Attention-Deficit Hyperactivity Disorder (ADHD) in Children: A Move towards Developmental Perspectives

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Abstract

The debate about diagnoses and treatment of attention deficit hyperactive disorder (ADHD) in children continue to range on between the developmental and biological perspectives. While there is increasing evidence that support the biological susceptibility of ADHD, a number of researches also emphasized the significant effect of environmental factors on the syndrome. This study used developmental perspectives to evaluate and bring together various bio-psychosocial factors that impact on development of children diagnosed with ADHD. To achieve this the study critically explored and integrated the existing and advancing study on ADHD to a more refined pattern that embraced developmental perspectives by organizing into sections; the clinical and social factors that associated with children diagnosed with ADHD. Also the study discussed how linkage in childhood ADHD fits within a developmental psychopathology perspective. Finally, the study revealed ADHD as a developmental disorder influenced by prenatal, biological and psychosocial environmental risk factors, and that better understanding of genomic susceptibilities, family environment, parental characteristics, and children's experiences can transform the pathway for its development in children.

Keywords: Attention deficit hyperactive disorder; Developmental perspectives; Childhood disorder; Genetic factors; Environmental factors

Introduction

Attention-deficit/hyperactivity disorder (ADHD) is a severe childhood disorder that affects many facet of human being, particularly young children populace (American Psychiatric Association) [1] and has been a subject of intensive research for decades [2]. While studies over the years demonstrated the advancement made on the syndrome, the intense interest in the disorder continue to produce a number of empirical data, such as on etiological factors, complex genetic and the neurobiological variables that underline the symptom particularly, its developmental cause and treatments in children diagnosed with ADHD. For example, studies like molecular and behaviour, have long offered considerable suggestions to support the significant effect of genetic factor on the symptoms [3,4]. Additionally, a quite number of models have long proposed to address the disorder in children, particularly on cognitive deficiency [5,6]. However, contrary to the progress on bio-cognitive development, theory on social and relational features of ADHD remains stagnant, as the general consensus on the disorder showed multiple casual pathways, with environmental factors primarily labelled as ameliorating the symptom in children [7].

While the research on family of children with ADHD has long been acknowledged [7], the debates about ADHD, particularly, its occurrence and origin in children continues to range on. More disturbing is the fact that developmental conceptualization of the syndrome has been neglected. Specifically, the clinical and social issues associated with the symptom have waned, to say the least, downplayed in most literature. This made it hard for practitioners and families of children diagnosed with ADHD to cope with the disorder, and projected the problems and discriminations experienced by families and children diagnosed of ADHD an important issue for consideration. Therefore, much is desire on the impact of gene-environment interactions on socio-cognitive development of children diagnosed of ADHD.

Objectives of the Study

This study provides a brief overview of the social and clinical issues associated with children diagnose of ADHD, and uses developmental perspectives to evaluate and bring together various bio-psychosocial factors that impact on children's development [8]. To achieve these objectives, the study critically examines and integrates the existing and advancing study on the ADHD to a more refined pattern that embraced developmental perspectives. The study also organized into sections, the clinical and social factors that relates to childhood ADHD and explain how these factors influenced children's development. Finally, the study discusses how linkage in childhood ADHD fits within a developmental psychopathology perspective and makes recommendations for future research.

Scientific Status of Attention Deficiency Hyperactive

ADHD is multidimensional disorder that exacts a significant effect on individual and society. This disorder has negative impact on families, as well as academic and vocational outcomes of vulnerable children [9]. As the most generally diagnosed neurobehavioral illness in children, the disorder is mostly treated with stimulant and non-stimulant drugs [10,11]. Even though the exact causes of the syndrome in children are still unknown, past and present research linked its origin to genetic and environmental factor [12,13]. Besides, research on ADHD emphasised more on the period of birth by establishing a strong correlation between the period of birth and children's psychological and behavioural disorders [14]. However, this is contrast...
to several other disorders where a reliable seasonal form is yet to be established [15,16].

