C2	C3	C4
18	M	11y 6m	102	0.4	1	6	60 – 51	64	63	61	66
19	M	8y 7m	107	0.73	3	5	50 – 41	64	66	70	n.a.
20	F	10 y	96	1.13	3 + dd	6	60 – 51	49	39	48	13

TIQ = Total Intelligence Quotient from WISC-IV; IRP z-score = Visual Perceptual Reasoning from WISC-IV; Subtype = 1 inattentive, 3 combined; dd = developmental dyslexia; CGI: Clinical Global Impression – Severity scale; CGAS: Children Global Assessment Scale; C1: Conners parents oppositionality, C2: Conners parents inattention, C3: Conners parents hyperactivity, C4: Conners parents ADHD index, n.a.= not available

Children with ADHD were compared to a neurotypical group of age matched children/adolescents. The inclusion criteria for the control group were: no medical history, negative neuro-psychiatric exam and no learning difficulties (reported by parents) and IQ (evaluated with by Raven Colored Progressive Matrix-CPM or Progressive Matrix-PM, depending on chronological age) > 5° percentile. The study was approved by the Ethics Committee of the Meyer's Hospital (n. 248/2020 ID-DNATN "Attention, Time and Numeracy in children and adolescents with neurodevelopmental disorders"). Informed parental consents were obtained for each participant before the study. All experiments were performed in accordance with relevant guidelines and regulation.

Time perception

Time sensory thresholds were psychophysically measured with an auditory categorization task (Figure 1A). On each trial, children listened to a single sound (500 Hz, 80 dB pure tone) and were asked to categorize it as “long” or “short”. To provide an experience of the tested ranges, before the testing phase, four initial “anchoring” trials were provided. In these trials, the lower and longer time durations were played twice each and the children were told that those sounds corresponded to the range extremes (no responses were required). In separate sessions (lasting around 4 minutes each), we measured two different timing regimes, one centered (geometric mean) around 0.5 s containing stimuli in the subsecond range (from 0.25 s to 1 s), and one centered (geometric mean) around 1.5 s with most of the stimuli belonging to the suprasecond range (from 0.75 to 3 s). Each time range was divided into 11 equal steps spanning 1 octave above and one below the geometric mean of that specific range. The stimuli in the 1.5 sec distribution were: 0.75, 0.86, 1, 1.13, 1.3, 1.5, 1.72, 1.98, 2.27, 2.61, 3 s. The stimuli in the 0.5 s distribution were: 0.25, 0.28, 0.33, 0.38, 0.43, 0.5, 0.57, 0.66, 0.75, 0.87, 1 s. In a single session, each duration was tested 4 times (randomly selected trial-by-trial) for a total of 44 trials for each range. Participants responded verbally (“long” or “short”), without any time pressure and the response was registered by the administrator with an appropriate key press. The proportion of “long” responses were

plotted against the stimuli duration (in log scale) and fitted with a cumulative Gaussian error function. The 50% point of the fit provided an estimate of the point of subjective equality (PSE). The difference in duration between the 50% and 75% points gives the just notable difference (JND), which was used to estimate Weber Fractions (10^{JND-1}), a dimension free index of sensory precision.

Data analysis

Data were analyzed by RM-ANOVA, ANCOVA, t-test and Pearson correlations and α values corrected for multiple comparisons when necessary (Bonferroni correction). Frequentist statistics were supplemented with Bayesian statistics, calculating Bayes Factors, the ratio of the likelihood of the alternative to the null hypothesis, and reporting them as base ten logarithms (LBF). For RM-ANOVA and ANCOVA with report LBF_{inclusion} indicating how much the data are likely to occur from a model including that specific factor (or interaction), compared to models not including them. By convention, LBF > 0.5 is considered substantial evidence in favour of the alternative hypothesis (difference between groups in this case) and LBF < -0.5 substantial evidence for the null hypothesis (no difference). Absolute values greater than 1 are considered strong evidence, and greater than 2 definitive.

To make the ADHD Visual Perceptual Reasoning (IRP) index comparable to the non-verbal reasoning skills measured by Raven matrices on the control group, the indexes were both converted into z-scores (according to the normative age-standardized data provided by the tests manuals). For technical reasons we did not collect data for two participants (one control and one ADHD) in the suprasecond timing task. Missing data were left empty and excluded analysis by analysis. Effects sizes were reported as Cohen-d and η^2 . Data were analyzed by JASP (Version 0.8.6), SPSS (Version 25) and R (Version 4.0.2) software.

Results

Demographical data

The ADHD and control groups did not differ in age ($t_{(38)}=0.035$, $p= 0.97$, $d=0.011$, LBF= -0.52) and male/female ratio ($X^2=0.93$, $p=0.33$). Non-verbal reasoning skills were slightly lower in the group with ADHD ($t_{(32)}=1.8$, $p= 0.083$, $d=0.6$, LBF= 0).

Groups difference on time perception

All participants were well able to perform the psychophysical timing task (depicted in Figure 1A) producing ordered psychometric functions. Figure 1 (panels B-C) shows psychometric functions obtained aggregating all the data together across participants. It is evident, even by inspection, that for the task measuring time perception in the subsecond range (B), the psychometric function of the control participants (black) was steeper than that produced by the sample with ADHD (red). This difference indicates less precision (higher thresholds, Weber Fractions) in individuals with ADHD. Regarding the

suprasecond stimuli (C) the psychometric functions have similar slopes between groups, indicating similar sensory precision levels.

