

This is an Open Access document downloaded from ORCA, Cardiff University's institutional repository: <https://orca.cardiff.ac.uk/id/eprint/138374/>

This is the author's version of a work that was submitted to / accepted for publication.

Citation for final published version:

Faraone, Stephen V., Banaschewski, Tobias, Coghill, David, Zheng, Yi, Biederman, Joseph, Bellgrove, Mark A., Newcorn, Jeffrey H., Gignac, Martin, Al Saud, Nouf M., Manor, Iris, Rohde, Luis Augusto, Yang, Li, Cortese, Samuele, Almagor, Doron, Stein, Mark A., Albatti, Turki H., Aljoudi, Haya F., Alqahtani, Mohammed M.J., Asherson, Philip, Atwoli, Lukoye, Bölte, Sven, Buitelaar, Jan K., Crunelle, Cleo L., Daley, David, Dalsgaard, Søren, Döpfner, Manfred, Espinet, Stacey, Fitzgerald, Michael, Franke, Barbara, Haavik, Jan, Hartman, Catharina A., Hartung, Cynthia M., Hinshaw, Stephen P., Hoekstra, Pieter J., Hollis, Chris, Kollins, Scott H., Sandra Kooij, J.J., Kuntsi, Jonna, Larsson, Henrik, Li, Tingyu, Liu, Jing, Merzon, Eugene, Mattingly, Gregory, Mattos, Paulo, McCarthy, Suzanne, Mikami, Amori Yee, Molina, Brooke S.G., Nigg, Joel T., Purper-Ouakil, Diane, Omigbodun, Olayinka O., Polanczyk, Guilherme V., Pollak, Yehuda, Poulton, Alison S., Rajkumar, Ravi Philip, Reding, Andrew, Reif, Andreas, Rubia, Katya, Rucklidge, Julia, Romanos, Marcel, Ramos-Quiroga, J. Antoni, Schellekens, Arnt, Scheres, Anouk, Schoeman, Renata, Schweitzer, Julie B., Shah, Henal, Solanto, Mary V., Sonuga-Barke, Edmund, Soutullo, César, Steinhausen, Hans-Christoph, Swanson, James M., Thapar, Anita, Tripp, Gail, van de Glind, Geurt, Brink, Wim van den, Van der Oord, Saskia, Venter, Andre, Vitiello, Benedetto, Walitza, Susanne and Wang, Yufeng 2021. The World Federation of ADHD International Consensus Statement: 208 evidence-based conclusions about the disorder. *Neuroscience and Biobehavioral Reviews* 128, pp. 789-818. 10.1016/j.neubiorev.2021.01.022

Publishers page: <http://dx.doi.org/10.1016/j.neubiorev.2021.01.022>

Please note:

Changes made as a result of publishing processes such as copy-editing, formatting and page numbers may not be reflected in this version. For the definitive version of this publication, please refer to the published source. You are advised to consult the publisher's version if you wish to cite this paper.

This version is being made available in accordance with publisher policies. See <http://orca.cf.ac.uk/policies.html> for usage policies. Copyright and moral rights for publications made available in ORCA are retained by the copyright holders.



Journal Pre-proof

The World Federation of ADHD International Consensus Statement: 208
Evidence-based Conclusions about the Disorder

Stephen V. Faraone, Tobias Banaschewski, David Coghill, Yi Zheng, Joseph Biederman, Mark A. Bellgrove, Jeffrey H. Newcorn, Martin Gignac, Nouf M. Al Saud, Iris Manor, Luis Augusto Rohde, Li Yang, Samuele Cortese, Doron Almagor, Mark A. Stein, Turki H. Albatti, Haya F. Aljoudi, Mohammed M.J. Alqahtani, Philip Asherson, Lukoye Atwoli, Sven Bölte, Jan K. Buitelaar, Cleo L. Crunelle, David Daley, Søren Dalsgaard, Manfred Döpfner, Stacey Espinet, Michael Fitzgerald, Barbara Franke, Jan Haavik, Catharina A. Hartman, Cynthia M. Hartung, Stephen P. Hinshaw, Pieter J. Hoekstra, Chris Hollis, Scott H. Kollins, J.J. Sandra Kooij, Jonna Kuntsi, Henrik Larsson, Tingyu Li, Jing Liu, Eugene Merzon, Gregory Mattingly, Paulo Mattos, Suzanne McCarthy, Amori Yee Mikami, Brooke S.G. Molina, Joel T. Nigg, Diane Purper-Ouakil, Olayinka O. Omigbodun, Guilherme V. Polanczyk, Yehuda Pollak, Alison S. Poulton, Ravi Philip Rajkumar, Andrew Reding, Andreas Reif, Katya Rubia, Julia Rucklidge, Marcel Romanos, J. Antoni Ramos-Quiroga, Arnt Schellekens, Anouk Scheres, Renata Schoeman, Julie B. Schweitzer, Henal Shah, Mary V. Solanto, Edmund Sonuga-Barke, César Soutullo, Hans-Christoph Steinhausen, James M. Swanson, Anita Thapar, Gail Tripp, Geurt van de Glind, Wim van den Brink, Saskia Van der Oord, Andre Venter, Benedetto Vitiello, Susanne Walitza, Yufeng Wang

PII: S0149-7634(21)00049-X

DOI: <https://doi.org/10.1016/j.neubiorev.2021.01.022>

Reference: NBR 4048

To appear in: *Neuroscience and Biobehavioral Reviews*

Received Date: 8 June 2020
Revised Date: 25 January 2021
Accepted Date: 25 January 2021

Please cite this article as: Faraone SV, Banaschewski T, Coghill D, Zheng Y, Biederman J, Bellgrove MA, Newcorn JH, Gignac M, Al Saud NM, Manor I, Rohde LA, Yang L, Cortese S, Almagor D, Stein MA, Albatti TH, Aljoudi HF, Alqahtani MMJ, Asherson P, Atwoli L, Bölte S, Buitelaar JK, Crunelle CL, Daley D, Dalsgaard S, Döpfner M, Espinet S, Fitzgerald M, Franke B, Haavik J, Hartman CA, Hartung CM, Hinshaw SP, Hoekstra PJ, Hollis C, Kollins SH, Sandra Kooij JJ, Kuntsi J, Larsson H, Li T, Liu J, Merzon E, Mattingly G, Mattos P, McCarthy S, Mikami AY, Molina BSG, Nigg JT, Purper-Ouakil D, Omigbodun OO, Polanczyk GV, Pollak Y, Poulton AS, Rajkumar RP, Reding A, Reif A, Rubia K, Rucklidge J, Romanos M, Ramos-Quiroga JA, Schellekens A, Scheres A, Schoeman R, Schweitzer JB, Shah H, Solanto MV, Sonuga-Barke E, Soutullo C, Steinhausen H-Christoph, Swanson JM, Thapar A, Tripp G, van de Glind G, Brink Wvd, Van der Oord S, Venter A, Vitiello B, Walitza S, Wang Y, The World Federation of ADHD International Consensus Statement: 208 Evidence-based Conclusions about the Disorder, *Neuroscience and Biobehavioral Reviews* (2021), doi: <https://doi.org/10.1016/j.neubiorev.2021.01.022>

This is a PDF file of an article that has undergone enhancements after acceptance, such as the addition of a cover page and metadata, and formatting for readability, but it is not yet the definitive version of record. This version will undergo additional copyediting, typesetting and review before it is published in its final form, but we are providing this version to give early visibility of the article. Please note that, during the production process, errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

© 2020 Published by Elsevier.

The World Federation of ADHD International Consensus Statement: 208 Evidence-based Conclusions about the Disorder

Short Title: The World Federation of ADHD International Consensus Statement

Stephen V. Faraone PhD^{a,b,c,*} sfaraone@childpsychresearch.org, Tobias Banaschewski MD, PhD^{d,e,f}, David Coghill MD^g, Yi Zheng MD^{h,i,j,k,l,m}, Joseph Biederman MD^{n,o}, Mark A. Bellgrove PhD^{16,17p,q}, Jeffrey H. Newcorn MD^{18,3r,c}, Martin Gignac FRCPC^{19,20,21s,t,u}, Nouf M. Al Saud^{22v}, Iris Manor MD^{23,24w,x}, Luis Augusto Rohde MD, PhD^{25y}, Li Yang MD, PhD^{z,A,l}, Samuele Cortese MD, PhD^{B,C,D,E,F}, Doron Almagor MD, FRCPC^{G,H}, Mark A. Stein PhD^{I,G}, Turki H. Albatti MD^K, Haya F. Aljoudi Psy.D^{L,M}, Mohammed M.J. Alqahtani PhD^{N,O}, Philip Asherson MRCPsych PD^P, Lukoye Atwoli MD, PhD^{Q,R,S,T}, Sven Bölte PhD^{U,V,W}, Jan K. Buitelaar MD, PhD^{5X}, Cleo L. Crunelle MD, PhD^{Y,Z}, David Daley PhD^{aa,ab}, Søren Dalsgaard MD, PhD^{ac,ad}, Manfred Döepfner PhD^{ae,af}, Stacey Espinet PhD^{ag}, Michael Fitzgerald MD^{ah}, Barbara Franke PhD^{ai,aj}, Jan Haavik MD, PhD^{ak,al}, Catharina A. Hartman PhD^{am,an,ao,ap}, Cynthia M. Hartung PhD^{aq}, Stephen P. Hinshaw PhD^{ar,as,at,au,av}, Pieter J. Hoekstra MD, PhD^{aw}, Chris Hollis PhD, FRCPsych^{E,ax,ay,az}, Scott H. Kollins PhD^{ba,bb}, J.J. Sandra Kooij MD, PhD^{bc,bd,be,bf}, Jonna Kuntsi PhD^{bg}, Henrik Larsson PhD^{bh,bi}, Tingyu Li MD^{bj,bk,bl}, Jing Liu MD^{z,A,l,bm,bn}, Eugene Merzon MD^{bo,bp,bq,br}, Gregory Mattingly MD^{bs}, Paulo Mattos MD, PhD^{bt,bu,bv}, Suzanne McCarthy PhD^{bw}, Amori Yee Mikami PhD^{bx}, Brooke S.G. Molina PhD^{by}, Joel T. Nigg PhD^{bz}, Diane Purper-Ouakil MD, PhD^{ca,cb}, Olayinka O. Omigbodun MBBS, MPH, FMCPsych^{cc,cd}, Guilherme V. Polanczyk MD, PhD^{ce}, Yehuda Pollak PhD^{cf,cg}, Alison S. Poulton MD^{ch,ci}, Ravi Philip Rajkumar MD^{cj}, Andrew Reding, Andreas Reif MD^{cl,cm}, Katya Rubia PhD^{cn,b,co}, Julia Rucklidge PhD^{cp}, Marcel Romanos PhD^{cq,cr,cs}, J. Antoni Ramos-Quiroga MD, PhD^{ct,cu,cv,cw,cx,cy,cz}, Arnt Schellekens MD, PhD^{da,db}, Anouk Scheres PhD^{dc}, Renata Schoeman MD, MBA, PhD^{dd,de,df,dg,dh,di}, Julie B. Schweitzer PhD^{dj}, Henal Shah MD^{dk}, Mary V. Solanto Ph.D. ^{di,dm,dn,do}, Edmund Sonuga-Barke PhD^{dp,dq}, César Soutullo MD, PhD^{dr,co,c}, Hans-Christoph Steinhausen MD, MSc, PhD, DMSc^{ds,dt,du,dv}, James M. Swanson PhD^{dw}, Anita Thapar FRCPsych, PhD^{dx}, Gail Tripp PhD^{dy}, Geurt van de Glind PhD^{dz}, Wim van den Brink MD, PhD^{ea}, Saskia Van der Oord PhD^{eb,ec}, Andre Venter PhD^{ed}, Benedetto Vitiello MD^{ee,ef}, Susanne Walitza MD, MSc^{eg}, Yufeng Wang MD, PhD^{z,A,l}

^aDepartments of Psychiatry and Neuroscience and Physiology, Psychiatry Research Division, SUNY Upstate Medical University, Syracuse, NY, USA

^bWorld Federation of ADHD

^cAmerican Professional Society of ADHD and Related Disorders (APSARD)

^dDepartment of Child and Adolescent Psychiatry and Psychotherapy, Central Institute of Mental Health, Medical Faculty Mannheim, University of Heidelberg, Mannheim, Germany

^eChild and Adolescent Psychiatrist's Representative, Zentrales-ADHS-Netz

^fThe German Association of Child and Adolescent Psychiatry and Psychotherapy

^gDepartments of Paediatrics and Psychiatry, Faculty of Medicine, Dentistry and Health Sciences, University of Melbourne, Melbourne, Australia

^hBeijing Anding Hospital, Capital Medical University, Beijing, China

ⁱThe National Clinical Research Center for Mental Disorders, Beijing, China

^jBeijing Key Laboratory of Mental Disorders, Beijing, China

^kBeijing Institute for Brain Disorders, Beijing, China

^lAsian Federation of ADHD

^mChinese Society of Child and Adolescent Psychiatry

- ⁿClinical & Research Programs in Pediatric Psychopharmacology & Adult ADHD, Massachusetts General Hospital, Boston, MA, USA
- ^oDepartment of Psychiatry, Harvard Medical School, Boston, MA, USA
- ^pTurner Institute for Brain and Mental Health and School of Psychological Sciences, Monash University, Clayton VIC, Australia
- ^qPresident, Australian ADHD Professionals Association (AADPA)
- ^rDepartments of Psychiatry and Pediatrics, Division of ADHD and Learning Disorders, Icahn School of Medicine at Mount Sinai, New York, NY, USA
- ^sDepartment of Child and Adolescent Psychiatry, Montreal Children's Hospital, MUHC, Montreal, Canada
- ^tChild and Adolescent Psychiatry Division, McGill University, Montreal, Canada
- ^uChair, Canadian ADHD Research Alliance (CADDRA)
- ^vChairman, Board of Directors, Saudi ADHD Society, Saudi Arabia
- ^wAssociate professor and head of the ADHD clinic, Geha MHC, Sackler School of Medicine, Tel Aviv University, Tel Aviv, Israel
- ^xChair, Israeli Society of ADHD (ISA)
- ^yDepartment of Psychiatry, Federal University of Rio Grande do Sul, Brazil
- ^zPeking University Sixth Hospital/Institute of Mental Health, National Clinical Research Center for Mental Disorders (Peking University Sixth Hospital), Beijing, China
- ^ANHC Key Laboratory of Mental Health (Peking University), Beijing, China
- ^BUniversity of Southampton, Southampton, UK
- ^CSolent NHS Trust, Southampton, UK
- ^DNew York University, New York, NY, USA
- ^EUniversity of Nottingham, Nottingham, UK
- ^FEuropean ADHD Guidelines (EAGG)
- ^GUniversity of Toronto, SickKids Centre for Community Mental Health, Toronto, Canada,
- ^HCanadian ADHD Research Alliance (CADDRA)
- ^IUniversity of Washington, Seattle, WA, USA
- ^JSeattle Children's Hospital, Seattle, WA, USA
- ^KConsultant Child and Adolescent Psychiatrist, Member, Saudi ADHD Society Medical and Psychological Committee, Saudi Arabia
- ^LConsultant Clinical Neuropsychologist, King Faisal Specialist Hospital & Research Center, Riyadh, Saudi Arabia
- ^MSaudi ADHD Society Medical and Psychological Committee
- ^NClinical Psychology, King Khalid University, Abha, Saudi Arabia
- ^OHead of the National Research Committee, Saudi ADHD Society
- ^PSocial Genetic & Developmental Psychiatry, Institute of Psychiatry, Psychology, and Neuroscience, King's College London, UK
- ^QDepartment of Mental Health, Moi University School of Medicine, Eldoret, Kenya
- ^RR&D Director, citiesRISE
- ^SHon Vice-President, Kenya Medical Association
- ^TSecretary-General, African Association of Psychiatrists
- ^UCenter of Neurodevelopmental Disorders (KIND) and Centre for Psychiatry Research, Karolinska Institutet & Stockholm Health Care Services, Region Stockholm, Sweden
- ^VChild and Adolescent Psychiatry, Stockholm Healthcare Services, Region Stockholm, Sweden

^WCurtin Autism Research Group, School of Occupational Therapy, Social Work and Speech Pathology, Curtin University, Perth, Western Australia

^XDepartment of Cognitive Neuroscience, Donders Institute for Brain, Cognition and Behaviour, Radboud University Medical Centre, Nijmegen, The Netherlands