As an unsatisfactory umbrella term, ADHD is apply to children with broadly differing temperaments and functional problems in school, home, and social settings. This group of children shared certain core features, such as limited sustained attention span, poor impulse control, and motor over activity. The children also developed abnormal syndromes, such as severe development, distraction and thoughtlessness that cause impairment in learning [16]. Research also established that ADHD has a strong genetic orientation. This means that the inattentiveness aspect of the disorder is related to fantasizing, distractibility, and problem with concentration on specific task for a lengthy period, while the hyperactivity element is pronounced as fidgeting, unnecessary talking, and restiveness [17]. Further, the signs of the disorder are also predisposed to accidents, strain interpersonal relationships, disruptions and improper conduct. However, apart from associated with clinically oriented disorder in children, ADHD also linked to characteristics in adulthood, such as drugs and alcohol misuse; socio-cognitive disorders; disruptive conduct and delinquency [18].

Despite the above illustrations and evidences on ADHD, the developments of the syndrome remain debatable. For instance, the symptom is regarded as multidimensional, and was generally linked to children and adults mental health [18]. This point to the fact that genetic factors contributed significantly to its development, and also its relations to ecological risk variables are complex. Based on this foregoing, there is a need to ponder on the evolving nature of the symptom and the differences in the phenotypic indicator, particularly, the influence of ecological factors on development of children with ADHD [18].

**Diagnosis Consideration of ADHD**

Research in the last 60 years has witnessed the use of several terminologies to describe attention deficit hyperactivity disorder (ADHD). Some of this terminology includes: hyperkinetic impulse disorder, minimal brain dysfunction, hyperactivity, attention deficit disorder. Moreover, the core characteristics of the ADHD are inattention, impulsivity, and hyperactivity. The syndrome affects about 4% of all children, and is more noticeable among young people [19,20]. Despite its occurrence in children, the origin of the disorder is yet to be identified. Thus, the differences in expression revealed the diverse conceptions of the primary symptoms and the assumed fundamental pathophysiology of the disorder.

Literature revealed that, the prospect of finding a diagnostic indicator for ADHD is not achievable. This is due in part, to the nature and complexity of the syndrome [21,22]. Research identified three subtypes of ADHD and each of this subtypes were differs on symptomatology. Therefore, for a child to be diagnosed of ADHD, and labelled with particular subtypes, he/she must display 6 symptoms for a period of 6 months. While achieving such diagnostic criteria is difficult, this method is used as a bench mark for diagnosed the disorder in children. Further, children diagnosed with ADHD also show some degree of functional impairment in multiple settings [19,23]. However, due to the parallel characteristics of the disorder, the comorbidities, such as anxiety disorders and ODD influenced its subtype in children.

Although the criteria listed in DSM-V for ADHD is more or less broader over DSM-IV-TR, the issue of sex differences in children hyperactive disorder continue to range on [19,1]. For instance, male child are 3 times potential of having ADHD and display hyperactive behaviour or combination of it, than female child [20]. Also, females are more expected to display predominantly absent-minded subtype and suffer from mental impairment and eating disorders [24]. Further, there is higher sense of aggressiveness and abuse of law among male than female children diagnosed of ADHD [24]. On the basis of this, there is a need for professionals working with children diagnosed of hyperactive disorder to be consciously aware of its sexual and developmental variances. This if observed, would prevent over-or under diagnosed of the disorder in children. Furthermore, there should be proper analysis and assessment of parents and teachers reports, so as not to mislabel the underline disorders [25,26].

**Etiological Model of ADHD**

While research established the main aetiology of ADHD as unknown [27]. It is also important to understand the aetiology and other associated disorders that relates with disorder. This understanding would assist in identifying the interactions between the genetic and environmental factors and its vulnerability in young children. The process also offered an understanding of the heterogeneity of the disorder in a meaningful manner, as research showed lack of systematic incorporation of the findings across multiple levels of analysis [28,27]. Etiological models of ADHD also emphasised the following: the impacts of genetic and environment factors; their correlations and interactions; its influence on brain composition and function, and the mediating role of the symptom expression. Based on this aforementioned, more investigation is needed in order to create a clear relationships between supposed fundamental genetic and neural processes, and behavioural manifestations of the disorder. This process would increase and encourage new and effective treatments (biological and non-biological), and offer necessary information on the framework that supports the management of ADHD particularly, in hypothesising, diagnostic of boundaries and the current arrangement of the illness.