The fitting procedure was applied to the data provided by each participant. Figure 2 reports between participants average thresholds (Weber Fractions, Wf) separately for the two perceptual tasks, while single subject data are reported in figure 2 B&C. From visual inspection, it is evident that only the time perceptual thresholds measured for short (subsecond) auditory stimuli were impaired in participants with ADHD, with a clear interaction between tasks and groups.

A RM ANOVA with task (2 levels: Wf subsecond, Wf suprasecond) as repeated measures factor and group (2 levels: ADHD, controls) as between participants factor, revealed a significant effect of task ($F_{(1,36)} = 7.38$, $p = 0.01$, $\eta^2 = 0.12$, LBF_{incl} = 2.31), suggesting different thresholds across tasks. Crucially the task*group interaction was highly statistically significant ($F_{(1,36)} = 18.04$, $p < 0.001$, $\eta^2 = 0.29$, LBF_{incl} = 2.59) indicating that the groups performed differently across the tasks. Post-hoc analyses confirmed that time thresholds for subsecond stimuli were higher in the ADHD group compared to controls ($t_{(38)} = 3.98$, $p < 0.001$, $d = 1.26$, LBF = 1.9). On this task, ADHD thresholds were, on average, almost double compared to controls (Wf = 0.23 and 0.12 for ADHD and controls respectively), indicating a severe impairment. Time thresholds for suprasecond stimuli ($t_{(36)} = 0.26$, $p = 0.79$, $d = 0.08$, LBF = -0.5) were statistically indistinguishable between the groups.

To explore the specificity of the impairment found for the time perception task in the subsecond range, we ran a separate ANCOVA with time thresholds as the dependent variable, groups (ADHD, controls) as the fixed factor and age, non-verbal reasoning, and sex as covariates. Even when partialling out the effect of these covariates, the result remained unchanged with a significant effect on the group ($F_{(1,29)} = 20.11$, $p < 0.001$, $\eta^2 = 0.33$, LBF_{incl} = 2.78).

Table 2						
Descriptive Statistics on time thresholds (Weber Fraction)						
Tasks	Groups	N	Mean (SD)	p-value	Cohen's d	LBF
Time subsecond (0.5s)	ADHD	20	0.23 (0.1)	< 0.001***	1.26	1.9
	Controls	20	0.12 (0.05)			
Time suprasecond (1.5s)	ADHD	19	0.15 (0.07)		0.08	-0.5
	Controls	19	0.14 (0.06)	0.79		
Two tailed t-tests, a Bonferroni corrected $0.05/2 = 0.025$						

To check the discriminant power of the subsecond auditory time thresholds, we ran a linear discriminant analysis with the group as the dependent variable and time thresholds (Wf) as independent variable. The results revealed 72.5% of cases correctly classified. The sensitivity was 60% while specificity was 85%. As

a sanity check, the same analysis on suprasecond thresholds (Wf) provides a near to chance level (53%) classification.

Developmental trajectories

To investigate whether the deficit was stable across the age range, we studied the developmental trajectories. Time thresholds for suprasecond stimuli had a similar and not significant dependency with age across both groups (ADHD: $r = -0.43$, $p = 0.062$, LBF = 0.15, Controls: $r = -0.42$, $p = 0.068$, LBF = 0.12), suggesting that both were near to a developmental plateau. The developmental trajectories of time thresholds in the subsecond range were, in contrast, different between the groups. While controls had reached an almost full developmental stage ($r = -0.34$, $p = 0.14$, LBF = -0.11), the age dependence for participants with ADHD was steeper ($r = -0.62$, $p = 0.003$, LBF = 1.2), suggesting a different developmental trend (Figure 3). Confirming partially independent mechanisms, regressing out age, thresholds for subsecond and suprasecond stimuli were not correlated with each other (ADHD: $r_{\text{partial}} = 0.367$, $p = 0.134$, LBF = 0.09 ; Controls: $r_{\text{partial}} = 0.287$, $p = 0.25$, LBF = -0.07).

Correlations with clinical symptoms

Within the sample with ADHD, we run correlations and between time thresholds (subsecond and suprasecond) and both general (CGI, CGAS, see Table 1) and specific clinical symptoms (the four parents Conners indexes, see Table 1 for details). For the CGAS test, which provides range scores, we transformed the ranges into categorical values reflecting the symptoms severity (following the test manual: from 1 to 10 with one indicating no symptoms and 10 indicating very severe symptoms). The analyses revealed no meaningful correlations (all $p > 0.05$, min LBF = -0.55, max LBF = 0.3).

Perceptual task reliability

It is theoretically possible that the different pattern of results provided by the two time tasks results from different reliability levels. To test this possibility, we measured and compared the reliability of the two psychophysical tasks. Following previous studies [20], we used a “sample-with-replacement” bootstrap technique [21]. For each participant, we calculated two separate thresholds in each task (0.5 or 1.5s), using a random sample of the data (44 trials, sampled with replacement), and then computed the correlation between those two measures, across participants. The process was reiterated 1,000 times. We found that mean correlations for subsecond (0.5s) and suprasecond (1.5s) stimuli were very similar (Pearson’s $r = 0.68$, $r = 0.64$, respectively) and not statistically different (bootstrap sign-test $p = 0.4$). This last control rules out the possibility that the different pattern of results was generated by different reliability levels.