^YVrije Universiteit Brussel (VUB), Universitair Ziekenhuis Brussel (UZ Brussel), Dept. of Psychiatry, Brussel, Belgium

^ZInternational Collaboration on ADHD and Substance Abuse (ICASA), Nijmegen, The Netherlands

^{aa}Division of Psychiatry and Applied Psychology, School of Medicine University of Nottingham, Nottingham, UK

^{ab}NIHR MindTech Mental Health MedTech Cooperative & Centre for ADHD and Neurodevelopmental Disorders Across the Lifespan (CANDAL), Institute of Mental Health, University of Nottingham, Nottingham, UK

^{ac}National Centre for Register-based Research, Aarhus University, Aarhus, Denmark

^{ad}The Lundbeck Foundation Initiative for Integrative Psychiatric Research, iPSYCH, Aarhus, Denmark

^{ae}Department of Child and Adolescent Psychiatry, Psychosomatics and Psychotherapy, School of Child and Adolescent Cognitive Behavior Therapy (AKiP), Faculty of Medicine and University Hospital Cologne, University Cologne, Cologne, Germany

^{af}Psychologist's Representative, Zentrales-ADHS-Netz, Germany

^{ag}Canadian ADHD Resource Alliance (CADDRA), Canada

^{ah}Former Professor of Child Psychiatry, Trinity College, Dublin, Ireland

^{ai}Departments of Human Genetics and Psychiatry, Donders Institute for Brain, Cognition and Behaviour, Radboud University Medical Center, Nijmegen, The Netherlands

^{aj}Professional Board, ADHD Europe

^{ak}Department of Biomedicine, University of Bergen, Bergen, Norway

^{al}Division of Psychiatry, Haukeland University Hospital, Bergen, Norway

^{am}University of Groningen, Groningen, The Netherlands

^{an}University Medical Center Groningen, Groningen, The Netherlands

^{ao}Interdisciplinary Center Psychopathology and Emotion regulation (ICPE), Groningen, The Netherlands

^{ap}ADHD across the Lifespan Network from European College of Neuropsychopharmacology (ECNP)

^{aq}Department of Psychology, University of Wyoming, Laramie, WY, USA

^{ar}University of California, Berkeley, CA, USA

^{as}University of California, San Francisco, CA, USA

^{at}American Psychological Association

^{au}Association for Psychological Science

^{av}Society for Research in Child Development

^{aw}University of Groningen, University Medical Center Groningen, Department of Child and Adolescent Psychiatry, Groningen, Netherlands

^{ax}Nottinghamshire Healthcare NHS Foundation Trust, Nottingham, UK

^{ay}NIHR MindTech MedTech Co-operative, Nottingham, UK

^{az}NIHR Nottingham Biomedical Research Centre, Nottingham, UK

^{ba}Duke University School of Medicine, Durham, NC, USA

^{bb}Duke Clinical Research Institute, Durham, NC, USA

- ^{bc}Associate Professor of Psychiatry, Amsterdam University Medical Center (VUMc), Amsterdam, The Netherlands
- ^{bd}Head of Dutch Expertise Center Adult ADHD, PsyQ, The Hague, The Netherlands
- ^{be}European Network Adult ADHD
- ^{bf}DIVA Foundation
- ^{bg}Social, Genetic and Developmental Psychiatry Centre, Institute of Psychiatry, Psychology and Neuroscience, King's College London, London, UK
- ^{bh}School of Medical Sciences, Örebro University, Örebro, Sweden
- ^{bi}Department of Medical Epidemiology and Biostatistics, Karolinska Institutet, Solna, Sweden,
- ^{bj}Growth, Development and Mental Health Center for Children and Adolescents, Children's Hospital of Chongqing Medical University, Chongqing, China
- ^{bk}National Research Center for Clinical Medicine of Child Health and Disease, Chongqing, China
- ^{bl}The Subspecialty Group of Developmental and Behavioral Pediatrics, the Society of Pediatrics, Chinese Medical Association.
- ^{bm}The Chinese Society of Child and Adolescent Psychiatry
- ^{bn}The Asian Society for Child and Adolescent Psychiatry and Allied Professions
- ^{bo}Assistant Professor, Department of Family Medicine, Sackler School of Medicine, Tel Aviv University, Tel Aviv, Israel
- ^{bp}Head of the Department of Managed Care, Leumit Health Services, Tel Aviv, Israel,
- ^{bq}Executive Board Member, Israeli Society of ADHD, Israel
- ^{br}Chair of ADHD Section, Israeli National Diabetes Council, Israel
- ^{bs}Washington University, St. Louis, MO; Midwest Research Group, St Charles, MO, St. Louis, MO, USA
- ^{bt}Federal University of Rio de Janeiro, Rio de Janeiro, Brazil
- ^{bu}D'Or Institute for Research and Education, Rio de Janeiro, Brazil
- ^{bv}Brazilian Attention Deficit Association (ABDA)
- ^{bw}School of Pharmacy, University College Cork, Cork, Ireland
- ^{bx}University of British Columbia, Vancouver, BC, Canada
- ^{by}Departments of Psychiatry, Psychology, Pediatrics, Clinical & Translational Science, University of Pittsburgh, Pittsburgh, PA, USA
- ^{bz}Center for ADHD Research, Department of Psychiatry, Oregon Health & Science University, Portland, OR, USA
- ^{ca}University of Montpellier, CHU Montpellier Saint Eloi, MPEA, Medical and Psychological Unit for Children and Adolescents (MPEA), Montpellier, France
- ^{cb}INSERM U 1018 CESP-Developmental Psychiatry, France
- ^{cc}Centre for Child & Adolescent Mental Health, College of Medicine, University of Ibadan, Ibadan, Nigeria
- ^{cd}Department of Child & Adolescent Psychiatry, University College Hospital, Ibadan, Nigeria,
- ^{ce}Faculdade de Medicina FMUSP, University of São Paulo, Brazil
- ^{cf}Seymour Fox School of Education, The Hebrew University of Jerusalem, Israel
- ^{cg}The Israeli Society of ADHD (ISA), Israel
- ^{ch}Brain Mind Centre Nepean, University of Sydney, Sydney, Australia
- ^{ci}Australian ADHD Professionals Association (AADPA), Australia

- ^{cj}Jawaharlal Institute of Postgraduate Medical Education and Research, Pondicherry, India,
^{ci}Department of Psychiatry, Psychosomatic Medicine and Psychotherapy, University Hospital Frankfurt, Frankfurt am Main, Germany
^{cm}German Psychiatric Association
^{cn}Department of Child & Adolescent Psychiatry, Institute of Psychiatry, Psychology & Neurosciences, King's College London, London, UK
^{co}European Network for Hyperkinetic Disorders (EUNETHYDIS)
^{cp}School of Psychology, Speech and Hearing, University of Canterbury, Christchurch, New Zealand
^{cq}Department of Child and Adolescent Psychiatry, Psychosomatics and Psychotherapy, Center of Mental Health, University Hospital Würzburg, Würzburg, Germany
^{cr}The German Association of Child and Adolescent Psychiatry and Psychotherapy, Germany,
^{cs}Zentrales-ADHS-Netz
^{ct}Department of Psychiatry, Hospital Universitari Vall d'Hebron, Barcelona, Catalonia, Spain,
^{cu}Group of Psychiatry, Mental Health and Addictions, Vall d'Hebron Research Institute (VHIR), Barcelona, Catalonia, Spain
^{cv}Biomedical Network Research Centre on Mental Health (CIBERSAM), Universitat Autònoma de Barcelona, Barcelona, Catalonia, Spain
^{cw}Department of Psychiatry and Forensic Medicine, Universitat Autònoma de Barcelona, Barcelona, Catalonia, Spain
^{cx}Chair of Neurodevelopmental Disorders Across Lifespan Section of European Psychiatric Association
^{cy}Board Member, International Collaboration on ADHD and Substance Abuse (ICASA), ^{cz}Board member of DIVA Foundation
^{da}Radboud University Medical Centre, Donders Institute for Brain, Cognition, and Behavior, Department of Psychiatry, Nijmegen, The Netherlands
^{db}International Collaboration on ADHD and Substance Abuse (ICASA), Nijmegen, The Netherlands
^{dc}Behavioural Science Institute, Radboud University, Nijmegen, The Netherlands
^{dd}Associate Professor and head of the MBA in Healthcare Leadership, University of Stellenbosch Business School, Cape Town, South Africa
^{de}South African Special Interest Group for Adult ADHD
^{df}The South African Society of Psychiatrists/Psychiatry Management Group management guidelines for ADHD
^{dg}World Federation of Biological Psychiatry
^{dh}American Psychiatric Association
^{di}Association for NeuroPsychoEconomics
^{dj}Department of Psychiatry and Behavioral Sciences and the Mind Institute, University of California, Davis, Sacramento, CA, USA
^{dk}Topiwala National Medical College & BYL Nair Ch. Hospital, Mumbai, India
^{dl}The Zucker School of Medicine at Hofstra-Northwell, Northwell Health, Hemstead, NY, USA
^{dm}Professional Advisory Board, Children and Adults with Attention-Deficit/Hyperactivity Disorder (CHADD)
^{dn}Professional Advisory Board, American Professional Society of ADHD and Related Disorders (APSARD)
^{do}Professional Advisory Board, National Center for Children with Learning Disabilities (NCLD)

^{dp}Department of Child and Adolescent Psychiatry, Institute of Psychiatry, Psychology & Neuroscience, King's College London, London, UK

^{dq}Department of Child & Adolescent Psychiatry, Aarhus University, Aarhus, Denmark

^{dr}Louis A. Faillace MD, Department of Psychiatry and Behavioral Sciences, University of Texas Health Science Center at Houston, Houston, TX, USA

^{ds}University of Zurich, CH, Switzerland

^{dt}University of Basel, CH, Switzerland

^{du}University of Southern Denmark, Odense, Denmark

^{dv}Centre of Child and Adolescent Mental Health, Copenhagen, Denmark

^{dw}Department of Pediatrics, University of California Irvine, Irvine, CA, USA

^{dx}Division of Psychological Medicine and Clinical Neurosciences, MRC Centre for Neuropsychiatric Genetics and Genomics, Cardiff University School of Medicine, Wales, UK,

^{dy}Human Developmental Neurobiology Unit, Okinawa Institute of Science and Technology Graduate University, Okinawa, Japan

^{dz}ICASA Foundation, Radboud University Medical Centre/ Nijmegen Institute for Scientist-Practitioners in Addiction, Nijmegen, The Netherlands

^{ea}Amsterdam University Medical Centers, Academic Medical Center, Amsterdam, The Netherlands

^{eb}Psychology and Educational Sciences, KU Leuven, Leuven, Belgium

^{ec}European ADHD Guidelines Group

^{ed}University of the Free State, Bloemfontein, South Africa

^{ee}University of Torino, Torino, Italy

^{ef}Johns Hopkins University School of Public Health, Baltimore, MD, USA

^{eg}Department of Child and Adolescent Psychiatry and Psychotherapy, University Hospital of Psychiatry Zurich, University of Zurich, Zurich, Switzerland

*Corresponding Author: Stephen V. Faraone, PhD Department of Psychiatry, SUNY Upstate Medical University, Institute for Human Performance, Room 3707, 505 Irving Ave. Syracuse, NY 13210.

Highlights

- ADHD occurs in 5.9% of youth and 2.5% of adults.
- Most cases of ADHD are caused by the combined effects of many genetic and environmental risks.
- There are small differences in the brain between people with and without ADHD.
- Untreated ADHD can lead to many adverse outcomes.
- ADHD costs society hundreds of billions of dollars each year, worldwide.

Abstract

Background: Misconceptions about ADHD stigmatize affected people, reduce credibility of providers, and prevent/delay treatment. To challenge misconceptions, we curated findings with strong evidence base.

Methods: We reviewed studies with more than 2,000 participants or meta-analyses from five or more studies or 2,000 or more participants. We excluded meta-analyses that did not assess publication bias, except for meta-analyses of prevalence. For network meta-analyses we required comparison adjusted funnel plots. We excluded treatment studies with waiting-list or treatment as usual controls. From this literature, we extracted evidence-based assertions about the disorder.

Results: We generated 208 empirically supported statements about ADHD. The status of the included statements as empirically supported is approved by 79 authors from 27 countries and 6 continents. The contents of the manuscript are endorsed by 362 people who have read this document and agree with its contents.

Conclusions: Many findings in ADHD are supported by meta-analysis. These allow for firm statements about the nature, course, outcome causes, and treatments for disorders that are useful for reducing misconceptions and stigma.

Key Words: ADHD, diagnosis, treatment, course, outcome, genetics, brain

Introduction

Nearly two decades ago, an international team of scientists published the first International Consensus Statement on attention-deficit hyperactivity disorder (ADHD) (Barkley, 2002). They sought to present the wealth of scientific data attesting to the validity of ADHD as a mental disorder and to correct misconceptions about the disorder that stigmatized affected people, reduced the credibility of health care providers, and prevented or delayed treatment of individuals challenged by the disorder (DosReis et al., 2010; Horton-Salway, 2013; McLeod et al., 2007; Mueller et al., 2012).

This paper updates the International Consensus Statement by cataloging important scientific discoveries from the last 20 years. We do not intend to present an encyclopedia of ADHD or guidelines for diagnosis and treatment. The latter can be found in the references cited. Our aim is to provide current and accurate information about ADHD supported by a substantial and rigorous body of evidence.

Methods

We identified evidence-based statements about ADHD through expert scrutiny of published high quality meta-analyses and very large studies. Expert scrutiny was provided by a project Steering Committee (Supplemental Table 1) which included representatives from the following professional groups dedicated to research and clinical care of ADHD: The World Federation of ADHD, European Network for Hyperkinetic Disorders (Eunethydis), the American Professional Society of ADHD and Related Disorders, the Canadian ADHD Resource Alliance, the Asian Federation of ADHD, the Latin American League of ADHD, the Australian ADHD Professionals Association, the Israeli Society of ADHD, the Saudi ADHD Society, Neurodevelopmental

Disorders Across Lifespan section of the European Psychiatric Association, the ADHD Guidelines Group of the Association of Medical Scientific Societies in Germany, the ADHD Network of European College of Neuropsychopharmacology, the Chinese Society of Child and Adolescent Psychiatry and the ADHD Section of the World Psychiatric Association.

For large cohort studies, we searched PubMed with these search criteria: ADHD [tiab] AND (nationwide [tiab] OR national [tiab] OR register [tiab] OR registry [tiab]) NOT review [Publication Type] NOT meta-analysis [Publication Type]. For meta-analyses, we searched PubMed with these search criteria: ADHD [All Fields] AND (meta-analysis [Title] OR meta-analysis [Title] OR meta-analytic [Title] OR systematic review [Title]). We excluded meta-analyses that did not assess publication bias, except for meta-analyses of prevalence. For network meta-analyses we required that comparison adjusted funnel plots be presented. For treatment studies, we excluded results of meta-analyses including comparisons of treatments with waiting-list or treatment as usual controls.

Apart from statements about the history of ADHD and its diagnostic criteria, we required each evidence-based statement to be supported by meta-analyses or by large registry studies with more than 2,000 participants. We required meta-analyses to report data from five or more studies or 2,000 or more participants.

We describe the magnitude of effect size findings using standard criteria as follows:

standardized mean difference: small = 0.20, medium = 0.50, large = 0.80; correlation

coefficient: small = 0.10, medium = 0.24, large = 0.37 (Ellis, 2010; Rosenthal and Rosnow, 1984).

“Moderate” is used as a synonym for “medium,” and “strong” for “large.” A “small” effect is

generally difficult to observe in an individual but may be very important for public health if it concerns a common exposure that affects many children. A “medium” effect is expected to be noticeable to a careful observer (Cohen, 1988). A “large” effect is generally relevant to clinical practice at the level of the individual.

If a topic is not included in this document, it does not mean the topic is unimportant; rather, it means the evidence found was insufficient to allow firm conclusions. This could be because there were insufficient studies of quality, because no attempt was made to assess publication bias, or because the data available did not support the claims made. After the document was completed, we invited additional colleagues to join as signatories to indicate their support of the document. In what follows, we use the term “evidence-based” to refer to evidence that meets the inclusion/exclusion criteria we used in our literature search. We recognize that other criteria could be applied, such as requiring the absence of severe heterogeneity in meta-analyses or increasing the numbers of research participants.