However, the hypotheses for reducing brain function in children diagnosed of ADHD were grounded on several observations that reduced the volume of gray and white matter in the brain. This causes shortfalls in cognitive processing, responsiveness, motor planning, speed of processing responses, and other related behaviour in children diagnosed of the disorder [29]. In addition, prefrontal cortex (PFC), caudate, and cerebellum were the primary source of shortfalls in children diagnosed of hyperactive disorder. This was formed by different neurons that together regulate attention, thoughts, emotions, behaviour, and negative actions in children [30,31]. Also, poor development of PFC11 reduced the activity of the PFC, caudate, or cerebellum [30]. The system activity between the regions is subtle to the neurochemical environs,” [30] and sustained by the combination of neurotransmitters (NTs), dopamine (DA), norepinephrine (NE) and multiple receptors [32,33].

Etiological model also identified aggressiveness, impairment and other related problems, (i.e. antisocial conduct) as the key goals of childhood ADHD. Though, medication was identified as a way of reducing hyperactive disorder in children, the long-term supports for the broader outcomes of ADHD are yet to be established. These underscore the importance of identifying the genetic-environment factors that caused the negativity and impairment in children with ADHD, and provide answer to the growth of active risk decline tactics in the long-term management of the disorder. Based on this
aftementioned, it is imperative for research to focus on understanding the genetic and environmental risk factors that associated with ADHD, as well as the clinical characteristic that project the outcomes of the disorder in children, as this would target resources and monitor children at risk of adverse concerns.

The Need for a New Model on ADHD

A decade of scientific study on ADHD has highlighted the need for a new theory that explains the syndrome. ADHD is confirmed as a disorder particularly, in respects to its basic nature. Most research on the ADHD is more or less investigative and descriptive, with exception of two. First, the work of Quay’s [34-36] who used the neuropsychological model of anxiety developed by Gray’s [37]. To describe the source of the poor inhibition manifested in ADHD. This model relates thoughtlessness to under-functioning of the brain’s behavioural inhibition system. The model also explained children with hyperactive disorder as subtle to the signs of conditioned punishment, and less sensitive to passive avoidance models [35]. However, the second model failed in its attempt to set up a concept similar to the one established in Quay-Gray theory. The model makes a comprehensive theory construction that offers coalescing explanation on various mental shortcomings that are related to children diagnosed with attention/ hyperactive disorder.

The Developmental Approach

The desire for a theory that embraced the clinical and social aspects of attention/hyperactive disorder has prompted the need for developmental approach to ADHD. Although a comprehensive neuropsychological model of ADHD has yet to be proposed, other models of psychopathologies was previously recommended [38]. Developmental approach entails the correlation between the etiological heterogeneity, high level of comorbidity, and biological and psychosocial/family of ADHD [27,39]. These interactions underscore the need to posit a multiple developmental pathways to treatment of children diagnosed of ADHD [27,28] and were mediated by a variety of within child and family contextual factors that associates with either the diminution or exacerbation of the symptoms over time.

For example, dynamic developmental psychopathology approach offers an explanation on how attention/hyperactive disorder evolved, and how the interactions between multiple risk and protective factors impact on children development [8]. The model proposed that some children in the course of their development were influenced by biological risk factors, with a relatively lesser impact from the ecological factors. Also, the model highlighted that across children and across time, there are variables that influence the development of attention/hyperactive disorder. The theory predicted that while precise symptom of ADHD at a particular time in life varies, they are influenced by factors that have positive or negative effects on the symptom development. The theory also explained that, individual differences in dopamine functioning have significant impact on motor functions and children learning and produced behaviours, such as attention problems, hyperactivity, and impulsiveness that associated with ADHD, and predict an increase in children’s behavioral variability. Overall, dynamic developmental theory proffer better explanation on how person predispositions interacted with the above mentioned conditions and relatively created behavioral, emotional, and cognitive effects that balanced the behavioral patterns in children with ADHD. Thus, a child’s characteristics coupled with the family situation exerted collaborating influence on ADHD and offered unique opportunity for analysing the disorder symptomatology.