Contextual effects

The paradigm used to measure time thresholds requires the ability to perceive both the stimuli mean and range extremes of the set. The group with ADHD could have had excessive contextual effects, which would have inflated thresholds. To check for this possibility, we measured (on aggregate data) the PSEs and thresholds as a function of the magnitude of the preceding stimulus (N-1). To this aim, separately for the two groups and the two tasks, we sorted the aggregated data into two categories in which the preceding stimulus (N-1) was shorter or longer than the stimulus tested in the current trial. The data were then fitted by psychometric functions providing PSEs and thresholds (Weber fraction). The analyses revealed very small effects on PSEs for both groups (ADHD 1.5s= shorter 1.4s, longer 1.5s ; ADHD 0.5s= shorter 0.56s, longer 0.61s ; Controls 1.5s= shorter 1.6s, longer 1.6s ; Controls 0.5s= shorter 0.55s, longer 0.56s), suggesting similar contextual effects. More importantly, for both subdivisions (shorter, longer), the difference between groups on subsecond thresholds remained evident and constant (Figure 4) confirming similar contextual effects.

Discussion

We found that children/adolescents with ADHD, compared to neurotypicals, had a severe sensory precision deficit in perceiving subsecond (0.25-1s) auditory stimuli. Time thresholds for relatively longer stimuli (0.75-3s) were completely unimpaired. Thresholds did not correlate between each other, and only subsecond thresholds showed a different developmental trajectory between groups, suggesting a developmental delay in participants with ADHD. Two control analyses ruled out the possibility that the pattern of results was driven by different tasks reliability levels and different use of contextual effects between groups. Consistently with previous studies [22], time processing abnormalities were equally reported in the various presentations of ADHD.

Time is not a unitary concept but encompasses many measurement scales, from a few milliseconds to several days. Even if there is not a defined boundary classifying a duration as "short" or "long", one of the most classic distinctions is that between subsecond and suprasecond stimuli. Clear dissociations have been previously found by pharmacological studies. [23] found that administering ethanol impaired auditory temporal discrimination thresholds for long (1s) but not shorter (50ms) intervals. A similar pharmacological dissociation was later found by Rammsayer [17] showing that auditory temporal processing of long durations (1s) was significantly impaired by administration of haloperidol (a dopamine receptor antagonist) and midazolam (benzodiazepine), whereas processing of extremely brief intervals (50ms) was only affected by haloperidol. The authors suggested that temporal processing of longer intervals is mediated by working-memory functions, while temporal processing of intervals in the range of milliseconds are more dependent on the effective level of dopaminergic activity in the basal ganglia. Psychophysical experiments also supported this differentiation. For example, it has been demonstrated that increasing cognitive load by dual-task procedures deteriorates discrimination thresholds for relatively long (1s), but not short (50ms) auditory stimuli, suggesting that the encoding of longer intervals requires cognitive control and working memory, while relatively shorter intervals are

automatically processed [15, 16]. Overall, these and many other results are difficult to explain, suggesting a unique system for timing perception.

By showing a specific impairment of auditory time thresholds for relatively short (0.5s) but not longer (1.5s) intervals in ADHD, the current results fit well with this idea, confirming different mechanisms for subsecond and suprasecond stimuli. The null correlation between thresholds across the two timing regimes is also consistent with this hypothesis. Given that ADHD is often associated with deficits on cognitive control and working memory, it is counterintuitive that timing thresholds for automatic, but not those under cognitive control were impaired in our sample of ADHD, suggesting a pure time perception deficit.

It is worth noting that our results are opposite to those found by [10]. In that study the authors measured auditory discrimination thresholds in controls and in children/adolescents with ADHD with short (550ms) and longer (4s) intervals. Results demonstrated worse thresholds for long but not short intervals.

Although we do not have a definitive explanation to account for this difference, the different tasks could, even partially, account for that. The paradigm used by Radonovich et al. presented two pairs of tones, one with fixed delay, the other with variable delays with participants asked to report whether the second delay was shorter or longer than the first. Our paradigm required the presentation and categorization of a single tone. We find it reasonable to speculate that the Radonovich et al. task, compared to the task used here, would charge relatively more additional resources such as working memory and/or attention, which may account for the different pattern of results.