Overview of Results

Our search strategy generated 208 empirically supported statements about ADHD. For details, see the PRISMA diagram in Supplemental Figure 1. The status of the included statements as empirically supported has been approved by the 79 authors from 27 countries and 6 continents (Supplemental Figure 2). It has been endorsed by 362 people who have read this document and agree with its contents (Supplemental Table 2). Table 1 summarizes our findings along with the item numbers that support each statement. A limitation of this consensus statement is that we do not report well-established research findings for which meta-analyses or very large

studies do not exist. The absence of such a study, is not always an indication of knowledge of absence of an effect.

Table 1: Summary of Findings	
Findings	Items
The syndrome we now call ADHD has been described in the medical literature since 1775.	1 - 13
When made by a licensed clinician, the diagnosis of ADHD is well-defined and valid at all ages, even in the presence of other psychiatric disorders, which is common.	14-19
ADHD is more common in males and occurs in 5.9% of youth and 2.5% of adults. It has been found in studies from Europe, Scandinavia, Australia, Asia, the Middle East, South America, and North America.	20-25
ADHD is rarely caused by a single genetic or environmental risk factor but most cases of ADHD are caused by the combined effects of many genetic and environmental risks each having a very small effect.	26-62
People with ADHD often show impaired performance on psychological tests of brain functioning, but these tests cannot be used to diagnose ADHD.	63-70
Neuroimaging studies find small differences in the structure and functioning of the brain between people with and without ADHD. These differences cannot be used to diagnose ADHD.	71-77
People with ADHD are at increased risk for obesity, asthma, allergies, diabetes mellitus, hypertension, sleep problems, psoriasis, epilepsy, sexually transmitted infections, abnormalities of the eye, immune disorders, and metabolic disorders.	78-100

People with ADHD are at increased risk for low quality of life, substance use disorders, accidental injuries, educational underachievement, unemployment, gambling, teenage pregnancy, difficulties socializing, delinquency, suicide, and premature death.	101-136
Studies of economic burden show that ADHD costs society hundreds of billions of dollars each year, worldwide.	137-147
Regulatory agencies around the world have determined that several medications are safe and effective for reducing the symptoms of ADHD as shown by randomized controlled clinical trials.	148-157
Treatment with ADHD medications reduces accidental injuries, traumatic brain injury, substance abuse, cigarette smoking, educational underachievement, bone fractures, sexually transmitted infections, depression, suicide, criminal activity and teenage pregnancy.	158-177
The adverse effects of medications for ADHD are typically mild and can be addressed by changing the dose or the medication.	178-188
The stimulant medications for ADHD are more effective than non-stimulant medications but are also more likely to be diverted, misused, and abused.	189-194
Non-medication treatments for ADHD are less effective than medication treatments for ADHD symptoms, but are frequently useful to help problems that remain after medication has been optimized.	195-208

A Brief History: ADHD is not a New Disorder

The concept of ADHD has a long history, starting with clinical reports from European countries.

The clinical significance of the signs and symptoms of the disorder has been recognized for over two centuries. Although these early reports did not use the term “ADHD”, they described children who showed the symptoms and impairments we now recognize as ADHD. For a detailed history see (Lange et al., 2010; Taylor, 2011; Weikard, 1799). Here are highlights from the early history of ADHD:

1. 1775: Melchior Adam Weikard, a German physician, wrote the first textbook description of a disorder with the hallmarks of ADHD.
2. 1798: Alexander Crichton from the Royal College of Physicians (United Kingdom) described a similar disorder in a medical textbook (Palmer and Finger, 2001).
3. 1845: Heinrich Hoffmann, who later became head of the first psychiatric hospital in Frankfurt am Main, Germany, described hyperactivity and attention deficits in a children’s book which documented ADHD-like behaviors and their associated impairments (Hoffmann, 1990).
4. 1887-1901: Désiré-Magloire Bourneville, Charles Boulanger, Georges Paul-Boncour, and Jean Philippe described an equivalent of ADHD in French medical and educational writings (Martinez-Badia and Martinez-Raga, 2015).
5. 1902: George Still, a physician in the United Kingdom, wrote the first description of the disorder in a scientific journal (Still, 1902a; Still, 1902b, c).

6. 1907: Augusto Vidal Perera wrote the first Spanish compendium of child psychiatry. He described the impact of inattention and hyperactivity among schoolchildren (Vidal Perera, 1907).
7. 1917: the Spanish neurologist and psychiatrist Gonzalo Rodriguez-Lafora described symptoms of ADHD in children and said they were probably caused by a brain disorder with genetic origins (Lafora, 1917).
8. 1932: Franz Kramer and Hans Pollnow, from Germany, described an ADHD-like syndrome and coined the term “hyperkinetic disorder”, which was later adopted as a term by the World Health Organization (Kramer and Pollnow, 1932; Neumarker, 2005).
9. 1937: Charles Bradley, from the USA, discovered that an amphetamine medication reduced ADHD-like symptoms (Bradley, 1937).
10. 1940s: ADHD-like symptoms in children described as “minimal brain dysfunction”.
11. 1956-1958: First hint in follow-up study of the persistence of minimal brain dysfunction-related behaviors into adulthood (Morris et al., 1956; O'Neal and Robins, 1958)
12. 1960s: U.S. Food and Drug Administration approved methylphenidate (Ritalin) for behavioral disorders in children.
13. 1970s to today: Diagnostic criteria for ADHD evolved based on research showing that the diagnosis predicts treatment response, clinical course, and family history of the disorder.

How is ADHD diagnosed?

ADHD can only be diagnosed by a licensed clinician who interviews the parent or caregiver and/or patient to document criteria for the disorder (American Psychiatric Association, 2013; Chinese Society of Psychiatry, 2001; Faraone et al., 2015; Feldman and Reiff, 2014; Pearl et al., 2001; Stein, 2008; World Health Organization, 2018). It cannot be diagnosed by rating scales alone, neuropsychological tests, or methods for imaging the brain.

The diagnosis of ADHD has been criticized as being subjective because it is not based on a biological test. This criticism is unfounded. ADHD meets standard criteria for validity of a mental disorder established by Robins and Guze (Faraone, 2005; 1970). The disorder is considered valid because: 1) well-trained professionals in a variety of settings and cultures agree on its presence or absence using well-defined criteria and 2) the diagnosis is useful for predicting a) additional problems the patient may have (e.g., difficulties learning in school); b) future patient outcomes (e.g., risk for future drug abuse); c) response to treatment (e.g., medications and psychological treatments); and d) features that indicate a consistent set of causes for the disorder (e.g., findings from genetics or brain imaging) (Faraone, 2005). Professional associations have endorsed and published guidelines for diagnosing ADHD (Alliance, 2011; Banaschewski T, 2018; Bolea-Alamanac et al., 2014; Crunelle et al., 2018; Flisher, 2013; Graham et al., 2011; Kooij et al., 2019; National Collaborating Centre for Mental Health, 2018; National Institute for Health Care and Excellence, 2018; Pliszka, 2007; Schoeman and Liebenberg, 2017; Seixas et al., 2012; Taylor et al., 2004; Wolraich et al., 2011).

The main features of the diagnosis are:

14. The diagnosis requires: 1) the presence of developmentally inappropriate levels of hyperactive-impulsive and/or inattentive symptoms for at least 6 months; 2) symptoms occurring in different settings (e.g., home and school); 3) symptoms that cause impairments in living; 4) some of the symptoms and impairments first occurred in early to mid-childhood; and 4) no other disorder better explains the symptoms (American Psychiatric Association, 2013; World Health Organization, 2018; Yi and Jing, 2015).
15. The clinical presentation of ADHD can be described as primarily inattentive, primarily hyperactive-impulsive, or combined, depending on the nature of their symptoms (American Psychiatric Association, 2013). Meta-analyses indicate that inattention is more strongly associated with academic impairment, low self-esteem, negative occupational outcomes and lower overall adaptive functioning. Hyperactive-impulsive symptoms are associated with peer rejection, aggression, risky driving behaviors and accidental injuries. Patterns of associated disorders also differ between the dimensions (Willcutt et al., 2012).
16. ADHD impairs the functioning of highly intelligent people, so the disorder can be diagnosed in this group. A population-based birth cohort study of over 5,700 children found no significant differences among children with high, average, or low IQ and ADHD in median age at which ADHD criteria were met, rates of learning disorders, psychiatric disorders, and substance abuse, and rates of stimulant treatment (Katusic et al., 2011; Rommelse et al., 2017).

17. In adolescence and young adulthood, many individuals with a history of childhood ADHD continue to be impaired by the disorder, although they often show reduced hyperactivity and impulsivity while retaining symptoms of inattention (Faraone et al., 2006).
18. Many large epidemiologic and clinical studies show that ADHD often co-occurs with other psychiatric disorders, especially depression, bipolar disorder, autism spectrum disorders, anxiety disorders, oppositional defiant disorder, conduct disorder, eating disorders, and substance use disorders (Bernardi et al., 2012; Chen et al., 2018c; Groenman et al., 2017; Nazar et al., 2016; Solberg et al., 2018; Tung et al., 2016; Yao et al., 2019). Their presence does not rule out a diagnosis of ADHD.
19. A meta-analysis comprising 25 studies with over eight million participants found that children and adolescents who are relatively younger than their classmates are more likely to have been diagnosed with ADHD (Caye et al., 2020)

How Common is ADHD?

ADHD occurs throughout the developed and developing world and is more common in males compared with females. It has not become more common over the past three decades although due to increased recognition by clinicians, the disorder is more likely to be diagnosed today than in prior decades.

20. A meta-analysis of 19 studies with over 55,000 participants found that 5.9% of youths meet diagnostic criteria for ADHD (Willcutt, 2012). Another meta-analysis, with 135 studies and about a quarter of a million youths, found no significant differences in prevalence between

North America and Europe, Asia, Africa, South America, and Oceania (Polanczyk et al., 2014).

21. The latter meta-analysis found no increase in the prevalence of ADHD in children and adolescents over the past three decades (Polanczyk et al., 2014). Although the prevalence of ADHD has not changed in this time period, large studies from the US and Sweden indicate that ADHD is more likely to have been diagnosed in recent years, which reflects changes in administrative and clinical practices (Rydell et al., 2018; Song et al., 2019; Xu et al., 2018).
22. A meta-analysis of six studies with over 5,300 participants estimated the prevalence of ADHD in adulthood to be 2.5% (Simon et al., 2009). A meta-analysis of 20 studies encompassing 13 countries and seven regions/metropolitan areas, involving more than 26,000 participants, estimated that 2.8% of adults meet criteria for ADHD (Fayyad et al., 2017). The lower prevalence in adults compared with youth is consistent with a meta-analysis of 21 studies with over 1,600 participants showing that only about one in six youths with ADHD still meet full diagnostic criteria for ADHD at age 25, and about half show signs of residual impairment (Faraone et al., 2006).
23. A meta-analysis of nine studies with a total of over 32,000 older adults found a prevalence of 2.2% based on ADHD rating scales, dropping to 1.5% when limited to persons at least fifty years old. Yet a meta-analysis of seven studies with over 11.7 million participants based on ADHD clinical diagnoses, performed by the same team, reported a prevalence of only 0.2% for persons at least fifty years old. A third meta-analysis performed by the same

researchers, of four studies with over 9.2 million participants, found an ADHD treatment rate of only 0.02% among persons at least fifty years old (Dobrosavljevic et al., 2020).

24. A meta-analysis of 19 studies encompassing over 150,000 U.S. Black youths under 18 years old reported an ADHD prevalence rate of 14 percent. The authors concluded, "Black individuals are at higher risk for ADHD diagnoses than the general US population. These results highlight a need to increase ADHD assessment and monitoring among Black individuals from different social backgrounds" (Cénat et al., 2020).
25. ADHD is more common in males. A meta-analysis of parent ratings of symptoms in 29 studies with over 42,000 participants, and teacher ratings in 24 studies with over 56,000 participants, found a roughly two-to-one male/female ratio in youth (Willcutt, 2012).

What causes ADHD?

For most people with ADHD, many genetic and environmental risk factors accumulate to cause the disorder (Faraone et al., 2015). The environmental risks for ADHD exert their effects very early in life, during the fetal or early postnatal period. In rare cases, however, ADHD-like symptoms can be caused by extreme deprivation early in life (Kennedy et al., 2016), a single genetic abnormality (Faraone and Larsson, 2018), or traumatic brain injury early in life (Stojanovski et al., 2019). These findings are helpful to understand the causes of ADHD but are not useful for diagnosing the disorder. The associations between aspects of the environment and the onset of ADHD have attained a very high level of evidential support. Some have strong evidence for a causal role but, for most, the possibility remains that these associations are due

to correlated genetic and environmental effects. For this reason, we refer to features of the pre- and post-natal environments that increase risk for ADHD as correlates, rather than causes. The genetic and environmental risks described below are not necessarily specific to ADHD.

Genetic Causes of ADHD

26. A review of 37 twin studies from the United States, Europe, Scandinavia, and Australia found that genes and their interaction with the environment must play a substantial role in causing ADHD (Faraone and Larsson, 2018; Larsson et al., 2014a; Pettersson et al., 2019).
27. In a genomewide study, an international team analyzed DNA from over 20,000 people with ADHD and over 35,000 without ADHD from the United States, Europe, Scandinavia, China, and Australia. They identified many genetic risk variants, each having a small effect on the risk for the disorder (Demontis et al., 2019). This study confirmed a polygenic cause for most cases of ADHD, meaning that many genetic variants, each having a very small effect, combine to increase risk for the disorder. The polygenic risk for ADHD is associated with general psychopathology (Brikell et al., 2020) and several psychiatric disorders (Lee PH, 2019).
28. Additional genes have been implicated by meta-analyses, but their status as risk genes remains uncertain until validated in a genomewide study. These genes are *ANKK1* (Pan et al., 2015) *DAT1* (Grunblatt et al., 2019b), *LRP5* and *LRP6* (Grunblatt et al., 2019a), *SNAP25* (Liu et al., 2017b), *ADGRL3* (Bruxel et al., 2020) *DRD4* and *BAIAP2* (Bonvicini et al., 2020; Bonvicini et al., 2016).

29. The polygenic risk for ADHD predicts ADHD symptoms in the population suggesting that the genetic causes of ADHD as a disorder, also influence lower levels of ADHD symptoms in the population (Demontis et al., 2019; Taylor et al., 2019).
30. In the population, those with a high polygenic risk for ADHD are more likely to have been diagnosed with ADHD (Li, 2019), anxiety or depression (Martin et al., 2018).
31. ADHD can also be the result of rare single gene defects (Faraone and Larsson, 2018) or abnormalities of the chromosomes (Cederlof et al., 2014). When the DNA of 8,000+ children with autism spectrum disorder (ASD) and/or ADHD and 5,000 controls was analyzed, those with ASD and those with ADHD had an increased rate of rare genetic mutations compared with controls (Satterstrom et al., 2019).
32. Family, twin, and DNA studies show that genetic and environmental influences are partially shared between ADHD and many other psychiatric disorders (e.g. schizophrenia, depression, bipolar disorder, autism spectrum disorder, conduct disorder, eating disorders, and substance use disorders) and with somatic disorders (e.g. migraine and obesity) (Demontis et al., 2019) (Faraone and Larsson, 2018) (Ghirardi et al., 2018) (Lee et al., 2019) (Lee et al., 2013) (Anttila et al., 2018; Tylee et al., 2018) (van Hulzen et al., 2017) (Vink and Schellekens, 2018) (Brikell et al., 2018) (Chen et al., 2019a) (Yao et al., 2019). However, there is also a unique genetic risk for ADHD. Evidence of shared genetic and environmental risks among disorders suggest that these disorders also share a pathophysiology in the biological pathways that dysregulate neurodevelopment and create brain variations leading to disorder onset.

33. Very large studies of families suggest that ADHD shares genetic or familial causes with autoimmune diseases (Li et al., 2019), hypospadias (Butwicka et al., 2015), and intellectual disability (Faraone and Larsson, 2018).