Psychosocial Adversity and its Developmental Course

Though, many studies have proposed significant evidence for the existence of psychosocial problems in children with hyperactive disorder, such evidences predicts the socio-cognitive and emotional development, rather than precise predictors of the disorder. Therefore, it remains uncertain whether experience of violence in infancy is a risk factor for ADHD, as there was no theoretical basis for observing this possible relationship. For example, exposure to violence in a household may act through psychosocial adversity and lead to permanent brain change that occurs as a result of prolonged exposure of the developing brain to steroid hormones [40].

However, Rutter et al. [41] reported that the combination of environmental factors (i.e., severe marital discord, low social class, paternal criminality, maternal mental disorder), rather than existence of a single factor, promote psychopathology in children. This argument was supported by a lot of scholars, such as Campbell; Faraone and Biederman; Rutter and Sroufe; and Taylor [8,42-44] as they linked genes-environment multiple interactions to attention/ hyperactive disorder in children. Similar findings by Biederman et al. [45] corroborated earlier work by Rutter and his colleagues to establish negative family-environment as significantly influenced children with ADHD. The finding also established that exposure to parental psychopathology (particularly maternal) is more pertinent to families of children with ADHD than the control families [45].

However, while some studies in the field of developmental approach established that children are born with a genetic predisposition that relates to hyperactive disorder [46], others maintained that heredities are rarely the sole reason for the development of attention/ hyperactive disorder, as MZ concordance rates is not near 100% [3,47]. For example, some scholars maintained that 50% of children with hyperactive disorder do not display the biological anomaly associated with congenital factors [48]. However, in situation where biological predisposition is strongly established, family characteristics was viewed as reflection of the indicator and consequence of the disorder in children.

Also, the categorization of relative contributions of shared versus non-shared hereditary and ecological menaces within the families of children diagnosed of ADHD is important for proper analysis of the disorder. For example, in a situation where there is a problem in a family, which is due to the disorder, or shared genetic susceptibilities, the family environments must be related to the child characteristics. On the other hand, when family breakdown is linked to the child empathy, the constancy of the disorder became aggravated. In this case, the family environment is associated with attention/hyperactive disorder not as a main cause, but as a factor that increased and influenced its development. Additionally, children with ADHD develops relatively little tendency to the disorder, as confusing and uncaring family setting increase their behaviours [49]. This means that, the degree of intellectual and physical stimulation that children received in their immediate environment impacted on their brain development and behaviour [50]. Therefore, a responsive and sensitive parenting would promote child self-regulation skills and parental difficulties that harmonize parents’ activities with child’s desires for development of disinhibited behaviour [49]. So, when the family and child characteristics work in tandem, child’s temperament antecedents

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of inattentiveness and impulsivity that create or exacerbate parents’ problems are moderated.

**Genetic Contributions to ADHD and Developmental Course**

ADHD is not a genetic disorder in a clear sense, but can be categorised as a genetic factor that was shaped by developmental pathways [51]. While past and present research continues to highlights the importance of genetic factors on ADHD [52,53], attempts to recognize its source of using a candidate gene method to detect common hereditary variant have been less successful [54]. Thus, genetic explanations of ADHD are determined by data, such as family and twin studies that shows ADHD as a familial and highly hereditary. This heritability was estimated to be in average of 76% [17]. While it was long established that attention/hyperactive disorder is a family oriented symptom, the first-degree families of affected persons displayed higher rates of the disorder (relative risk 4-5). Therefore, the threats of the disorder are higher in families of those with history of hyperactive disorder [17]. This highlighted the significant agreement between early studies of children diagnosed with hyperactive syndrome (Morrison and Stewart) successive studies that uses DSM-III and DSM-III-R definitions of ADHD [55].

For instance, a meta-analysis study conducted by Faraone et al. [17] revealed a small but significant impacts for a number of assumed functional variants in genes controlling brain neurochemistry particularly, in the dopamine system (e.g., D4 and the dopamine transporter (DAT1)). The common variants in genes of other neuromodulator systems (i.e., serotonin and norepinephrine) was also related with genes that control the general brain function and growth [56,57].