Regarding brain areas involved in the perception of subsecond and suprasecond intervals, the literature, despite suggesting different networks, is not definitive. [24] compared auditory time discrimination thresholds between controls and patients with focal lesions in the frontal cortex or in the cerebellum. The results indicates that frontal lesions impaired timing thresholds for long (4s) but not short (0.4s) intervals, while cerebellar lesions impairing both. Harrington et al. [25], however, cast doubts on the involvement of the cerebellum in the auditory time perception of subsecond intervals (0.3s, 0.6s), with patients with cerebellar lesions showing similar thresholds compared to controls. On the other hand, cerebellar lesions have been demonstrated to have detrimental effects on visual time thresholds for suprasecond (8–21 s) stimuli [26]. As mentioned before, basal ganglia have also been found to play a role on time perception. Gouvêa et. al [27] trained rats to categorize sounds as belonging to a long or short category. Animals made few errors when categorizing the shortest and longest intervals (the extremes), but performance become worse for intervals near to the 1.5s categorical boundary (range: 0.6–2.4 s). Recording from populations of single striatal neurons, the authors found cells firing at different times within the interval period, suggesting short and long preferring neurons. The causal role of striatal neurons was also demonstrated by injecting muscimol. As a result, the duration thresholds worsened significantly, compared to the control group injected with saline. The parietal cortex has also been shown to play a critical role in time perception [28–31]. Hayashi et al. [32] correlated time discrimination thresholds measured in a sample of neurotypical adults, with gray matter (GM) volume in different parts of cortical and subcortical areas. Results showed that GM volume in the cerebellum but not in the parietal cortex

correlated with subsecond thresholds whilst, contrarily, GM volume in the parietal cortex but not in the cerebellum correlated with suprasecond thresholds. Moreover (as in the current study) threshold for subsecond and suprasecond stimuli did not correlate with each other. In the same line, a meta-analysis of imaging studies also suggests that the parietal cortex, compared to the cerebellum, is more likely activated by supra second stimuli [33]. With the current psychophysical data, we cannot say much on the brain networks involved in subsecond and supracesond timing, but our findings are largely in line with the idea that those two functions engage distinct neural mechanisms.

Another final interesting point emerges from the current results. The analysis of developmental trajectories (Fig. 3A) suggests that, rather than a generalized deficit across all ages, there could be a developmental delay for subsecond thresholds in ADHD. Despite having few participants in the higher age range, the results seem to suggest that above about 12/13 years old, the difference between groups would gradually decrease. Future studies might replicate this finding and expand the age range to quantitatively define the magnitude of such a possible developmental delay.

Conclusions

A precise subsecond timing perception is crucial in perceiving and acting on the continuously changing environment. It has been suggested that this ability is automatic and allows complex human behaviours including speech perception and speech performance, music, driving, and many sports. The results of this study showed that children and adolescents with ADHD have a severe precision (thresholds) deficit in the duration perception of subsecond auditory stimuli. Time perception for relatively longer and more cognitively controlled stimuli was unimpaired. Although, with the current results, we cannot explain why the temporal perception of short stimuli is impaired in our sample with ADHD, it is surprising how such a simple and fast (4-minute) psychophysical test succeeds in differentiating the two groups ($LBF = 1.9$). Moreover, while timing thresholds had a relatively low sensitivity power (60%), they turned out to have a good specificity (85%) level and were able to correctly classify 72.5% of cases. Finally, the timing thresholds deficit was resistant to the statistical controls of general but important covariates such as age, sex and non-verbal IQ, promoting it as a potential easy-to-administer tool for future studies. A better comprehension of time perception abnormalities in ADHD may give new insights in the neurological bases of the disorder. Furthermore, it may have relevant diagnostic and treatment implications. Understanding the functional consequences of these specific deficits in everyday life, in combination with other ADHD symptoms (i.e. impulsivity), may help to focus interventions on more specific goals, improving the clinical care of youth with ADHD.

References

1. Barkley, R.A., *Behavioral inhibition, sustained attention, and executive functions: constructing a unifying theory of ADHD*. Psychol Bull, 1997. **121**(1): p. 65-94.
2. Toplak, M.E., C. Dockstader, and R. Tannock, *Temporal information processing in ADHD: findings to date and new methods*. J Neurosci Methods, 2006. **151**(1): p. 15-29.