Environmental Correlates of ADHD: Exposure to Toxicants

34. A pair of meta-analyses found small correlations between lead burden and inattention symptoms (27 studies, over 9,300 youths) and hyperactivity-impulsivity symptoms (23 studies, over 7,800 youths) (Goodlad et al., 2013). A more recent meta-analysis of 14 studies with over 17,000 children reported that higher blood lead levels were associated with quadrupled odds of ADHD (Nilsen and Tolve, 2020). A study of over 2,500 youths from the National Health and Nutrition Examination Survey, a cross-sectional, nationally representative sample of the U.S. population, found that those with blood lead levels in the top third were 2.3 times more likely to have ADHD compared with those in the bottom third (Froehlich et al., 2009). A similar study, with over 4,700 youths from the same national survey, found that those with blood lead levels in the highest fifth were four times more likely to have ADHD compared with those in the bottom fifth (Braun et al., 2006).
35. Three meta-analyses with over twenty studies covering more than three million persons have found prenatal exposure to maternal smoking associated with a greater than 50% increase in incidence of ADHD (Huang et al., 2017) (Dong et al., 2018; Nilsen and Tolve, 2020). Although this association has also been seen in large population studies (Joelsson et al., 2016; Obel et al., 2016; Skoglund et al., 2014), it disappears after adjusting for family history of ADHD which indicates that the association between maternal smoking during

pregnancy and ADHD is due to familial or genetic factors that increase the risk for both smoking and ADHD.

36. A meta-analysis of nine studies spanning three continents and over 100,000 participants found that childhood exposure to secondhand cigarette smoke was associated with a 60% greater likelihood of ADHD. It was unclear to what extent the association was causal versus due to confounders (Huang et al., 2020).
37. In a meta-analysis of 15 double-blind, placebo-controlled trials with 219 participants, artificial food dyes were associated with a small increase in hyperactivity in children (Schab and Trinh, 2004). Another meta-analysis, covering 20 studies with a combined total of 794 individuals, found a very small increase in ADHD symptoms, but only when rated by parents, not by teachers or other observers (Nigg et al., 2012).
38. In a Taiwanese study of over 10,000 births, maternal use of acetaminophen during pregnancy was associated with a 33% greater likelihood of ADHD in their children (Chen et al., 2019b). Another study, examining 113,000 offspring from the Norwegian Mother and Child Cohort Study and the Norwegian Patient Registry, including 2,246 with ADHD, found a dose-response relationship between maternal prenatal use of acetaminophen and ADHD (Ystrom et al., 2017).
39. A nationwide study using the Danish national registers looked at 913,000 children born between 1997 and 2011. Prenatal exposure to the anti-epileptic drug valproate was associated with a 50% greater risk of ADHD. No associations were found for other anti-epileptic drugs (Christensen et al., 2019).

40. In a Norwegian registry study, 297 children with ADHD and 553 controls were randomly sampled from an eligible population of over 24,000. Children of mothers in the highest quintile of phthalate metabolite levels were three times more likely to have had ADHD as children compared with those in the bottom quintile, after adjusting for confounders, such as maternal age at delivery, sex of the child, maternal education, marital status, and prenatal maternal smoking (Engel et al., 2018).
41. Organophosphate pesticides are potent neurotoxins. In a sample of 1,139 children from the U.S. population, a tenfold increase in the organophosphate metabolite dimethyl alkylphosphate (DMAP) was associated with 55% increase in the probability of having ADHD. Children with detectable levels of the most-commonly detected DMAP metabolite were twice as likely to have ADHD compared with those with undetectable levels (Bouchard et al., 2010).
42. A meta-analysis found no significant effect of three classes of air pollutants –particulate matter (six studies, over 51,000 persons) and nitrogen oxides (five studies, over 51,000 persons) (Zhang et al., 2020b). A Taiwan-wide longitudinal cohort study geolinking over 16,000 mother-infant pairs to levels of air pollutants found no association between small particulate matter levels, sulphur dioxide levels, or nitrogen dioxide levels during gestation and ADHD diagnoses in the first eight years of their offsprings' lives. It did find a 25 percent greater odds for having ADHD with exposures to nitric oxide, a common traffic pollutant (Shih et al., 2020).

43. A nationwide cohort study used the South Korean national health insurance registry to identify all 7,200 hospital admissions of adolescents with a primary diagnosis of ADHD from 2013 to 2015, and daily readings of three air pollutants from 318 monitoring stations distributed across the country over the same period. It found that spikes in nitrogen dioxide, sulphur dioxide, and particulate matter were associated, respectively, with 47%, 27%, and 12% increases in ADHD related hospital admissions in succeeding days. There were no significant differences between male and female adolescents, or between older and younger adolescents (Park et al., 2020).
44. A meta-analysis of nine European population studies encompassing 4,826 mother-child pairs examined the relationship between exposure to Perfluoroalkyl Substances (PFAS) via breast milk in infancy and development of ADHD. No associations were found with ADHD in offspring (Forns et al., 2020).
45. A meta-analysis of seven studies encompassing a total of over 25,000 participants from six countries on three continents found no evidence of an association between sugar consumption and ADHD in youth (Farsad-Naeimi et al., 2020)

Environmental Correlates of ADHD: Nutrient Deficiencies

46. A pair of meta-analyses found no difference in serum iron levels in youths with ADHD (six studies, 617 participants) but small-to-moderate reductions in serum ferritin, a protein that stores iron (ten studies, over 2,100 participants) (Wang et al., 2017). Another pair of meta-analyses likewise found no difference in serum iron levels (six studies, over 1,700

participants) but small-to-moderate reductions in serum ferritin (12 studies, over 6,000 participants) (Tseng et al., 2018).

47. A meta-analysis of nine studies and 586 people found moderately lower overall blood levels of omega-3 PUFAs in ADHD than non-ADHD youth (Hawkey and Nigg, 2014).
48. A nationwide population-based case-control study using the Finnish national registers compared 1,067 patients with ADHD born between 1998 and 1999 with 1,067 matched controls. Lower maternal vitamin D levels were associated with a roughly 50% greater likelihood of ADHD in their children (Sucksdorff et al., 2019).

Environmental Correlates of ADHD: Events During Pregnancy and Birth

49. A meta-analysis of twelve studies with over 6,000 participants found a threefold increase in the rate of ADHD among very/extremely preterm or very/extremely low birth weight babies (Franz et al., 2018). Another meta-analysis, combining 85 studies with a total of over 4.6 million births, found a small-to-moderate correlation between low birth weight and ADHD (Momany et al., 2018). A Swedish national register study of 1.2 million children found a stepwise increase in the likelihood of ADHD with increasing prematurity. Results were not due to having an ADHD relative or socioeconomic stress (Lindstrom et al., 2011). Similar results were reported from the Finnish national registers when comparing over 10,000 people with ADHD with over 38,000 controls (Sucksdorff et al., 2015).
50. A meta-analysis of six studies combining 1.4 million people found that children whose mothers had hypertensive disorders during pregnancy had a 25% increase in the rate of ADHD (Maher et al., 2018).

51. A nationwide population-based cohort study using Swedish registers and covering more than two million children, 115,000 of them with ADHD, found that maternal preeclampsia during pregnancy is associated with a 15% greater subsequent likelihood of ADHD in offspring, rising to over 40% when the fetus is small for gestational age and exposed to preeclampsia. This pattern in families showed that it is not due to genetic or other family influences (Maher et al., 2020).
52. Two meta-analyses, one with seven studies with over 28,000 participants and another with three studies and over 1.4 million participants, found that children of obese mothers were roughly 60% more likely to develop ADHD (Jenabi et al., 2019; Sanchez et al., 2018). A study of over 80,000 mother-child pairs participating in the Danish National Birth Cohort reported an almost 50% elevated risk of ADHD in children of obese mothers and a doubled risk in children of severely obese mothers (Andersen et al., 2018).
53. A meta-analysis of two large cohort studies with a combined total of over 3.1 million persons found a slight but significant association between maternal hyperthyroidism during pregnancy and ADHD in offspring. A second meta-analysis of four cohort studies encompassing over 3.4 million participants likewise found a slight but significant association between maternal hypothyroidism and ADHD in offspring. No attempt was made to assess the role of confounders (Ge et al., 2020).
54. A nationwide cohort study using Danish national registers examined over a million births, comparing offspring of mothers with a single prior miscarriage and mothers with more than one prior miscarriage with mothers with no history of miscarriage. It found that after adjusting for a wide range of possible confounders which turned out to have little effect,

children of mothers with a single prior miscarriage were 9% more likely to develop ADHD than those of mothers without any miscarriage. Children of mothers with two or more prior miscarriages were 22% more likely to be diagnosed with ADHD. This upward exposure-response trend was statistically significant (Wang et al., 2020).

Environmental Correlates of ADHD: Deprivation, Stress, Infection, Poverty and Trauma

55. A Taiwan-wide longitudinal cohort study based on the country's universal coverage National Health Insurance Research Database compared over 14,000 enterovirus patients (ER71) with an equal number of controls matched by age and sex. After further adjusting for paternal occupation and urbanization level of residence it found the enterovirus patients were 25 percent more likely to subsequently be diagnosed with ADHD (Tseng et al., 2020).
56. A nationwide population-based cohort study using Danish registers compared over 29,000 children born to women who lost a close relative during pregnancy with a million other children in the same cohort and found that boys born to these women were twice as likely to have ADHD (Li et al., 2010).
57. A U.S. population-based study of over 14,000 participants in the National Longitudinal Study of Adolescent Health found that after adjusting for demographic, socioeconomic, and familial risk factors for child maltreatment, ADHD inattentive type was associated with having been exposed to sexual abuse and physical neglect (Ouyang et al., 2008).
58. A nationwide population-based cohort study of over 18,000 children from the South Korean National Health Insurance database found that lower levels of family income were associated with increased rates of ADHD (Choi et al., 2017). A Swedish study of over

800,000 people reported similar results even after adjusting for shared familial/genetic risk factors in families (Larsson et al., 2014b).

59. A Danish national register longitudinal cohort study of a million people found that Rutter's indicators of adversity were predictive of ADHD. Out-of-home care was strongly predictive; low social class, paternal criminality, maternal mental disorder, and severe marital discord were moderately predictive. Large family size had no effect (Ostergaard et al., 2016).
60. A countrywide population study using Danish national registers looked at over 630,000 youths and found dose-response relationships between lower parental educational attainment, parental unemployment, and parental relative poverty and higher risk of ADHD in offspring. Combinations of social disadvantages had cumulative risks. For instance, parental relative income poverty plus completion of no more than compulsory education plus unemployment was associated with a roughly five percent higher risk of ADHD in their offspring (Keilow et al., 2020).
61. A Swedish national register cohort study of over 540,000 people found a dose-response relationship between cumulative indicators of adversity in the family and ADHD. A death in the family increased the subsequent likelihood of ADHD by 60%. Substantial parental substance abuse, criminality, or psychiatric disorder each more than doubled the likelihood as did residential instability and household public assistance (Bjorkenstam et al., 2018).
62. In a sample of 4,122 U.S. youths with ADHD from the 2016 U.S. National Survey of Children's Health, greater family cohesion and community support decreased the risk for moderate to severe ADHD (Duh-Leong et al., 2020).

What Have We Learned from Studying the Brains of People with ADHD?

There are two broad classes of research findings about the brains of people with ADHD. The first comes from studies of the performance of patients on psychological tests that study mental processes. The second comes from methods that directly examine brain structure or function with neuroimaging scans. Although many of these studies have found differences between groups of people who are and are not diagnosed with ADHD, the differences are typically small and do not dramatically differ between people with ADHD and those with other disorders. They are, therefore, not useful for diagnosing the disorder (Thome et al., 2012). These differences are not caused by drug treatment and, for some patients, diminish or change as patients grow out of the disorder.

Performance Deficits in Psychological Processes

63. A meta-analysis of 137 studies with over 9,400 participants of all ages found ADHD to be associated with moderately lower IQ and reading scores and larger decreases in spelling and arithmetic scores (Frazier et al., 2004). Another meta-analysis, spanning 21 studies with over 1,900 adults, concluded that ADHD-associated IQ deficits were small and not clinically meaningful (Bridgett and Walker, 2006).
64. A series of meta-analyses found that people with ADHD had small to moderate difficulties with abstract problem solving and working memory (12 studies, 952 persons), focused attention (22 studies, 1,493 persons), sustained attention (13 studies, 963 persons), and verbal memory (8 studies, 546 persons) (Schoechlin and Engel, 2005). Another meta-

analysis, with 11 studies with 829 participants, reported people with ADHD were moderately more prone to cognitive errors known as “rule violations” (Patros et al., 2019).

65. Two meta-analyses, one with 21 studies and over 3,900 participants, the other with 15 studies with over a thousand participants, found that those diagnosed with ADHD have a moderate tendency to favor small immediate rewards over large delayed rewards (Jackson and MacKillop, 2016; Marx et al., 2018).
66. A meta-analysis of 37 studies with more than 2,300 participants found a small-to-moderate association between ADHD and risky decision-making (Dekkers et al., 2016). Another meta-analysis, combining 22 studies with 3,850 children and adolescents, found those with ADHD exhibited moderately greater impulsive decision-making overall on delay discounting and delay of gratification tasks (Patros et al., 2016).
67. A recent meta-meta-analysis included 34 meta-analyses of neurocognitive profiles in ADHD (all ages) concerning 12 neurocognitive domains. Those with ADHD had moderate impairments in multiple domains (working memory, reaction time variability, response inhibition, intelligence/achievement, planning/organization). Effects were larger in children and adolescents than in adults (Pievsky and McGrath, 2018).
68. A meta-analysis of 49 studies and over 8,200 children and adolescents found moderate impairments in working memory in those with ADHD. These deficits declined with age (Ramos et al., 2020).
69. Among youths with ADHD, a series of meta-analyses found no significant sex differences in either total ADHD symptoms (15, studies, over 3,400 youths), inattention symptoms (26

studies, over 5,900 youths), or hyperactivity-impulsivity symptoms (24 studies, over 4,900 youths) (Loyer Carbonneau et al., 2020).

70. A meta-analysis of randomized controlled trials (RCTs) with preschoolers found that cognitive training led to moderate improvement in working memory (23 studies, over 2,000 participants) and small-to-moderate improvement in inhibitory control (26 studies, over 2,200 participants) (Pauli-Pott et al., 2020).

Differences in the Brain Found by Neuroimaging Studies

71. An analysis of structural magnetic resonance imaging (MRI) data from 36 cohorts with a total of over 4,100 participants found slightly reduced total cortical surface area in children with ADHD. The same team found some subcortical regions of the brain were smaller in children with ADHD, mainly in frontal, cingulate and temporal regions with some reductions in cortical thickness in temporal regions. The same team found some subcortical regions of the brain, i.e., basal ganglia, amygdala, hippocampus and intracranial volumes were smaller in children with ADHD in 23 cohorts of 3,242 participants. The differences seen in children were not seen in adolescents or adults (Hoogman et al., 2017; Hoogman et al., 2019). All of the differences observed were small to very small and subtle.
72. Comparative meta-analyses show that structural grey matter volume reductions in basal ganglia and insula are disorder-specific relative to OCD in 30 data sets with 1,870 participants (Norman et al., 2016) while medial frontal reductions were specific to ASD in 66 data sets with 3,610 participants (Lukito et al., 2020). An analysis of structural magnetic resonance imaging (MRI) data from 48 cohorts with a total of over 12,000 participants

showed that ADHD participants had smaller hippocampus volume relative to OCD which was related to IQ differences and smaller intracranial volume relative to ASD and OCD patients (Boedhoe et al., 2020). The functional under-activations in right inferior frontal cortex and basal ganglia during tasks of cognitive control were disorder-specific relative to OCD in 1,870 participants (Norman et al., 2016), while the inferior frontal dysfunction was specific relative to autism in 3,610 participants (Lukito et al., 2020).

73. A meta-analysis of ten diffusion tensor imaging studies with 947 participants found that the most consistent white matter differences between those with and without ADHD were located in the splenium of the corpus callosum extending to the right cingulum, the right sagittal stratum, and left tapetum, suggesting problems with the connections between the two hemispheres in posterior parieto-temporal attention regions and in long-range fronto-posterior association tracts (connecting inferior frontal, temporal, parietal and occipital regions) involved in attention and perception (Chen et al., 2016).

74. A meta-analysis of 21 functional MRI studies with 607 participants found that those with ADHD showed consistent and replicable under-activation in typical regions of inhibitory control such as right inferior frontal cortex, supplementary motor area and the basal ganglia relative to typically developing individuals (Hart et al., 2013). The inferior frontal under-activation findings were replicated in two further fMRI meta-analyses of inhibitory control with 33 datasets/1,161 participants, and 42 datasets/2,005 participants, respectively (Lukito et al., 2020; Norman et al., 2016). Another meta-analysis including 130 fMRI studies with 1,914 participants found no convergence except for aberrant function in basal ganglia for neutral fMRI tasks and inferior frontal under-function in males only (Samea et al., 2019).