Also, the analysis of comorbid psychiatric disorders supported the inherent heterogeneity of the ADHD in children. This established a significant degree of ADHD among families of adults with ADHD [58]. For example, the independent samples of children with DSM-III attention-deficit disorder and DSM-III-R ADHD are related to familial susceptibilities [55,59,60], while attention/hyperactive disorder and bipolar conditions was established as a separate familial subtype of ADHD in children [61]. Attention/hyperactive disorder were also found to be familial free from anxiety disorders and learning disabilities [62]. Based on this foregoing, one can conclude by suggest that stratification by behaviour and bipolar disorders divides the life of children diagnosed of ADHD into more familial related subgroups, and that major depressive disorder is a generic expression of different subtypes of ADHD in children. Persistent attention/hyperactive disorder are a useful phenotype for molecular genetic studies [52]. However, despite the inaccessible findings in literature on ADHD, individual gene relationships account for modest variation in children diagnosed of ADHD [17,54].

**Twin and Adoption Studies**

Due to the genetic nature of ADHD, twins studies are consistently used to establish the heritability, or the level at which genetic characteristics influence attention/hyperactive disorder [63-65]. The studies also offered a reliable evidence to support that, hereditary factors add to the aetiology of ADHD i.e., (60-91%) [51]. Twin studies confirmed that inherited factors are the main source of continuousness of attention/ hyperactive disorder particularly, the relationship between the disorder and disruptive behaviour [18] and revealed that inherited factors impacted on ADHD and its developmental progression. Also, research on twins and adoption studies established extra risk factors that do not have any significant influence on the origin of ADHD, but contributed to its clinical developmental outcomes. However, this notion was condemned because children genetic factor are chosen with a priori notion of genetic involvement in the syndrome, while in neuropsychiatric illnesses, the pathophysiology is typically unidentified.

**Biological Adversity**

Research suggested that some biologic factors, such as lead contamination, food additives/diet, cigarette and alcohol exposure, to mention a few, contributed to the development of attention/ hyperactive disorder in children. Though, Feingold Diet for ADHD was promoted by the media and acknowledged by most parents as a contributing factor, scientific enquiry showed that the idea is ineffectual, as addictiveness to food cannot cause attention/hyperactive disorder. Also, research argued that exposure to lead pollution causes restlessness, hyperactivity, distractibility, and lower intellectual ability in children diagnosed with hyperactive disorder. This idea was opposed by other studies that established that lead account for only few of the majority of ADHD issues in children. This means that exposure to high lead environment does not necessary lead to hyperactive disorder in children. Research also identified complications during pregnancy and delivery (i.e., maternal age, poor maternal health, and duration of labour) to mention a few as influenced development of ADHD in children [66] and confirmed maternal smoking as related to the pathophysiology of ADHD and causes disruption to nicotinic receptors and change dopaminergic activity.

**Gene-Environment Interaction and ADHD**

Though, studies on children with ADHD revealed a significant relationship between heredity and attention/hyperactive disorder, there are quite a number of environmental factors that connected with ADHD symptoms. Two of these factors, i.e. exposure to maternal smoking in pregnancy [67], and low birth weight/prematurity [68] have been systematically analysed and reported as contributed to the development of ADHD. However, not all the vulnerable children that are exposed to environmental severity developed attention hyperactive disorder. For example, effects of gene-environment interaction on ADHD occurs when genes responds to environmental adversity. This is documented as important features of attention/hyperactive disorder in children.

However, only few works have probe the influence of G6E on development of children hyperactive disorder. For instance, a recent research on the issue established strong link between a DAT1 haplotype (combination of risk alleles) and attention/hyperactive disorder when mother is alcoholic during pregnancy [69], while others studies reported the DAT1 risk allele earlier found to be related with attention/hyperactive disorder as associated with hyperactive-impulsive symptoms found in children exposed to maternal smoking during pregnancy. Furthermore, studies that focus on childhood behavioural disorder symptoms reported that children who carried the COMT gene risk variant are more vulnerable to the negative effects of lower birth weight [51]. While all these findings require replication, the indication so far showed that, some genetic factors influenced children sensitivity to ecological adversity and the developmental sequence of attention/hyperactive disorder.
Discussion