3. Zheng, Q., et al., *Time Perception Deficits in Children and Adolescents with ADHD: A Meta-analysis*. J Atten Disord, 2020: p. 1087054720978557.
4. Nejati, V. and S. Yazdani, *Time perception in children with attention deficit-hyperactivity disorder (ADHD): Does task matter? A meta-analysis study*. Child Neuropsychol, 2020. **26**(7): p. 900-916.
5. Smith, A., et al., *Evidence for a pure time perception deficit in children with ADHD*. J Child Psychol Psychiatry, 2002. **43**(4): p. 529-42.
6. Toplak, M.E. and R. Tannock, *Time perception: modality and duration effects in attention-deficit/hyperactivity disorder (ADHD)*. J Abnorm Child Psychol, 2005. **33**(5): p. 639-54.
7. Rubia, K., A. Smith, and E. Taylor, *Performance of children with attention deficit hyperactivity disorder (ADHD) on a test battery of impulsiveness*. Child Neuropsychol, 2007. **13**(3): p. 276-304.
8. Taş Dölek, G., et al., *Impaired auditory and visual time reproduction in adult patients with attention deficit-hyperactivity disorder*. J Clin Exp Neuropsychol, 2021. **43**(2): p. 176-186.
9. Plummer, C. and N. Humphrey, *Time perception in children with ADHD: the effects of task modality and duration*. Child Neuropsychol, 2009. **15**(2): p. 147-62.
10. Radonovich, K.J. and S.H. Mostofsky, *Duration judgments in children with ADHD suggest deficient utilization of temporal information rather than general impairment in timing*. Child Neuropsychol, 2004. **10**(3): p. 162-72.
11. Gooch, D., M. Snowling, and C. Hulme, *Time perception, phonological skills and executive function in children with dyslexia and/or ADHD symptoms*. J Child Psychol Psychiatry, 2011. **52**(2): p. 195-203.
12. Barkley, R.A., et al., *Executive functioning, temporal discounting, and sense of time in adolescents with attention deficit hyperactivity disorder (ADHD) and oppositional defiant disorder (ODD)*. J Abnorm Child Psychol, 2001. **29**(6): p. 541-56.
13. Bauermeister, J.J., et al., *Time estimation and performance on reproduction tasks in subtypes of children with attention deficit hyperactivity disorder*. J Clin Child Adolesc Psychol, 2005. **34**(1): p. 151-62.
14. Meaux, J.B. and J.J. Chelonis, *Time perception differences in children with and without ADHD*. J Pediatr Health Care, 2003. **17**(2): p. 64-71.
15. Rammsayer, T.H. and R. Ulrich, *Elaborative rehearsal of nontemporal information interferes with temporal processing of durations in the range of seconds but not milliseconds*. Acta Psychol (Amst), 2011. **137**(1): p. 127-33.
16. Rammsayer, T.H. and S.D. Lima, *Duration discrimination of filled and empty auditory intervals: cognitive and perceptual factors*. Percept Psychophys, 1991. **50**(6): p. 565-74.
17. Rammsayer, T.H., *Neuropharmacological evidence for different timing mechanisms in humans*. Q J Exp Psychol B, 1999. **52**(3): p. 273-86.
18. Ptacek, R., et al., *Clinical Implications of the Perception of Time in Attention Deficit Hyperactivity Disorder (ADHD): A Review*. Med Sci Monit, 2019. **25**: p. 3918-3924.

19. Wechsler, D., ed. *WIS-IV Wechsler Intelligence Scale for Children Fourth edition*, Giunti OS editor. 2012.
20. Anobile, G., et al., *Numerosity but not texture-density discrimination correlates with math ability in children*. Dev Psychol, 2016. **52**(8): p. 1206-16.
21. Efron, B. and R.J. Tibshirani, *An introduction to the bootstrap* (Vol. 57) New York, NY: Chapman & Hall. Vol. 57. 1993.
22. Valko, L., et al., *Time processing in children and adults with ADHD*. J Neural Transm (Vienna), 2010. **117**(10): p. 1213-28.
23. Rammsayer, T.H. and W.H. Vogel, *Pharmacologic properties of the internal clock underlying time perception in humans*. Neuropsychobiology, 1992. **26**(1-2): p. 71-80.
24. Mangels, J.A., R.B. Ivry, and N. Shimizu, *Dissociable contributions of the prefrontal and neocerebellar cortex to time perception*. Brain Res Cogn Brain Res, 1998. **7**(1): p. 15-39.
25. Harrington, D.L., et al., *Does the representation of time depend on the cerebellum? Effect of cerebellar stroke*. Brain, 2004. **127**(Pt 3): p. 561-74.
26. Malapani, C., et al., *Cerebellar dysfunctions of temporal processing in the seconds range in humans*. Neuroreport, 1998. **9**(17): p. 3907-12.
27. Gouvêa, T.S., et al., *Striatal dynamics explain duration judgments*. Elife, 2015. **4**.
28. Leon, M.I. and M.N. Shadlen, *Representation of time by neurons in the posterior parietal cortex of the macaque*. Neuron, 2003. **38**(2): p. 317-27.
29. Bueti, D. and V. Walsh, *The parietal cortex and the representation of time, space, number and other magnitudes*. Philos Trans R Soc Lond B Biol Sci, 2009. **364**(1525): p. 1831-40.
30. Bueti, D., et al., *Learning about time: plastic changes and interindividual brain differences*. Neuron, 2012. **75**(4): p. 725-37.
31. Harvey, B.M., et al., *A Network of Topographic Maps in Human Association Cortex Hierarchically Transforms Visual Timing-Selective Responses*. Curr Biol, 2020. **30**(8): p. 1424-1434.e6.
32. Hayashi, M.J., et al., *Dissociable neuroanatomical correlates of subsecond and suprasecond time perception*. J Cogn Neurosci, 2014. **26**(8): p. 1685-93.
33. Lewis, P.A. and R.C. Miall, *Distinct systems for automatic and cognitively controlled time measurement: evidence from neuroimaging*. Curr Opin Neurobiol, 2003. **13**(2): p. 250-5.