75. A meta-analysis of nine studies with over 1,250 research participants found that elevations in the theta/beta on the electroencephalogram cannot be considered a reliable diagnostic measure for ADHD although it may have prognostic value in some patients (Arns et al., 2013).
76. A meta-analysis of six studies with 148 participants examined mismatch negativity, which assesses the integrity of auditory sensory memory and involuntary attention switching. It reported that ADHD children had small-to-moderate reductions in mismatch negativity amplitude compared with healthy controls (Cheng et al., 2016).
77. Meta-analyses and systematic reviews showed that the medications used to treat ADHD are not associated with observed deficits in brain structure (Hoogman et al., 2017; Hoogman et al., 2019; Lukito et al., 2020; Norman et al., 2016; Spencer et al., 2013), but with improved brain function, most prominently in inferior frontal and striatal regions (Hart et al., 2013; Lukito et al., 2020; Norman et al., 2016; Rubia et al., 2014; Spencer et al., 2013).

What kinds of Non-Psychiatric Medical Problems Commonly Occur among People with ADHD?

A relatively new area of research into ADHD is examining what types of medical problems are more common than expected among people with ADHD. As you read this section, keep in mind that not all people with ADHD will suffer from all, or even only one, of these disorders.

Obesity

78. A Swedish national register study of over 2.5 million people found ADHD patients had a threefold greater risk of obesity relative to their non-ADHD siblings and cousins. It also found a familial co-aggregation of ADHD and clinical obesity, the strength of which varied directly with the degree of genetic relatedness (Chen et al., 2018c).
79. A meta-analysis found that compared with typically developing people, children and adolescents with unmedicated ADHD were about 20% more likely to be overweight or obese (15 studies, over 400,000 participants), and adults with unmedicated ADHD almost 50% more likely to be overweight or obese (9 studies, over 45,000 participants) (Nigg et al., 2016). Meta-analyses of twelve studies with over 180,000 participants found that people with unmedicated ADHD were about 40% more likely to be obese, whereas those who were medicated were indistinguishable from typically developing people (Cortese et al., 2016b).

Allergies and Asthma

80. A Swedish national register study of over 1.5 million people found that those with asthma were 45% more likely to have ADHD even after adjustment for relevant variables (Cortese et al., 2018b). A cohort study of almost a million births using the Danish national registers found that children born to asthmatic mothers were 40% more likely to develop ADHD (Liu et al., 2019b).
81. In a meta-analysis of six longitudinal studies with over 50,000 participants, those with asthma or atopic eczema were a third more likely to have ADHD than controls. A meta-

analysis of three studies with over 48,000 participants found that those with allergic rhinitis were about 50% more likely to have ADHD (van der Schans et al., 2017).

Diabetes Mellitus:

82. A retrospective analysis of over 650,000 children and adolescents in German diagnosis and prescription databases found ADHD was 40% more likely to be diagnosed among children with type 1 diabetes (T1DM) (Kapellen et al., 2016).
83. A German multi-center registry study of over 56,000 children and adolescents found that those with both ADHD and T1DM suffered twice as often from diabetic ketoacidosis compared with diabetic patients without ADHD. They also found significant differences in HbA1c, and concluded, "Pediatric patients with ADHD and T1DM showed poor metabolic control compared with T1DM patients without ADHD" (Hilgard et al., 2017).
84. A longitudinal study using the Taiwan National Health Insurance Research Database enrolled over 35,000 patients with ADHD and over 70,000 age- and sex-matched controls. Adolescents and young adults with ADHD were about three times more likely to develop type 2 diabetes mellitus (Chen et al., 2018b).
85. A cohort study using multiple Swedish national registers looked at over 1.6 million adults aged 50 to 64 years. Prevalence of type 2 diabetes mellitus was 70% greater among adults with ADHD (Chen et al., 2018c).
86. A meta-analysis found that maternal pre-existing type 1 diabetes was associated with a small increased risk of ADHD in offspring (4 studies, over five million people). So was

paternal pre-existing type 1 diabetes (3 studies, 4.7 million people), and maternal pre-existing type 2 diabetes (2 studies, 2.6 million people) (Zeng et al., 2019). A Swedish study looked at all 15,615 children born after their parents were diagnosed with type 1 diabetes. After controlling for confounders, it found that these children had a 30% greater chance of being diagnosed with ADHD (Ji et al., 2018).

Other Somatic Disorders:

87. A meta-analysis of 18 studies with more than 2,500 children and adolescents found a moderate association between sleep-disordered breathing and ADHD (Sedky et al., 2014).
88. A meta-analysis of sleep in adults with ADHD found no significant differences with normally developing adults, as measured by polysomnography. In four studies with 178 participants, sleep onset latency, stage 1 sleep, stage 2 sleep, slow wave sleep, REM, and sleep efficiency were all comparable. Same with total sleep time (3 studies, 130 persons), and with REM latency and wake after sleep onset (3 studies, 121 persons). As measured by actigraphy, there were no significant differences for time in bed and actual wake time (3 studies, 159 persons), true sleep (4 studies, 222 persons). However, sleep onset latency was much greater for those with ADHD, and sleep efficiency was moderately lower (4 studies, 222 persons). Nevertheless, subjective evaluations by those with ADHD reported moderately greater difficulty in falling asleep (8 studies, over 1,700 persons), moderately greater frequency of night awakenings and moderately lesser likelihood of being rested at wake-up (5 studies, over 1,100 persons), and moderately worse sleep quality (5 studies, over 800 persons)(Lugo et al., 2020).

89. In a Norwegian national registry study of over 1.2 million males and over 1.2 million females, males with ADHD were 30% more likely to be diagnosed with psoriasis, and women with ADHD more than 50% more likely to be diagnosed with psoriasis, than normally developing controls (Hegvik et al., 2018).
90. A Taiwan nationwide population cohort study of over 8,000 people with ADHD and 32,000 matched controls explored associations with autoimmune diseases. It reported that those with ADHD had well over twice the prevalence of ankylosing spondylitis, ulcerative colitis, and autoimmune thyroid disease, and over 50% greater likelihood of asthma, allergic rhinitis, and atopic dermatitis (Chen et al., 2017a) .
91. A population-based cohort study of over 900,000 Danish children found that epilepsy was associated with a 2.7-fold increased risk for ADHD (Bertelsen et al., 2016). Another population-based cohort study, of over 12,000 Taiwanese, reported that epilepsy was associated with a 2.5-fold increased risk for ADHD. Conversely, a linked cohort study of over 18,000 Taiwanese found ADHD was associated with a fourfold increase in epilepsy (Chou et al., 2013).
92. A countrywide registry study of 1.9 million Swedes reported that those with epilepsy were three and a half times more likely to have ADHD. The risk of having ADHD was 85% greater if the person's mother had epilepsy, 50-60% greater if the father or a brother or sister did, 15% greater for cousins. Genetics explained 40% of the variance, with non-shared environmental factors explaining another 50% (Brikell et al., 2018).

93. A longitudinal study using the Taiwan Health Insurance Research Database compared almost 18,000 adolescents and young adults with ADHD with over 70,000 age- and sex-matched controls. Those with ADHD were over three times more likely to develop sexually transmitted infections, after adjusting for demographic data, other psychiatric disorders, and ADHD medications (Chen et al., 2018a).
94. A Danish national register cohort study of 1.1 million people found that hospitalization for serious infections was associated with a subsequent doubling in the rate of ADHD diagnosis. Among those treated with anti-infective agents, the risk of subsequent diagnosis with ADHD was halved (Kohler-Forsberg et al., 2019).
95. A Danish national register study of almost a million people found that children with autoimmune disease were 24% more likely to develop ADHD. Maternal autoimmune disease was associated with a 12% greater likelihood of ADHD in their offspring. Paternal autoimmune disease was not associated with any significant effect (Nielsen et al., 2017).
96. Using Taiwan's nationwide population-based dataset, over 116,000 children with ADHD were compared with the same number of randomly selected children without ADHD. Those with ADHD were much more likely to have significant abnormalities of the eye: almost 90% more likely to have amblyopia ("lazy eye"), over 80% more likely to have astigmatism, and twice as likely to have heterotropia, in which the eyes diverge at rest (Ho et al., 2020). A study using the same database matched 6,817 youths diagnosed with amblyopia to over 27,000 age- and sex-matched controls. Those in the amblyopia group had 1.8 times the risk of developing ADHD (Su et al., 2019).

97. In a study of over 2.5 million German youth, those with ADHD were nine times more likely to have metabolic disorders, five times more likely to develop viral pneumonia, four times more likely to have white blood cell disorders, three times more likely to have kidney failure, high blood pressure, or be obese, two and a half times more likely to have type 2 diabetes or migraines, twice as likely to have asthma or atopic dermatitis, and 50% more likely to have glaucoma (Akmatov et al., 2019). A Brazilian population-based study including 5,671 children found those with migraine about four times more likely to have ADHD (Arruda et al., 2020).
98. A study of over 59,000 boys diagnosed with ADHD and over 52,000 healthy boys in Taiwan reported that those in the ADHD group were twice as likely to develop testicular dysfunction (Wang et al., 2019).
99. A nationwide population cohort study using the Swedish national registers compared over 19,000 children with a diagnosis of biopsy-verified celiac disease with over 95,000 matched child controls. It found a subsequent 29% increased risk of ADHD in the celiac patients, rising to 39% when restricting to adult diagnoses of ADHD. However, when comparing 13,000 children diagnosed with celiac disease to their 18,000 non-celiac siblings, the increases became nonsignificant, suggesting the increases were primarily attributable to confounders (Lebwohl et al., 2020).
100. A Swedish nationwide study using national registers examined medical records of all individuals aged 18 to 64 years who were residing in Sweden during 2013 and identified 41,840 who filled at least one prescription for ADHD medicines. Young adults with ADHD

were four times more likely to have somatic co-prescriptions and fifteen times more likely to have psychotropic co-prescriptions than normally developing controls. For middle-aged adults (30-49) the odds were six and 21 times greater, respectively, and for older adults, seven and 18 times greater. Respiratory medications (primarily for allergies reactions and asthma) were the most likely to be dispensed for somatic purposes, followed by alimentary tract and metabolic medications (most frequently proton pump inhibitors indicated for gastric/duodenal ulcers and gastroesophageal reflux disease), then cardiovascular system medications (primarily for hypertension and arrhythmias) (Zhang et al., 2020a).

What is the Impact of ADHD on Patients and Families?

ADHD is a disorder associated with serious distress and/or impairments in living. Although, as documented below, many severe adverse outcomes have been associated with ADHD, the typical patient does not experience all, or even most, of these problems. Many patients live enjoyable and productive lives, especially if they receive treatment.

Quality of Life

101. A meta-analysis of seven studies with over 5,000 youths and their parents reported large impairments in the quality of life of youths with ADHD relative to typically developing peers, regardless of whether evaluated by the youths themselves or by their parents.

Physical functioning was only moderately impaired, but emotional functioning and social functioning was strongly impaired. School functioning was strongly impaired. As youths with

ADHD grew older, their quality of life compared with typically developing peers grew worse in physical, emotional, and school domains. (Lee et al., 2016).

102. A meta-analysis of 17 studies encompassing 647 families (over 2,300 participants) evaluated the quality of life of parents whose children had ADHD relative to parents with typically developing children. Parents of the former reported a moderate deficit in quality of life relative to parents of the latter (Dey et al., 2019).

Emotional and Social Impairment

103. A study of over 8,600 youths from the US National Health Interview Survey found that those with ADHD were six times as likely to have a high level of emotional, conduct, and peer problems, and nine times as likely to manifest a high level of impairment including interference with home life, friendships, classroom learning, and leisure activities (Strine et al., 2006).
104. A meta-analysis of 22 studies with over 21,000 participants found that youth with ADHD were strongly impaired in the ability to modulate their reactivity to novel or stressful events (Graziano and Garcia, 2016). Another meta-analysis, combining twelve studies with over 1,900 participants, found that adults with ADHD had very elevated levels of emotional dysregulation compared with normally developing controls (Beheshti et al., 2020).
105. A meta-analysis found that children with ADHD had medium-to-large impairments in socializing with peers as measured by rejection/likability, popularity, and friendships (61 studies, over 24,000 children). They also had moderate impairments in social skills such as sharing, cooperating, turn-taking, reciprocity (68 studies, over 148,000 children), and social-

information processing, such as recognizing social cues, identifying problems, generating solutions, and avoiding biases (23 studies, over 3,750 children) (Ros and Graziano, 2018).

106. A study of over 53,000 U.S. children from the National Survey of Children's Health found that those with ADHD were 2.4 times as likely to engage in bullying (Montes and Halterman, 2007). A more recent study of some 64,000 children using the same database confirmed this finding, reporting that those with ADHD were 2.8 times more likely to engage in bullying (Benedict et al., 2015).

Accidental Injuries

107. A nationwide cohort study of over 50,000 youths with ADHD and an equal number of age-, sex-, and comorbidity-matched controls drawn from Taiwan's National Health Insurance Research Database reported that having ADHD was associated with a more than three-quarters greater likelihood of burn injury. For those under six years old, the risk was doubled. For youths between six and seventeen years old, the increase in risk was about 70 percent. There were no significant differences between boys and girls (Yeh et al., 2020).
108. A meta-analysis of 32 studies covering more than four million people found that those with ADHD had a 40 to 50% greater risk of accidental physical injuries (Ruiz-Goikoetxea et al., 2018a).
109. A Swedish national registers study followed 17,408 individuals with ADHD from 2006 to 2009 and found that patients with ADHD had an almost 50% greater risk of serious transport accidents (Chang et al., 2014b).

110. A U.S. study of over 8,000 high school and collegiate athletes (predominantly male football players) found that those with ADHD were three times as likely to have had three or more reported concussions (Nelson et al., 2016).
111. A meta-analysis of 16 studies encompassing over 175,000 people estimated that controlling for mileage driven, those with ADHD were 23% more likely to be involved in vehicular crashes (Vaa, 2014).
112. A retrospective cohort study of over 18,000 New Jersey drivers found that the crash risk for those with ADHD was a third greater than for those without (Curry et al., 2017).
113. A meta-analysis of five studies, comprising over three thousand patients with minor traumatic brain injury (mTBI) and over nine thousand controls found that those with mTBI were twice as likely to have ADHD than those without mTBI (Adeyemo et al., 2014).

Premature Death and Suicide

114. A Danish study of almost two million people found ADHD is associated with a small risk for premature death, mostly due to accidents. When ADHD was accompanied by other psychiatric and substance use disorder, the chances of premature death increased (Dalsgaard et al., 2015b).
115. A cohort study of more than 2.2 million Taiwanese found no increased risk of death from natural-causes associated with ADHD. But people with ADHD had twice the rate of suicide, twice the rate of death by homicide, and a 30% greater rate of death from unintentional injury (Chen et al., 2019c).

116. Using nationwide registers in Denmark, a cohort study of 2.9 million people reported a fourfold higher rate of suicide attempts and deaths in patients with ADHD. The risk was over tenfold in those with ADHD plus another psychiatric diagnosis (Fitzgerald et al., 2019).
117. A meta-analysis found that persons with ADHD attempted suicide at twice the rate of typically developing people (six studies, over 65,000 persons), had over three times the rate of suicidal ideation (23 studies, over 70,000 persons), and over six times the rate of completed suicide (four studies, over 130,000 persons) (Septier et al., 2019).
118. A Taiwanese study of over 20,000 adolescents and young adults with ADHD and over 61,000 age- and sex-matched non-ADHD individuals found that those with ADHD were almost four times as likely to attempt suicide, and over six times as likely to repeat suicide attempts. Methylphenidate or atomoxetine treatment did not increase the risk of suicide attempts or repeated suicide attempts. Long-term methylphenidate treatment was associated with a lower risk for repeated suicide attempts among men (Huang et al., 2018).
119. In a prospective cohort study of more than 2.6 million Swedes, adults with ADHD had a small increase in premature death, mostly due to accidents and suicide. There was no significant association for children with ADHD (Sun et al., 2019b).

Crime and Delinquency

120. A study of the Danish population using nationwide registers found that, compared with other youth, those diagnosed with ADHD were more than twice as likely to be convicted of criminal offenses and were three times as likely to be incarcerated. After adjusting for other

risk factors, those with ADHD were 60% more likely to have been convicted of a crime, and 70% more likely to have been incarcerated (Mohr-Jensen et al., 2019).