This systematic review used developmental perspectives to address the clinical and social factors associated with children diagnosed of ADHD. Specifically, it demonstrates that gene-environment interactions are important factors in the development of attention/hyperactive disorder in children. By focusing on developmental perspective, this paper provided considerable evidence to support the influence of bio-psychosocial factors on behaviour of children diagnosed with attention/hyperactive disorder. Also, the study supported the growing body of research that emphasised the use of developmental perspective as opposed to clinical treatment of children with attention/hyperactive disorder. Further, the study charted a developmental framework as bases for conceptualizing the effect of gene-environment interaction on children diagnosed with ADHD, and reviewed the consequences and limitations of existing studies on the symptom by exemplifying the areas where untimely deductions have been obtained and where further effort is desirable.

In addition, the study established parent-child interactions and gene-environment interaction as impacted on the development of children with attention/hyperactive disorder. This shows that, the stressful demanding and intrusive nature of children diagnosed with attention/hyperactive disorder evoked negative reactions from other family members and disrupted family relationships. The review of literature in this present study also revealed that children with attention/hyperactive disorder influenced their parent's behaviour and adjustment, and that parent's behaviour impacted on development of children diagnosed with the disorder [69]. This confirmed family characteristics and histories as the cause of attention/hyperactive disorder in children, as parent behaviour was linked to children conduct problems [18].

Overall, there is a general concession about the continuum association between genetic and environmental factors in children diagnosed with ADHD, as family factors were mentioned as the most influential variable that promote attention/hyperactive disorder in children. Though, the number of unsupportive or inconclusive studies actually limits these conclusions, this present review motivated research and hastens full informed conclusions about the clinical and social factors associated with children diagnosed of ADHD. The study also indicates that the dynamism of social and biological variables in children diagnosed with attention/hyperactive disorder is not only influenced by environmental factors, but by common genetic characteristics of the parent and the child [58].

Conclusion and Recommendation

Though, attention deficit hyperactive disorder is a predominant neurobehavioral illness in children, the symptom is characterised by factors such as hereditary, ecological, and biologic aetiologies that begin from conception to adulthood. Although its aetiology remains indeterminate, the developing evidence on the symptom documented its strong neurobiological and hereditary foundations and emphasised the phenotypic difficulty of disorder on children development. Therefore, there is a need to understand how genomic susceptibilities, family environment, parental characteristics, and children's experiences interrelate and modify its developmental pathway in children; as such efforts would prospectively enlighten and proffers intervention strategy that support its diagnose. Based on these assumptions, the following recommendations are suggested:

| References |
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| 1. Powers AD, Gleason ME, Oltmanns TF (2013) Symptoms of borderline personality disorder predict interpersonal (but not independent) stressful life events in a community sample of older adults. J Abnorm Psychol 122: 469-474. |
| 2. Barkley RA (2006) Attention deficit/hyperactivity disorder: A handbook for diagnosis and treatment (3rds ed). New York: Guilford. |
| 3. Kuntsi J, Stevenson J (2000) Hyperactivity in children: a focus on genetic research and psychological theories. Clin Child Fam Psychol Rev 3: 1-23. |
| 4. Sunohara GA, Roberts W, Malone M, Schachar R, Tannock R, et al. (2000) Linkage of the dopamine D4 receptor gene and attention-deficit/hyperactivity disorder. Journal of the American Academy. |
| 5. Berger A, Posner MI (2000) Pathologies of brain attentional networks. Neurosci Biobehav Rev 24: 3-5. |
| 6. Livesey D, Keen J, Rouse J, White F (2006) The relationship between measures of executive function, motor performance and externalising behaviour in 5- and 6-year-old children. Hum Mov Sci 25: 59-64. |
| 7. Sonuga-Barke EJ, Auerbach J, Campbell SB, Daley D, Thompson M (2005) Preschool varieties of hyperactive and dysregulated behaviour: Multiple pathways between risk and disorder. Developmental Science 8: 141-150. |
| 8. Rutter M, Sroufe LA (2000) Developmental psychopathology: concepts and challenges. Dev Psychopathol 12: 265-296. |
| 9. Biederman J, Faraone SV, Monuteaux M, Spencer T, Winters T, et al. (2004) Gender effects of attention deficit hyperactivity disorder in adults, revisited. Biol Psychiatry 55: 692-700. |
| 10. United States, 2003 and 2007 (2010) Increasing prevalence of parent-reported attention-deficit/hyperactivity disorder among children, MMWR Morb Mortal Wkly Rep 59: 1349-1443. |
| 11. Pastor PN, Reuben CA (2008) Diagnosed attention deficit hyperactivity disorder and learning disability: United States, 2004-2006. Vital Health Stat 10: 1-14. |
| 12. Nigg J, Nikolas M, Burt SA (2010) Measured gene-by-environment interaction in relation to attention-deficit/hyperactivity disorder. J Am Acad Child Adolescence Psychiatry 49: 863-73. |
behavioral aggression and cognitive impulsivity in children with attention-deficit/hyperactivity disorder (ADHD): Findings from a family-based association test (FBAT) analysis. Behavioral and Brain Functions 4: 4-48.