Figures

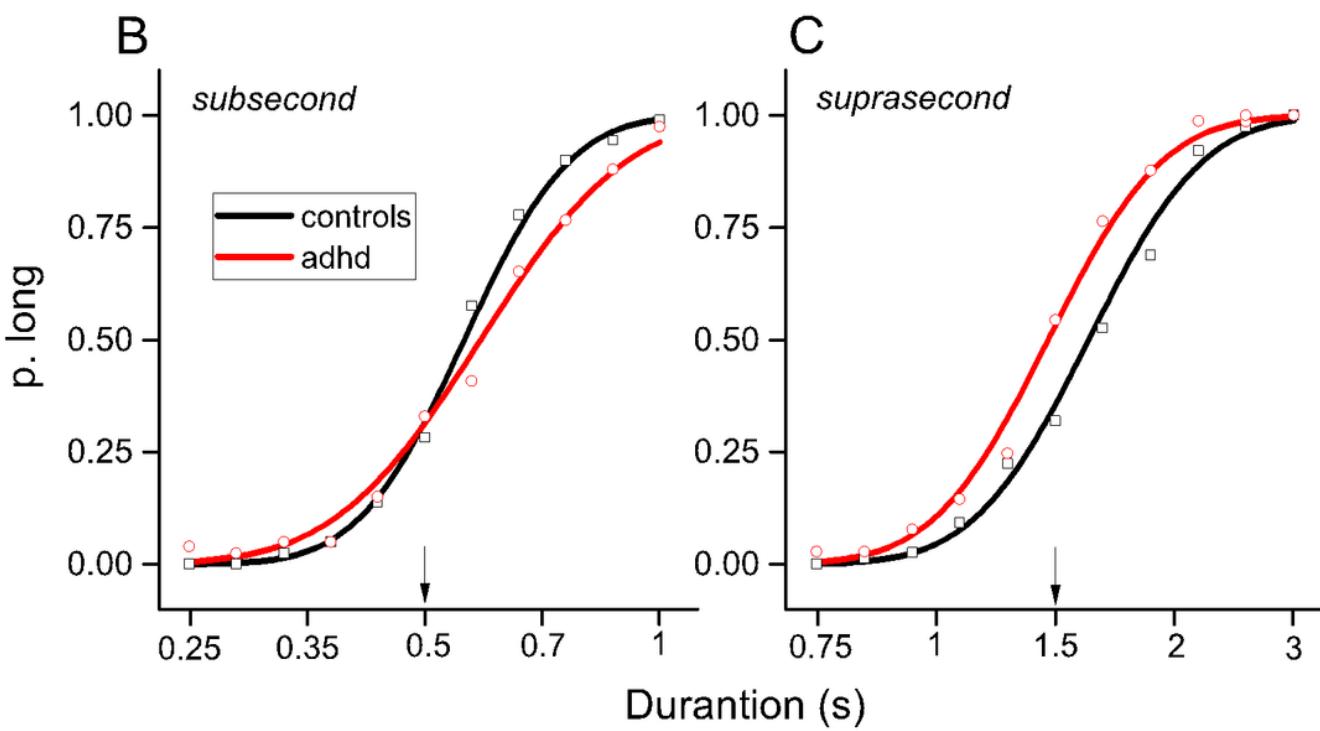
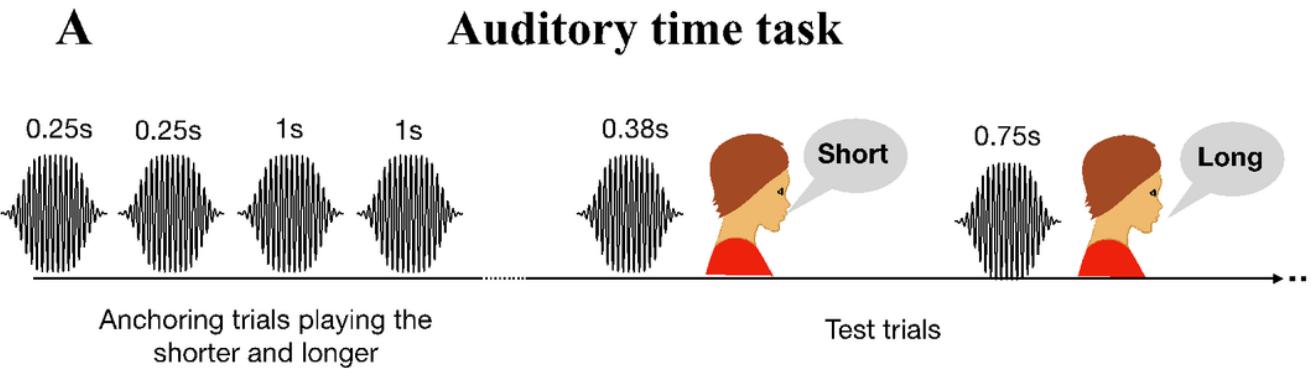


Figure 1

A) A block consisted of four initial “anchoring” trials in which the shorter (0.25 s or 0.75 s depending on the condition) and the longer (1 s or 3 s depending on the condition) stimuli were presented (participants were told that those sounds correspond to the range extremes). After this phase, the testing phase started and participants were asked to categorize as “long or short” a sound randomly drawn from a pre-defined distribution. B-C) Psychometric functions (aggregate data) for controls (black, squares) and participants with ADHD (red, circles) for subsecond (B) and suprasecond (C) ranges.