121. A meta-analysis comprising 21 studies and over 19,500 prison inmates found that the prevalence of ADHD in prisons based on interview diagnoses was 20.5% with no differences observed between males and females or adolescents and adults (Young et al., 2015). Another meta-analysis reported the prevalence of ADHD among adolescents in juvenile detention to be just over 17%, both for males (24 studies, over 24,000 individuals) and females (13 studies, over 3,900 individuals), which is much higher than the prevalence in the population (Beaudry et al., 2020).
122. A study using a nationally representative American sample of over 5,000 adults found that those with ADHD were over twice as likely to be perpetrators of physical dating violence, and 65% more likely to be victims of such violence (McCauley et al., 2015).
123. In a nationwide study of over 21,000 Icelandic adolescents and young adults, 14% reported having been interrogated at a police station. Of these, 15% reported making a false confession. Those with ADHD were twice as likely to make a false confession (Gudjonsson et al., 2016).
124. A study using the Danish national registries looked at violent crimes against youth aged 7-18 years, among a total of 678,000 individuals. Children with ADHD were 2.7 times more likely to be victims of violent crimes than their typically developing peers, after adjusting for confounding risk factors (Christoffersen, 2019).

Educational Underachievement

125. A study of a U.S. sample of almost 30,000 adults found that those with ADHD were twice as likely not to have graduated from high school on time, after adjusting other psychiatric disorders (Breslau et al., 2011).
126. A nationwide cohort study of over 750,000 Scottish school children using linked national registers identified those who had been prescribed medicine for ADHD. Even while receiving medication, these children were more than three times as likely as typically developing peers to have low educational achievement, more than twice as likely to drop out of school before age 16, more than eight times as likely to have a record of special educational needs, 50% more likely to get injured, 40% more likely to be unemployed. These results were adjusted for socioeconomic confounders and other psychiatric conditions (Fleming et al., 2017).
127. A meta-analysis of ten studies and 830 youths found that ADHD was strongly associated with poorer performance on measures of overall, expressive, receptive, and pragmatic language (Korrel et al., 2017).

Substance Use Disorders

128. A meta-analysis of twelve studies covering over 5,400 people found that those with ADHD were almost three times more likely to be nicotine-dependent. Combining eleven studies with almost 2,400 participants, those with ADHD were 50% more likely to develop a drug or alcohol use disorder than those without ADHD (Lee et al., 2011).

129. A meta-analysis found that ADHD was associated with a more than twofold greater odds of alcohol-use disorders (13 studies, over 20,000 participants) and nicotine-related disorder (14 studies, over 1,800 participants) (Groenman et al., 2017).
130. A Swedish study of over half a million people found a more than threefold association between ADHD and subsequent drug use disorders after adjusting for sex and parental education (Sundquist et al., 2015).

Other

131. Studies of 2.7 million girls from Denmark (Ostergaard et al., 2017), 380,000 from Sweden (Skoglund et al., 2019) and 7,500 from Taiwan (Hua et al., 2020) found that those with ADHD were more likely to have teen pregnancies than those without ADHD. Consistent with these results, large studies from Sweden (Chang et al., 2014a), Finland (Chudal et al., 2015) and a consortium of eight European countries (Pohlabeln et al., 2017) each found ADHD to be more likely among children of teenage mothers than among children of older mothers.
132. A study of over 36,000 people from the US reported that ADHD increased the risks for problem gambling, spending too much money, reckless driving, and quitting a job without a plan for what to do next (Bernardi et al., 2012).
133. A nationwide study using Taiwan's National Health Insurance Research Database compared 675 adults with ADHD and 2,025 without ADHD, matched by age and sex. After adjusting for other psychiatric disorders, urbanization level of residence, and monthly income, those with ADHD had 3.4 times the risk of developing dementia (Tzeng et al., 2019).

134. A meta-analysis of nine studies encompassing almost a million and a half people found that ADHD is associated with a threefold greater risk of poisoning in children (Ruiz-Goikoetxea et al., 2018b). In a study from Taiwan comparing 3,685 children with ADHD with 36,000 controls, those with ADHD had a more than fourfold greater risk of deliberate self-poisoning (Chou et al., 2014).
135. A longitudinal study of some 15,000 U.S. adolescents reported that those with ADHD had a 12% reduction in employment and a 34% reduction in earnings relative to non-ADHD siblings (Fletcher, 2014).
136. Using Danish registers, a nationwide population study of over 675,000 youths between the ages of 7 and 18 found that youths with ADHD were 3.7 times as likely to be reported as victims of sexual crimes than normally developing controls. After adjusting for covariates, such as parental violence, parental inpatient mental illness, parental suicidal behavior or alcohol abuse, parental long-term unemployment, family separation, and child in public care outside the family, youths with ADHD remained almost twice as likely to be reported as victims of sexual crimes (Christoffersen, 2020).

What is the Economic Burden of ADHD?

Given the many adverse outcomes associated with ADHD, it will come as no surprise to readers that these effects have a substantial economic cost to individual patients, families, and society.

137. A systematic review of seven European studies of hundreds of thousands of participants estimated total ADHD-related costs in the Netherlands as €9,860 to €14,483 per patient per year, with annual national costs more than €1 billion (Le et al., 2014).

138. A review of the costs of child, youth and adult ADHD in Australia estimated the total annual costs to be over \$20 billion Australian dollars, or \$25,000 per person with ADHD. This includes financial costs of \$12.8 billion, well-being losses of \$7.6 billion, and productivity losses of \$10.2 billion (Australian ADHD Professionals Association, 2019).
139. A systematic review of 19 U.S. studies of hundreds of thousands of people found that ADHD was associated with overall national annual costs from \$143 to \$266 billion, mostly associated with adults (\$105 to \$194 billion). Costs borne by family members of people with ADHD ranged from \$33 - \$43 billion (Doshi et al., 2012).
140. A study with over 7,000 workers in ten nations found that those with ADHD had an average of 22 annual days of lost role performance compared with those without ADHD (de Graaf et al., 2008).
141. A study of a U.S. national Fortune 100 company's database of over 100,000 beneficiaries compared healthcare costs for youths with ADHD with matched controls without ADHD. The annual average cost per family member was \$2,728 for non-ADHD family members of ADHD patients, almost double the \$1,440 for family members of matched controls (Swensen et al., 2003).
142. German health insurance records, including over 25,000 patients with ADHD, indicate that patients with ADHD cost roughly €1,500 more annually than those without ADHD. Main cost drivers were inpatient care, psychiatrists, and psychotherapists. Mood, anxiety, substance use disorders, and obesity were significantly more frequent in patients with

ADHD. The additional costs resulting from these conditions added as much as €2,800 per patient (Libutzki et al., 2019).

143. Using the National Health Insurance Service claims data for the population aged 19 years or younger in South Korea (69,353 diagnosed with ADHD), the total annual economic burden due to ADHD was estimated to be \$47.55 million (Hong et al., 2020).
144. Using the Danish national registers, over 5,000 adults with a diagnosis of ADHD in adulthood who had not received a diagnosis in childhood were identified. Excluding cases with missing data, other psychiatric diagnoses, and cases without a same-sex sibling free of any diagnosed psychiatric diagnoses, a final cohort was formed consisting of 460 sibling pairs. On average, adults with ADHD had an annual economic burden of just over €20,000 compared with their normally developing siblings (Daley et al., 2019).
145. A nationwide cohort study of over 445,000 people in the Swedish national registers compared healthcare costs for three groups: those with childhood ADHD that persisted into adulthood, those whose ADHD remitted in adulthood, and those who never had ADHD. Those who never had ADHD had average annual healthcare costs of €304. Those in remission had double the cost, and those with persistent ADHD over triple the cost (Du Rietz et al., 2020).
146. A nationwide population study of over 83,000 persons with ADHD and 334,446 non-ADHD controls matched by age and sex used Danish national registries to calculate the net socioeconomic cost of ADHD. Relative to controls, and summing net direct health costs and net losses from lower income and employment, the yearly average cost per individual with

ADHD came to just over €16,000. Including additional social transfers, the total rose to just over €23,000. For partners of persons with ADHD, the additional yearly average cost per individual was almost €5,500. With additional social transfers, the total rose to €8,000 (Jennum et al., 2020).

147. Using a database that tracks more than sixty German nationwide health insurance programs, a study of five million member records identified 2,380 individuals first diagnosed with ADHD as adults. Their direct healthcare costs in the year following diagnosis averaged €4,000. Despite explicit German guidelines recommending ADHD medication, only a third were prescribed medication, dropping to one eighth four years later. Two-thirds received psychotherapy. The authors concluded that "guideline recommendations are not yet comprehensively implemented in everyday routine care" (Libutzki et al., 2020).

Which Medications are Safe and Effective for Treating ADHD?

As determined by governmental regulatory agencies around the world, several medications are safe and effective for treating ADHD symptoms as determined by randomized controlled clinical trials that typically study patients for several weeks. These medications, which are as efficacious, or more efficacious, than many medications used for non-psychiatric disorders (Leucht et al., 2012), are classified as either stimulants (methylphenidate and amphetamine) or non-stimulants (atomoxetine, extended release guanfacine, and extended release clonidine).

Effects of Medications on Symptoms: Results from Randomized, Controlled Clinical Trials

148. Protocols for using medications for ADHD are well described in detailed guidelines prepared by professional health care associations (Alliance, 2011; Banaschewski T, 2018;

Bolea-Alamanac et al., 2014; Crunelle et al., 2018; Flisher, 2013; Graham et al., 2011; Kooij et al., 2019; National Collaborating Centre for Mental Health, 2018; National Institute for Health Care and Excellence, 2018; Pliszka, 2007; Schoeman and Liebenberg, 2017; Seixas et al., 2012; Taylor et al., 2004; Wolraich et al., 2011).

149. A network meta-analysis found stimulants to be highly effective in reducing the symptoms of ADHD. Compared with placebo, as rated by clinicians, amphetamines were associated with large improvements in all age groups (youths 6 studies with 2179 participants, adults 5 studies with 1521 participants), methylphenidate with large improvements in youths (9 studies, 2677 participants) and moderate ones in adults (11 studies, 2909 participants). Extended release guanfacine (7 studies, 1930 participants) led to moderate improvements in children. Atomoxetine led to moderate improvements in all age groups (youths 21 studies with 3812 participants, adults 11 studies with 3377 participants). Taking side effects into account, the medications with the best benefit-to-risk ratios were methylphenidate for children and adolescents, and amphetamines for adults (Cortese et al., 2018a).
150. A meta-analysis of 18 studies with over 2,000 adults with ADHD found three amphetamine derivatives (dextroamphetamine, lisdexamfetamine, and mixed amphetamine salts) to be associated with moderate reductions in ADHD symptoms (Castells et al., 2011). Another meta-analysis, combining four studies with 216 youths, found mixed amphetamine salts to be slightly more effective at reducing ADHD symptoms than methylphenidate (Faraone et al., 2002).

151. A meta-analysis of 19 parallel group trials with over 1,600 participants, found methylphenidate produced moderate to large improvements in teacher-rated ADHD symptoms, teacher-rated behavior and parent-rated quality of life. There was no evidence of serious adverse events, and just a slightly elevated risk of non-serious side effects (Storebø et al., 2015).
152. A meta-analysis found dexamethylphenidate strongly reduced youth ADHD symptoms relative to placebo (seven studies, almost 1,500 participants), and had three times the clinical response rate (four studies, over 600 participants) (Maneeton et al., 2015). Another meta-analysis, covering six RCTs with 253 participants, reported that methylphenidate strongly reduced adult ADHD symptoms, with higher doses resulting in greater improvement (Faraone et al., 2004).
153. A meta-analysis of seven studies with over 1,600 participants reported that atomoxetine moderately reduced ADHD symptoms. (Cheng et al., 2007).
154. A meta-analysis found that methylphenidate (13 studies, over 2,200 adults) and lisdexamfetamine (five studies, over 2,300 adults) led to small-to-moderate reductions in symptoms of emotional dysregulation; for atomoxetine (three studies, 237 adults) the reductions were small (Lenzi et al., 2018). Another meta-analysis covering nine studies with over 1,300 youths reported atomoxetine to be associated with small reductions in emotional symptoms (Schwartz and Correll, 2014).

155. A meta-analysis reported moderate-to-strong improvements in ADHD symptoms with methylphenidate in ADHD patients with borderline intellectual functioning or intellectual disability (8 studies, 423 children). (Sun et al., 2019a).
156. A meta-analysis of 23 studies with over 2,900 children with ADHD reported that stimulant medications reduced anxiety by 14% relative to placebo (Coughlin et al., 2015).
157. A meta-analysis of nine studies with over 1,300 participants found stimulants to be highly effective in reducing aggression, oppositional behavior, and conduct problems in youth with ADHD (with and without oppositional defiant disorder) and conduct disorder, as measured by teachers, and moderately effective as measured by parents (Pringsheim et al., 2015).

Effects of Medications on Impairments Associated with ADHD: Results from Naturalistic Studies

158. A Swedish registry study of over 650,000 students found that treatment with ADHD medication for three months resulted in a more than nine-point gain in grade point sum (on a scale of 0 to 320); treatment was associated with an increase in the probability of completing upper secondary school by two-thirds (Jangmo et al., 2019).
159. A Swedish national register study of over 61,000 youths with ADHD found that their test scores were higher during periods they were taking medication vs non-medicated periods (Lu et al., 2017). A Danish study of over half a million children (over 6,400 with ADHD) found that discontinuation of ADHD medication was associated with a small but significant decline in grade point averages (Keilow et al., 2018). A meta-analysis of nine RCTs comprising 1,463

patients found that discontinuing medications led to a worsening in quality of life for children and adolescents but not adults. (Tsujii et al., 2020)

160. A Swedish cohort study of over 25,000 ADHD patients found a one-third reduction in criminality among men receiving ADHD medication, and a 40% reduction for women (Lichtenstein et al., 2012). A Danish national registry study of over 4,200 individuals with childhood ADHD found that crime rates in adulthood were 30-40% lower during periods of taking ADHD medication (Mohr-Jensen et al., 2019).
161. A Danish cohort study of over 700,000 people, including 4,557 with ADHD, found that among teenagers with ADHD, stimulant treatment was associated with a decrease in rates of injuries (30% for ten-year olds and 40% for twelve-year olds) (Dalsgaard et al., 2015a).
162. Using the Swedish national registries, a study followed 9,421 youths with ADHD and 2,986 youths with both ADHD and other psychiatric diagnoses from 2006 to 2013. It compared periods when they were taking ADHD medication with periods when they were not. During medicated periods both groups had a greater than 10% reduction in unintended injuries, and a greater than 70% reduction in traumatic brain injuries (Ghirardi et al., 2020).
163. A Taiwanese study of over 124,000 youths with ADHD found that methylphenidate treatment decreased the risk for traumatic brain injuries, after adjusting for confounders (Liao et al., 2018).
164. A nationwide study compared 7,200 Taiwanese youths with ADHD with 36,000 children without ADHD. After adjusting by age, sex, urbanization level, and geographic region, boys with ADHD were almost 40% more likely and girls with ADHD 60% more likely to suffer bone

fractures (Guo et al., 2016). Another study from Taiwan identified over 6,200 youths newly diagnosed with ADHD and assessed the effect of methylphenidate treatment. The risk of bone fractures was 20% lower in those who had over half a year of methylphenidate treatment (Chen et al., 2017b).

165. A population-based, electronic medical records database in Hong Kong identified over 17,000 individuals aged 6-19 years who had been prescribed methylphenidate. Of these, almost 5,000 had at least one trauma-related emergency room admission. Researchers found a 9% reduction in such admissions during periods covered by a methylphenidate prescription compared with periods with no active prescription (Man et al., 2015).

166. A meta-analysis of five studies with over 13,000 participants found that ADHD medications (primarily stimulants) were associated with a greater than 10% reduction in unintentional injuries (Ruiz-Goikoetxea et al., 2018a).

167. Using Swedish national registers, a study of over 17,000 people with ADHD found that medication for ADHD was associated with a greater than 50% reduction in the risk of serious transport accidents among males but not females. Over 40% of crashes by male patients would have been avoided if they had been receiving treatment during the entire period (Chang et al., 2014b). A U.S. national cohort study of 2.3 million people with ADHD examined emergency room visits for motor vehicle crashes over ten years. Males with ADHD had a 38% lower risk of crashes in months when receiving ADHD medication compared with months when not receiving medication, and females a 42% lower risk in months when receiving ADHD medication. About a fifth of crashes would have been

avoided if they had been on medication throughout the period of the study (Chang et al., 2017).