57. Biederman J, Milberger S, Faraone SV, Kiely K, Guite J, et al. (1995) Familienvironment risk factors for attention deficit hyperactivity disorder: A test of Rutter's indicators of adversity. Archives of General Psychiatry 52: 464-470.

58. Biederman J, Faraone SV, Keenan K, Tsuang MT (1991) Evidence of familial association between attention deficit disorder and major affective disorders. Arch Gen Psychiatry 48: 633-642.

59. Biederman J, Faraone SV, Keenan K, Benjamin J, Krifcher B, Moore C, et al. (1992) Further evidence for family-genetic risk factors in attention deficit/hyperactivity disorder. Patterns of comorbidity in probands and relatives psychiatrically and pediatrically referred samples. Arch Gen Psychiatry 49: 728-738.

60. Faraone SV, Biederman J, Monuteaux MC (2001) Attention deficit hyperactivity disorder with bipolar disorder in girls: further evidence for a familial subtype? J Affect Disord 64: 19-26.

61. Faraone S, Biederman J, Krifcher Lehman B, Keenan K, et al. (1993) Evidence for the independent familial transmission of attention-deficit hyperactivity disorder and learning disabilities: Results from family genetic study. Am J Psychiatry 150: 891-895.

62. Hudziak JJ, Rudiger LP, Neale MC, Heath AC, Todd RD (2000) A twin study of inattentive, aggressive, and anxious/depressed behaviors. J Am Acad Child Adolesc Psychiatry 39: 469-476.

63. Kuntsi J, Stevenson J (2001) Psychological mechanisms in hyperactivity: II. The role of genetic factors. J Child Psychol Psychiatry 42: 211-219.

64. Martin N, Scourfield J, McGuffin P (2002) Observer effects and heritability of childhood attention-deficit hyperactivity disorder symptoms. Br J Psychiatry 180: 260-265.

65. Sprich-Buckminster S, Biederman J, Milberger S, Faraone SV, Lehman BK (1993) Are perinatal complications relevant to the manifestation of ADDP? Issues of comorbidity and familiality. J Am Acad Child Adolesc Psychiatry 32: 1032-1037.

66. Langley K, Rice F, van den Bree MB, Thapar A (2005) Maternal smoking during pregnancy as an environmental risk factor for attention deficit hyperactivity disorder behaviour. A review. Minerva Pediatr 57: 359-371.

67. Bhutta AT, Cleves MA, Casey PH, Craddock MM, Anand KJ (2002) Cognitive and behavioral outcomes of school-aged children who were born preterm: a meta-analysis. JAMA 288: 728-737.

68. Brookes Kill, J Guindalini C, et al. (2006) A common haplotype of the dopamine transporter gene associated with attention-deficit/hyperactivity disorder and maternal prenatal smoking during pregnancy. Archives of General Psychiatry 63: 74-81.

69. Kahn RS, Khoury J, Nichols WC, et al. (2003) Role of dopamine transporter genotype and maternal prenatal smoking in childhood hyperactive-impulsive,inattentive, and oppositional behaviors. Journal of Pediatrics 143: 104-110.