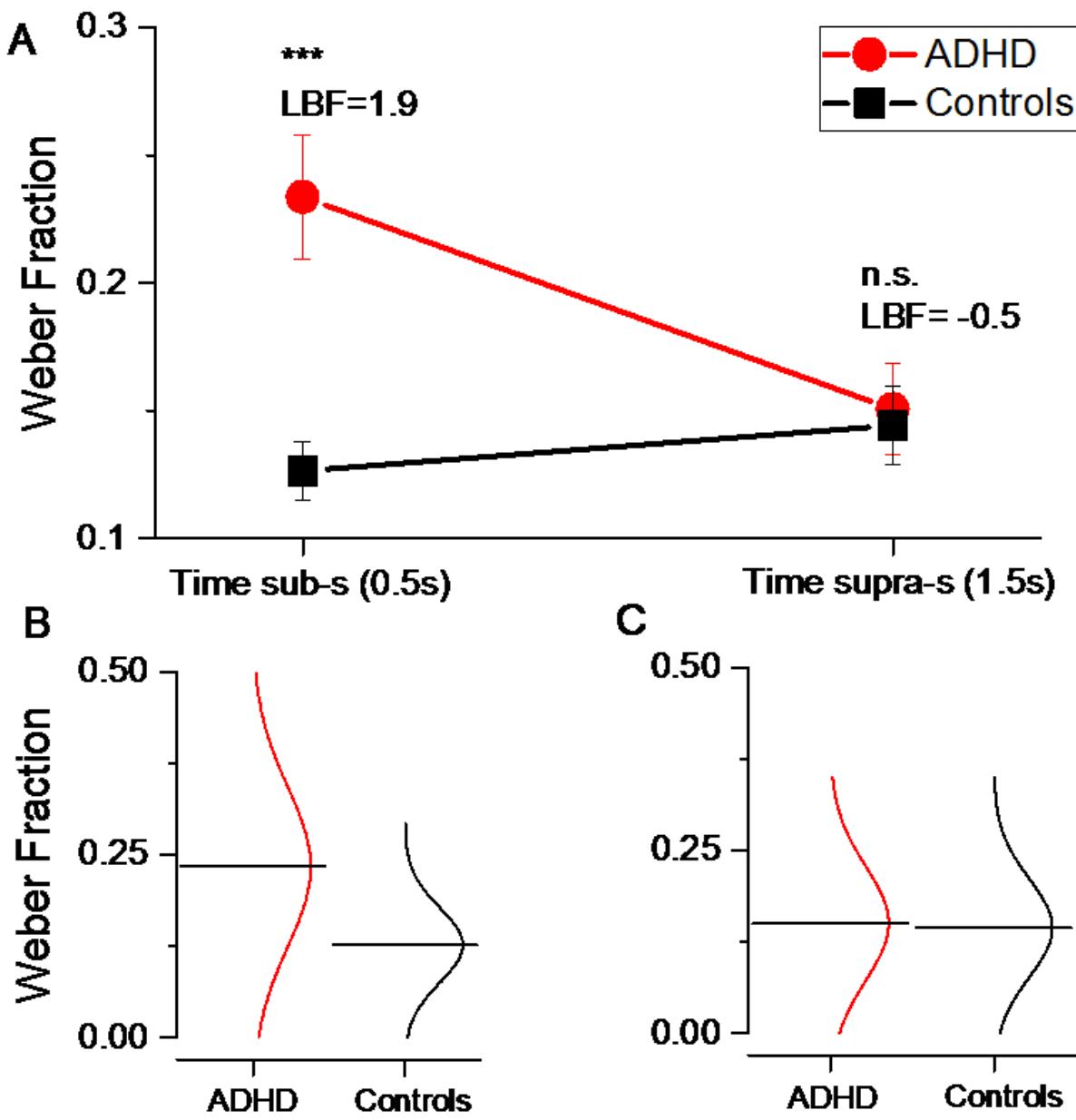


Figure 2

A) Between participants average time thresholds divided by the two groups (ADHD: red circles and Controls: black squares). Error bars are standard errors of the mean; *** $p<0.001$; n.s. not statistically different. B-D) Individual data reporting time thresholds distributions. Horizontal lines report data average.