168. A longitudinal study using the Taiwan Health Insurance Research Database compared almost 18,000 adolescent and young adults with ADHD with over 70,000 age- and sex-matched controls. Short-term use of ADHD medications was associated with a 30% reduction in sexually transmitted infections, and long-term use with a 40% reduction, though these reductions were only among males (Chen et al., 2018a).
169. A nationwide longitudinal cohort study using the Swedish national registers found that among more than 38,000 individuals with ADHD, ADHD medication was associated with a greater than 40% reduction in the risk for depression three years later. The risk decreased with the duration of ADHD medication use. Depression was 20% less common when patients received ADHD medication compared with periods when they did not (Chang et al., 2016).
170. A Swedish population-based study of 38,000 people with ADHD found a 20% decline in suicide related events among those prescribed stimulants during periods when they were under treatment as opposed to during periods when they were not under treatment. No such benefit was found for non-stimulant medications (Chen et al., 2014).
171. A Taiwanese study identified 85,000 youths with ADHD using National Health Insurance data to examine whether methylphenidate use affected suicide attempts. After adjusting for relevant variables, it found a 60% lower risk of suicide in those using methylphenidate

for 3 months to half a year, and a 70% reduction among those using methylphenidate for more than half a year (Liang et al., 2018b).

172. A study using the Swedish national registers investigated the association between prescription stimulant medication for ADHD in 2006 and substance abuse during 2009 among all 38,753 people born between 1960 and 1998 and diagnosed with ADHD. After controlling for relevant variables, it found a greater than 30% reduction in indicators of substance abuse among those prescribed stimulants. The longer the duration of medication, the lower the rate of substance abuse (Chang et al., 2014c). A meta-analysis of 14 studies with over 2,300 participants found that people with ADHD were about half as likely to smoke cigarettes when regularly treated with stimulant medications (Schoenfelder et al., 2014). A meta-analysis found that stimulants did not increase the risk for alcohol (11 studies, over 1,300 participants, nicotine (6 studies, 884 participants), cocaine (7 studies, 950 participants), or cannabis abuse or dependence (9 studies, over 1,100 participants) (Humphreys et al., 2013).
173. A nationwide study of over 7,500 Taiwanese adolescents with ADHD and over 30,000 matched controls found that long-term use of ADHD medication use was associated with a 30% decrease in teenage pregnancy (Hua et al., 2020).
174. A nationwide population-based cohort using Taiwan's National Health Insurance Research Database identified over 68,000 children and adolescents with a diagnosis of ADHD and who were prescribed methylphenidate and compared them with an identical number of controls matched on age, gender and year of first ADHD diagnosis. After

controlling for potential confounders, ADHD individuals prescribed methylphenidate had a one-fifth lower rate of all-cause mortality than ADHD individuals not prescribed methylphenidate. Delayed use of methylphenidate, on the other hand, was associated with slightly higher (5%) mortality. Long-term methylphenidate use was associated with a one-sixth lower rate of all-cause mortality. The authors caution, however, that "information lacking in the database precluded the measurement of other possible confounders, such as family history, psychosocial stressors, effect of behavioural therapy or severity of comorbidities," and thus unmeasured confounding cannot be excluded (Chen et al., 2020a).

175. A nationwide population-based cohort using Taiwan's National Health Insurance Research Database identified over 90,000 individuals younger than 18 years with a diagnosis of ADHD, and compared risk of burn injury between those not on methylphenidate, those on methylphenidate for less than 90 days, and those on methylphenidate for more than 90 days. The data suggested that fully half the incidence of burn injuries could have been prevented by taking methylphenidate. Compared with patients not taking methylphenidate, those taking it for less than 90 days had a 30% lesser risk of burn injuries, and those taking it for 90 days or more a 57% reduction in risk, after adjusting for confounders (Chen et al., 2020b).

Effects of Medications for ADHD on the Brain

176. A meta-analysis of methylphenidate treatment for ADHD found moderate improvements in response inhibition (25 studies, 787 participants) and sustained attention

(29 studies, 956 participants), but no significant effect on working memory (13 studies, 559 participants) (Tamminga et al., 2016).

177. A meta-analysis of 14 fMRI studies with 212 participants reported that medication treatment for ADHD made the brains of youth with ADHD function in a way that was more like the brains of people without ADHD in brain areas involved in the control of cognition, which is typically disrupted in ADHD (Rubia et al., 2014). Medication treatment for ADHD had no effect on brain structure in studies of 4,180 ADHD patients in the ENIGMA-ADHD Working Group set of 36 cohorts from around the world (Hoogman et al., 2017; Hoogman et al., 2019).

Adverse Effects of ADHD Medications

178. A meta-analysis found that stimulants moderately reduced total sleep time (7 studies, 223 children), delayed the onset of sleep (7 studies, 171 children), and slightly-to-moderately decreased sleep efficiency (7 studies, 155 children) (Kidwell et al., 2015). A meta-analysis found that children and adolescents on methylphenidate were 50% more likely to report abdominal pain (46 studies, over 4,600 youths) and over three times more likely to experience decreases in appetite (52 studies, over 4,800 youths) and weight (7 studies, over 850 youths) (Holmskov et al., 2017). An umbrella review of network meta-analyses and meta-analyses of RCTs and cohort studies examined 78 adverse events across 19 categories of 80 psychotropic medications in children and adolescents with mental disorders including data from nine network meta-analyses, 39 meta-analyses, 90 individual RCTs, and eight cohort studies with a total of 337,686 children and adolescents included

(Solmi et al., 2020). Five medications for ADHD were associated with significantly worse anorexia (atomoxetine, d-amphetamine, lisdexamphetamine, methylphenidate, modafinil), four with insomnia (d-amphetamine, lisdexamphetamine, methylphenidate, modafinil), three with weight loss (atomoxetine, methylphenidate, modafinil), two each with abdominal pain (methylphenidate, guanfacine), discontinuation due to adverse event (lisdexamphetamine, guanfacine), hypertension (atomoxetine, lisdexamphetamine), and sedation (clonidine, guanfacine), and one with QT prolongation (guanfacine)..

179. A meta-analysis of twelve studies with over 3,300 adults found that those taking atomoxetine were about 40% more likely to discontinue treatment due to adverse events than those on placebo (Cunill et al., 2013). A meta-analysis found that methylphenidate was more than twice as likely to induce insomnia as atomoxetine (10 studies, over 3,000 youths), but about half as likely to cause nausea (8 studies, over 2,750 youths) and vomiting (9 studies, over 2,500 youths), and about a sixth as likely to cause drowsiness (9 studies, over 2,800 youths) (Liu et al., 2017a). A meta-analysis of methylphenidate treatment studies reported a 55% increase in adverse events relative to placebo, none life-threatening (11 studies, over 2,100 youths), but a fivefold increase in anorexia (3 studies, 613 youths), and more than fourfold increase in insomnia (4 studies, 749 youths) (Ching et al., 2019).
180. Children treated with stimulants may show delays in expected height gains averaging two centimeters over one or two years. These sometimes attenuate over time and often reverse when treatment is stopped (Faraone et al., 2008). A medical records study from the USA comparing 32,999 stimulant-treated ADHD children with 11,515 controls found continuing declines in expected height over a four-year period. A study from Germany,

however, specifically addressed whether stimulants predicted patients being very short (i.e., being less than or equal to the third percentile of the population). After comparing 3,806 boys not treated with methylphenidate with 118 treated boys, the results did not indicate that methylphenidate increased the probability of this adverse outcome (McCarthy et al., 2018).

181. A study using Danish national registers followed over 700,000 individuals for an average period of almost a decade. Looking at 8,300 people with ADHD, stimulant users had more than twice the rate of cardiovascular events (primarily hypertension) than nonusers. These events were rare (Dalsgaard et al., 2014).
182. A meta-analysis of five studies with over 43,000 children and adolescents found no significant difference in adverse cardiac events between methylphenidate and atomoxetine, and a meta-analysis of three studies with 775 adults found no significant difference in adverse cardiac events between methylphenidate and placebo (Liang et al., 2018a).
183. A meta-analysis covering people of all ages reported methylphenidate was not associated with a higher risk of all-cause death (3 studies, over 1.4 million people), heart attack or stroke (3 studies, over half a million people) (Liu et al., 2019a).
184. A cohort study of over 1.8 million pregnancies in the United States and over 2.5 million pregnancies in the health registries of Denmark, Finland, Sweden, Norway, and Iceland reported that use of methylphenidate (but not amphetamines) by pregnant woman was associated with a higher risk for cardiac malformations from 12.9 per thousand infants to 16.5 per thousand infants (Huybrechts et al., 2018). A meta-analysis of four studies of three

million women also found that intrauterine exposure to methylphenidate was associated with a higher risk of cardiac malformations (Koren et al., 2020).

185. A meta-analysis examining the safety of atomoxetine found no significant increase in risk of irritability (3 studies, over 1,100 children) (Pozzi et al., 2018). Two others, one combining twenty studies with over 3,000 participants, and another combining 37 studies with over 3,800 participants, found no increase in risk of all-cause treatment discontinuation in youths (Catala-Lopez et al., 2017; Schwartz and Correll, 2014). However, a meta-analysis of twelve studies with over 3,300 adults found 40% greater rate of all-cause treatment discontinuation, leading to a conclusion that “atomoxetine has a poor benefit–risk balance for the treatment of adults with ADHD” (Cunill et al., 2013).
186. The Hong Kong Clinical Data Analysis & Reporting System, a population-based, electronic medical records database, was used to examine over 25,000 people receiving methylphenidate for ADHD. During the 90-day period prior to initiation of treatment, individuals with ADHD were greater than six times more likely to attempt suicide than after treatment. After ongoing treatment, the risk for attempted suicide was no longer elevated among patients with ADHD (Man et al., 2017).
187. Using the same Hong Kong database, the risk for psychosis did not differ between periods when patients were on and off methylphenidate treatment (Man et al., 2016).
188. A Swedish registry study of over 23,000 adolescents and young adults treated with methylphenidate for ADHD found no evidence for an association between psychosis and methylphenidate treatment. A year after initiation of methylphenidate treatment, the

incidence of psychotic events was 36% lower in those with a history of psychosis and 18% lower in those without a history of psychosis relative to the period immediately before the beginning of treatment (Hollis et al., 2019).

Misuse and Diversion of Stimulant Medications

189. A systematic review of 109 studies concluded that the non-medical use of prescribed stimulants is a significant public health problem, especially in college students. Most non-medical use is associated with zero or minor medical effects, but adverse medical outcomes, including death, occur in some individuals, particularly when administered by non-oral routes. Academic and occupational performance enhancement were the most commonly cited motivations for non-medical use of stimulants, but there is little evidence that academic performance is improved by non-medical use in individuals without ADHD (Faraone et al., 2020).
190. The non-medical use of prescribed stimulants in individuals without ADHD is associated with lower educational attainment. A U.S. prospective study followed a nationally representative sample of over 8,300 high school seniors from age 18 to age 35. Those who used prescription stimulants non-medically were 17% less likely to earn a bachelor's degree than those who had neither medical or non-medical use (McCabe et al., 2017).
191. A retrospective study compared 4.4 million people dispensed ADHD medications with 6.1 million people dispensed asthma medications. Obtaining prescriptions from multiple prescribers or filling prescriptions at multiple pharmacies was highly correlated with abuse, misuse, and diversion. These "shopping" behaviors were four times more frequent in the

ADHD group than in the asthma group. Those dispensed stimulant medications were more than eight times as likely to engage in shopping behavior than those dispensed nonstimulants, but only one in 250 people with stimulant prescriptions engaged in shopping behavior (Cepeda et al., 2014).

192. A U.S. study of over 440,000 respondents found that use of illegal drugs or other non-medical use of prescription drugs preceded non-medical use of ADHD medication in more than three out of four cases (Sweeney et al., 2013).

193. A study examined Swedish national pharmacy dispensing data for all 56,922 individuals who filled a methylphenidate prescription between 2010 and 2011. 4,304 of the methylphenidate users (7.6%) overused medication as measured by dispensed prescriptions. Overuse was 17 times more frequent for ages 46-65 compared with ages 6-12 year. It was also twice as frequent among those with previous alcohol and drug misuse (Bjerkeli et al., 2018).

194. Large studies of calls to U.S. poison control centers related to ADHD medications find that intentional exposures, including suspected suicide and medication abuse and/or misuse is associated with admission to critical care units and, rarely, death especially when snorted or injected (Faraone et al., 2019a; King et al., 2018).

Which Non-Medication Treatments are Safe and Effective for ADHD?

Many non-medical treatments have been proposed for ADHD. Most of those offered on the Internet have not been tested or have been shown not to be effective. In this section, we distinguish between the effects of a treatment for ADHD symptoms and other benefits it may

confer. Due to the way these therapies are implemented and recorded in the medical record, large scale naturalistic studies of longer-term outcomes are not possible.

Behavioral and Cognitive-Behavioral Therapies

Behavioral treatments for ADHD are diverse in nature and have a different content and focus depending on the age of the patient. For preschoolers and grade-school children, parents are trained to improve their method of disciplining and interacting with their children. For adolescents and adults, therapy helps patients improve their organizational skills. For some patients, teachers contribute to a program aimed at improving the child's behavior. Some of these therapies focus on improving social behaviors and developing practical skills. In this section, however, we focus only on the ability of such treatments to improve ADHD symptoms. Readers should keep in mind that the failure of a treatment to substantially improve ADHD symptoms does not mean it is not useful for other purposes.

195. A meta-analysis found parent training for preschool children with ADHD to be associated with a moderate reduction in parent-reported ADHD symptoms (15 studies, few with active controls, over a thousand participants) and conduct problems (14 studies, few with active controls, over a thousand participants), but no significant results for independently assessed ADHD symptoms (6 studies, 403 participants) and conduct problems (6 studies, 311 participants). Independent assessments reported a small reduction in negative parenting (10 studies, 771 participants) (Rimestad et al., 2019).
196. A meta-analysis of 19 studies of cognitive behavior therapy (CBT) for adults with ADHD included 896 participants. It found associations with moderate improvements in self-

reported ADHD symptoms and self-reported functioning. But when limited to the two studies with active controls and blind assessors (N = 244 participants), it found only small improvements (Knouse et al., 2017). In another meta-analysis of 160 patients with adult ADHD, CBT led to large to moderate improvements compared with waiting list controls. In three studies of 191 patients CBT led to small to moderate improvements compared with active controls (Young et al., 2020).

197. A meta-analysis of 32 studies with over two thousand participants found that cognitive training resulted in small to moderate improvements in executive functioning in preschoolers with ADHD (Scionti et al., 2019).

198. A meta-analysis explored the effectiveness of meditation-based therapy. It found moderate reductions in ADHD symptoms in both children and adolescents (6 RCTs, 240 participants) and adults (6 RCTs, 339 participants), but half the studies did not use active controls. Removing studies with waiting list controls made results nonsignificant. The authors concluded “there is insufficient methodologically sound evidence to support the recommendation of meditation-based therapies as an intervention aimed to target ADHD core symptoms or related neuropsychological dysfunctions in children/adolescents or adults with ADHD” (Zhang et al., 2018).

199. A meta-analysis found that social skills training for youth with ADHD did not improve teacher-assessed social skills (11 studies, over 1,200 youths), general behavior (8 studies,

over 1,000 youths), or school performance and grades (5 studies, over 600 youths) (Storebo et al., 2019).

200. A meta-analysis of ten studies with 893 youths reported that organizational skills interventions led to moderate reductions in parent-reported inattention symptoms (Bikic et al., 2017).

Computer-based Cognitive Training and Neurofeedback

201. A meta-analysis of five randomized controlled trials (RCTs) with 263 participants exploring the efficacy of neurofeedback found a small reduction in inattention, but no significant reduction in hyperactivity-impulsivity or overall ADHD symptoms with ratings by probably blinded evaluators (researchers measuring outcomes did not know if patients were receiving the active or control treatment) (Micoulaud-Franchi et al., 2014). A more recent meta-analysis of ten RCTs with 256 participants found no effect on inattention symptoms, but a small-to-medium reduction in hyperactivity-impulsivity symptoms (Van Doren et al., 2019).
202. The European ADHD Guidelines Group published meta-analyses of cognitive training and neurofeedback for youth. Probably blinded cognitive training studies with active controls (6 studies, 287 youths) reported no significant reduction in ADHD symptoms. But they did find moderate improvements in verbal working memory (5 studies, 263 youths). There were no significant effects on academic outcomes in math and reading (95 studies, 290 youths) (Cortese et al., 2015). Blinded neurofeedback studies with active/sham controls (6 studies, 251 participants) found no significant reduction in ADHD symptoms (Cortese et al., 2016a).