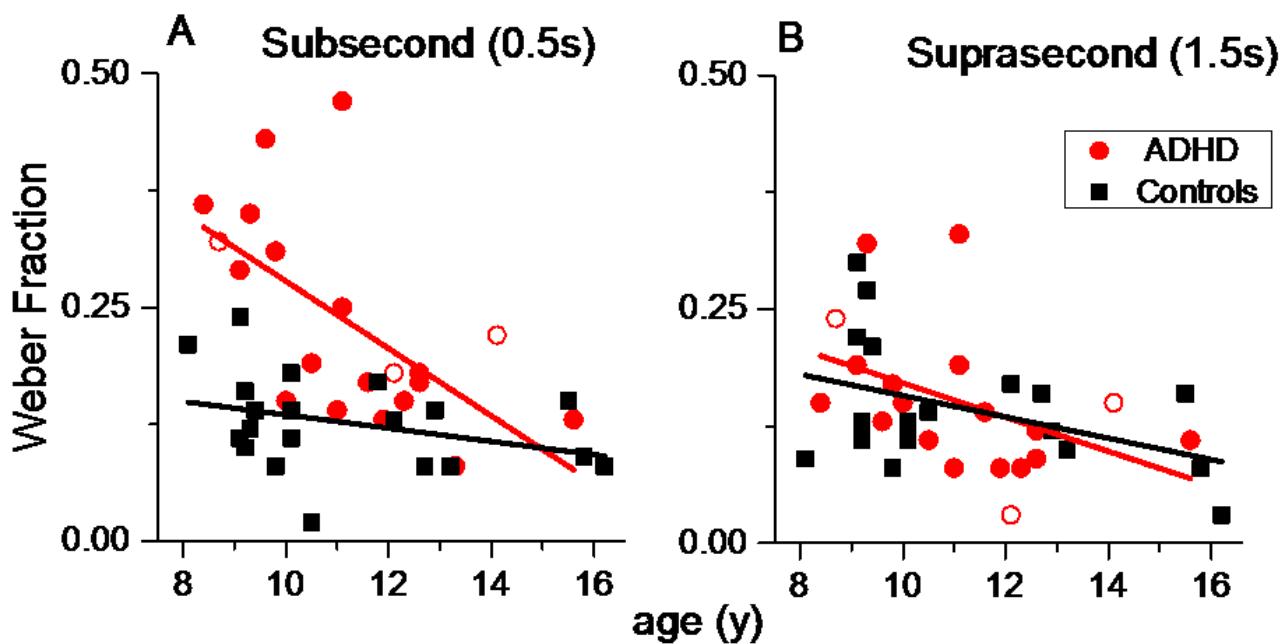


Figure 3

Auditory time thresholds for stimuli in the subsecond (A) and suprasecond (B) ranges as a function of age divided by the two groups (ADHD: red filled circles, ADHD+dyslexia: red open circles, Controls: black squares).

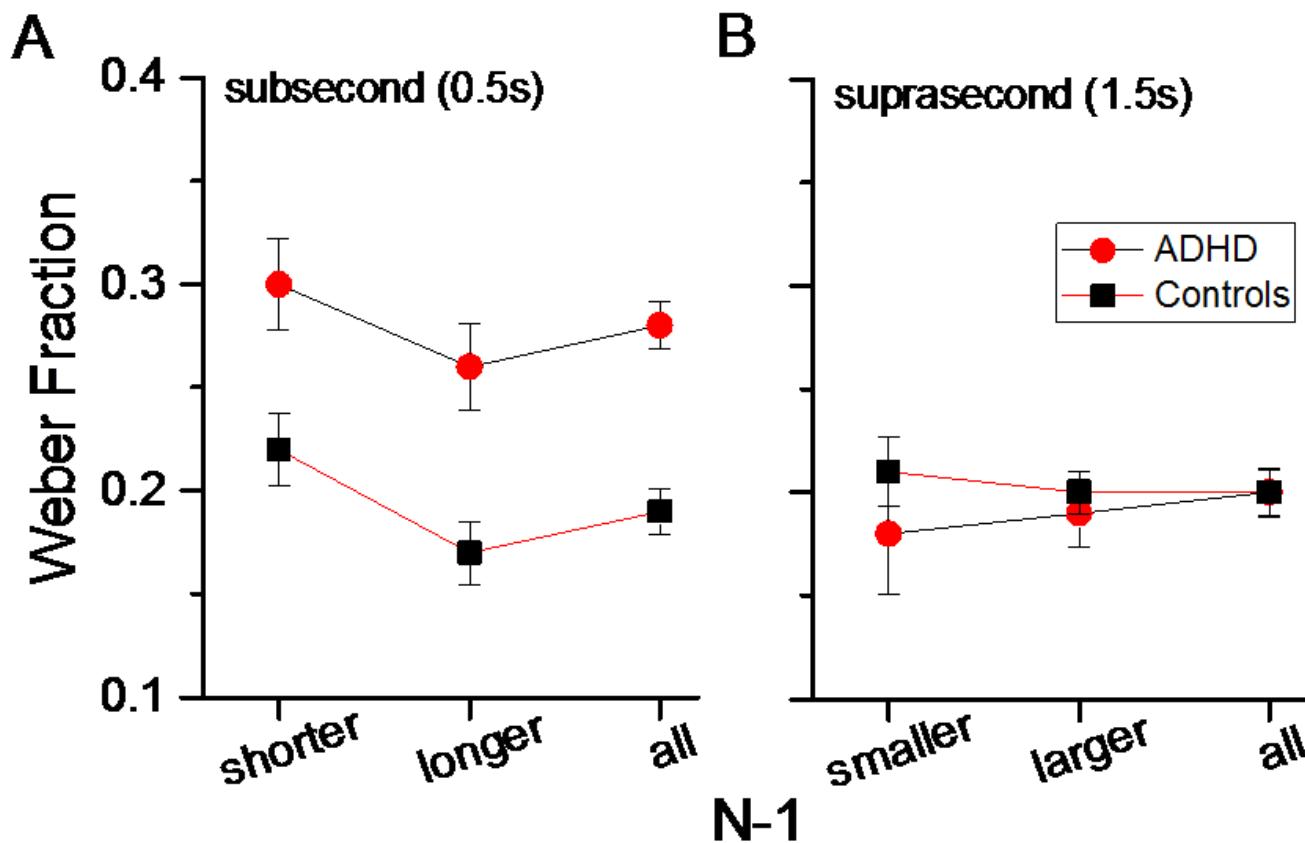


Figure 4

Auditory time thresholds for stimuli in the subsecond (A) and suprasecond (B) range measured on all trials (all) or on data sorted as a function of the preceding stimulus duration ($N-1$) that could be shorter or longer compared to the stimulus judged in the current trial.