203. A meta-analysis found that working memory training led to short-term improvements in both verbal working memory (21 studies, over 1,300 participants) and visuospatial working memory (18 studies, over 1,000 participants), with “no convincing evidence that even such near-transfer effects are durable.” Moreover, most of the studies lacked active controls (Melby-Lervag and Hulme, 2013).

Supplements, Diet, and Exercise

204. Omega-3 fatty acid supplementation was associated with small-to-medium improvements in ADHD symptoms in three meta-analyses (ten studies with 699 participants, 16 studies with 1,408 participants, 7 studies with 534 participants) (Bloch and Qawasmi, 2011; Chang et al., 2018; Hawkey and Nigg, 2014). Another meta-analysis, with 18 studies and 1,640 participants, found tiny improvements (Puri and Martins, 2014).
205. A meta-analysis found no evidence of any effect of omega-3 fatty acid supplements on parent-rated (5 studies, 650 children) or teacher-rated (3 studies, 598 children) emotional lability symptoms, or parent-rated (8 studies, 875 children) or teacher-rated (6 studies, 805 children) oppositional symptoms in children with ADHD (Cooper et al., 2016).
206. A meta-analysis of five double-blind crossover studies with 164 participants found that restricting synthetic food colors from children’s diets was associated with a small reduction in ADHD symptoms (Nigg et al., 2012).
207. A meta-analysis of ten studies (300 children) found exercise was associated with a moderate reduction in ADHD symptoms, but had no significant effect after adjusting for publication bias (Vysniauske et al., 2020). Another meta-analysis found no significant effect

of exercise on either hyperactivity/impulsivity (4 studies, 227 participants) or inattention symptoms (6 studies, 277 participants), but significant reductions in anxiety and depression (5 studies, 164 participants) (Zang, 2019).

208. A nationwide population study using the Swedish Twin Register identified almost 18,000 twins who completed a web-based examining the relationship between inattention and hyperactivity/impulsivity subtypes and dietary habits. The two subtypes of ADHD exhibited very similar associations. Both had significant associations with unhealthy diets. Both were more likely to be eating foods high in added sugar and neglecting fruits and vegetables while eating more meat and fats. After adjusting for degree of relatedness of twins (whether monozygotic or dizygotic) and controlling for the other ADHD subtype, the associations remained statistically significant for inattention, but diminished to negligible levels or became statistically nonsignificant for hyperactivity/impulsivity. Even for persons with inattention symptoms, adjusted correlations were small (never exceeding $r = 0.10$), with the strongest associations being for overall unhealthy eating habits and eating foods high in added sugar. Among over 700 pairs of monozygotic (“identical”) twins, it found small but robust associations between inattention symptoms and unhealthy eating habits, and especially with consumption of foods high in added sugar. For hyperactivity/impulsivity symptoms, the association with unhealthy eating habits was weaker, and the association with consumption of foods high in added sugar became statistically insignificant (Li et al., 2020).

Discussion

This work has curated evidence-based statements about ADHD which paint a picture of the disorder that we summarize as follows:

ADHD is a chronic disorder in which developmentally inappropriate symptoms of inattention and/or hyperactivity/impulsivity lead to impairments in many aspects of living. The disorder, which starts in childhood or early adolescence and is more common in boys than girls, affects 5.9% of youth and 2.8% of adults worldwide. There are multiple genetic and environmental risk factors that accumulate in various combinations to cause ADHD. These risk factors lead to subtle changes in multiple brain networks and in the cognitive, motivational, and emotional processes they control. People diagnosed with ADHD have an elevated risk for school failure, antisocial behavior, other psychiatric problems, somatic disorders, drug and alcohol abuse, accidental injuries, and premature death, including attempted and completed suicide. As a result, ADHD costs society hundreds of billions of dollars each year. Several medications are safe and effective for treating ADHD and for preventing many adverse outcomes. Non-medication treatments are available but, compared with medications, are less effective for reducing inattention, hyperactivity, and impulsivity.

Despite this large body of evidence, we have much more to learn about the disorder and its various manifestations. Epidemiologic studies have taught us that ADHD occurs around the world, but we know little about how culture affects the expression of ADHD symptoms or the response to treatment. Because most research about ADHD is based on Caucasian and East

Asian samples, we must be cautious in generalizing our assertions to other groups. In addition, far more research pertains to males than females. We also need to learn more about ADHD in older adults. Future research into ADHD should examine more diverse samples from a wider range of cultural contexts.

We have learned much about the causes of ADHD but are only beginning to understand how genes and environment combine to cause the disorder and affect the brain to produce symptoms and impairments. Some of these causes may be shared with ADHD's somatic comorbidities. Examples include oxidative stress, inflammation, and insulin resistance. Future work should focus on biological and psychological causal mechanisms to find points of intervention that will improve the effectiveness of medical and non-medical treatments and, eventually, prevent onset of the disorder. Although the medications that treat ADHD are highly effective, we need better methods to prevent the misuse and diversion of these medications, especially among adolescents and young adults (Faraone et al., 2020).

Many decades of research have led to a method of diagnosing ADHD that is highly valid as a predictor of treatment response, family history of ADHD, many clinical features, measures of brain structure and function, and adverse outcomes. Nevertheless, there are several new directions for diagnosis. One is to better understand the nature and causes of emotional symptoms in ADHD and whether these should be incorporated into diagnostic criteria (Faraone et al., 2019b). Another is to determine if and how mild or sub-threshold cases of ADHD should be diagnosed and treated (Kirova et al., 2019). Different trajectories of ADHD across the life-cycle need to be further investigated.

Many researchers are trying to develop computerized or biological tests using information about the patient's behavior, brain and/or genetic makeup. The hope is that such tests will one day diagnose the disorder, predict a personalized approach to treatment or assist clinicians in these areas. Others are working on methods that use the vast data available from medical records to predict which patients with ADHD are at greatest risk for adverse outcomes later in life. Such work may someday allow healthcare systems to allocate resources to the highest risk patients.

Although we have good treatments for ADHD, even the best treatments are only partially effective. The future of treatment for ADHD will include new medications currently in development and a stronger evidence base for novel non-medication treatments for treating ADHD symptoms or associated impairments, such as trigeminal nerve stimulation (McGough et al., 2019) and game-based treatments (Craven and Groom, 2015; Dosis et al., 2015). And more data are needed to improve existing non-medication treatments and to test the efficacy of traditional therapies such as acupuncture, yoga, and Ayurvedic therapies. Also, little is known about how the somatic disorders that co-occur with ADHD interact with treatments for ADHD and how the symptoms of the disorder affect somatic outcomes. We need to learn more about how duration of treatment affects outcomes over longer periods of time.

We also know little about stigma and ADHD. Stigmatizing attitudes toward ADHD are common and may play a role in socially and clinically important outcomes. These negative attitudes affect patients at all stages of their life. Such attitudes have been documented among

individuals at all ages and in all groups, including family, peers, teachers, clinicians, and even individuals with ADHD themselves (Lebowitz, 2016).

Despite these and other gaps in our knowledge about ADHD, nearly two and a half centuries after the first textbook description of an ADHD-like syndrome, the statements about ADHD which we have curated, make us confident that the contemporary diagnosis of the disorder is a valid and useful category that can be used around the world to improve the lives of the many people who suffer from the disorder and its complications.

References

- Adeyemo, B.O., Biederman, J., Zafonte, R., Kagan, E., Spencer, T.J., Uchida, M., Kenworthy, T., Spencer, A.E., Faraone, S.V., 2014. Mild Traumatic Brain Injury and ADHD: A Systematic Review of the Literature and Meta-Analysis. *J Atten Disord* 18, 576-584.
- Akmatov, M.K., Ermakova, T., Batzing, J., 2019. Psychiatric and Nonpsychiatric Comorbidities Among Children With ADHD: An Exploratory Analysis of Nationwide Claims Data in Germany. *J Atten Disord*, 1087054719865779.
- Alliance, C.A.R., 2011. Canadian ADHD Practice Guidelines, 3rd ed., 3rd ed.
- American Psychiatric Association, 2013. Diagnostic and statistical manual of mental disorders (5th ed.), 5th ed. American Psychiatric Publishing, Arlington, VA.
- Andersen, C.H., Thomsen, P.H., Nohr, E.A., Lemcke, S., 2018. Maternal body mass index before pregnancy as a risk factor for ADHD and autism in children. *Eur Child Adolesc Psychiatry* 27, 139-148.
- Anttila, V., Bulik-Sullivan, B., Finucane, H.K., Walters, R.K., Bras, J., Duncan, L., Escott-Price, V., Falcone, G.J., Gormley, P., Malik, R., Patsopoulos, N.A., Ripke, S., Wei, Z., Yu, D., Lee, P.H., Turley, P., Grenier-Boley, B., Chouraki, V., Kamatani, Y., Berr, C., Letenneur, L., Hannequin, D., Amouyel, P., Boland, A., Deleuze, J.-F., Duron, E., Vardarajan, B.N., Reitz, C., Goate, A.M., Huentelman, M.J., Kamboh, M.I., Larson, E.B., Rogaeva, E., St George-Hyslop, P., Hakonarson, H., Kukull, W.A., Farrer, L.A., Barnes, L.L., Beach, T.G., Demirci, F.Y., Head, E., Hulette, C.M., Jicha, G.A., Kauwe, J.S.K., Kaye, J.A., Leverenz, J.B., Levey, A.I., Lieberman, A.P., Pankratz, V.S., Poon, W.W., Quinn, J.F., Saykin, A.J., Schneider, L.S., Smith, A.G., Sonnen, J.A., Stern, R.A., Van Deerlin, V.M., Van Eldik, L.J., Harold, D., Russo, G., Rubinsztein, D.C., Bayer, A., Tsolaki, M., Proitsi, P., Fox, N.C., Hampel, H., Owen, M.J., Mead, S., Passmore, P., Morgan, K., Nöthen, M.M., Schott, J.M., Rossor, M., Lupton, M.K., Hoffmann, P., Kornhuber, J., Lawlor, B., McQuillin, A., Al-Chalabi, A., Bis, J.C., Ruiz, A., Boada, M., Seshadri, S., Beiser, A., Rice, K., van der Lee, S.J., De Jager, P.L., Geschwind, D.H., Riemenschneider, M., Riedel-Heller, S., Rotter, J.I., Ransmayr, G., Hyman, B.T., Cruchaga, C., Alegret, M., Winsvold, B., Palta, P., Farh, K.-H., Cuenca-Leon, E., Furlotte, N., Kurth, T.,

Ligthart, L., Terwindt, G.M., Freilinger, T., Ran, C., Gordon, S.D., Borck, G., Adams, H.H.H., Lehtimäki, T., Wedenoja, J., Buring, J.E., Schürks, M., Hrafnisdóttir, M., Hottenga, J.-J., Penninx, B., Artto, V., Kaunisto, M., Vepsäläinen, S., Martin, N.G., Montgomery, G.W., Kurki, M.I., Hämäläinen, E., Huang, H., Huang, J., Sandor, C., Webber, C., Muller-Myhsok, B., Schreiber, S., Salomaa, V., Loehrer, E., Göbel, H., Macaya, A., Pozo-Rosich, P., Hansen, T., Werge, T., Kaprio, J., Metspalu, A., Kubisch, C., Ferrari, M.D., Belin, A.C., van den Maagdenberg, A.M.J.M., Zwart, J.-A., Boomsma, D., Eriksson, N., Olesen, J., Chasman, D.I., Nyholt, D.R., Anney, R., Avbersek, A., Baum, L., Berkovic, S., Bradfield, J., Buono, R., Catarino, C.B., Cossette, P., De Jonghe, P., Depondt, C., Dlugos, D., Ferraro, T.N., French, J., Hjalgrim, H., Jamnadas-Khoda, J., Kälviäinen, R., Kunz, W.S., Lerche, H., Leu, C., Lindhout, D., Lo, W., Lowenstein, D., McCormack, M., Møller, R.S., Molloy, A., Ng, P.-W., Oliver, K., Privitera, M., Radtke, R., Ruppert, A.-K., Sander, T., Schachter, S., Schankin, C., Scheffer, I., Schoch, S., Sisodiya, S.M., Smith, P., Sperling, M., Striano, P., Surges, R., Thomas, G.N., Visscher, F., Whelan, C.D., Zara, F., Heinzen, E.L., Marson, A., Becker, F., Stroink, H., Zimprich, F., Gasser, T., Gibbs, R., Heutink, P., Martinez, M., Morris, H.R., Sharma, M., Ryten, M., Mok, K.Y., Pulit, S., Bevan, S., Holliday, E., Attia, J., Battey, T., Boncoraglio, G., Thijs, V., Chen, W.-M., Mitchell, B., Rothwell, P., Sharma, P., Sudlow, C., Vicente, A., Markus, H., Kourkoulis, C., Pera, J., Raffeld, M., Silliman, S., Boraska Perica, V., Thornton, L.M., Huckins, L.M., William Rayner, N., Lewis, C.M., Gratacos, M., Rybakowski, F., Keski-Rahkonen, A., Raevuori, A., Hudson, J.I., Reichborn-Kjennerud, T., Monteleone, P., Karwautz, A., Mannik, K., Baker, J.H., O'Toole, J.K., Trace, S.E., Davis, O.S.P., Helder, S.G., Ehrlich, S., Herpertz-Dahlmann, B., Danner, U.N., van Elburg, A.A., Clementi, M., Forzan, M., Docampo, E., Lissowska, J., Hauser, J., Tortorella, A., Maj, M., Gonidakis, F., Tziouvas, K., Papezova, H., Yilmaz, Z., Wagner, G., Cohen-Woods, S., Herms, S., Julià, A., Rabionet, R., Dick, D.M., Ripatti, S., Andreassen, O.A., Espeseth, T., Lundervold, A.J., Steen, V.M., Pinto, D., Scherer, S.W., Aschauer, H., Schosser, A., Alfredsson, L., Padyukov, L., Halmi, K.A., Mitchell, J., Strober, M., Bergen, A.W., Kaye, W., Szatkiewicz, J.P., Cormand, B., Ramos-Quiroga, J.A., Sánchez-Mora, C., Ribasés, M., Casas, M., Hervas, A., Arranz, M.J., Haavik, J., Zayats, T., Johansson, S., Williams, N., Elia, J., Dempfle, A., Rothenberger, A., Kuntsi, J., Oades, R.D., Banaschewski, T., Franke, B., Buitelaar, J.K., Arias Vasquez, A., Doyle, A.E., Reif, A., Lesch, K.-P., Freitag, C., Rivero, O., Palmason, H., Romanos, M., Langley, K., Rietschel, M., Witt, S.H., Dalsgaard, S., Børghlum, A.D., Waldman, I., Wilmot, B., Molly, N., Bau, C.H.D., Crosbie, J., Schachar, R., Loo, S.K., McGough, J.J., Grevet, E.H., Medland, S.E., Robinson, E., Weiss, L.A., Bacchelli, E., Bailey, A., Bal, V., Battaglia, A., Betancur, C., Bolton, P., Cantor, R., Celestino-Soper, P., Dawson, G., De Rubeis, S., Duque, F., Green, A., Klauck, S.M., Leboyer, M., Levitt, P., Maestrini, E., Mane, S., De-Luca, D.M.-, Parr, J., Regan, R., Reichenberg, A., Sandin, S., Vorstman, J., Wassink, T., Wijsman, E., Cook, E., Santangelo, S., Delorme, R., Rogé, B., Magalhaes, T., Arking, D., Schulze, T.G., Thompson, R.C., Strohmaier, J., Matthews, K., Melle, I., Morris, D., Blackwood, D., McIntosh, A., Bergen, S.E., Schalling, M., Jamain, S., Maaser, A., Fischer, S.B., Reinbold, C.S., Fullerton, J.M., Grigoriou-Serbanescu, M., Guzman-Parra, J., Mayoral, F., Schofield, P.R., Cichon, S., Mühleisen, T.W., Degenhardt, F., Schumacher, J., Bauer, M., Mitchell, P.B., Gershon, E.S., Rice, J., Potash, J.B., Zandi, P.P., Craddock, N., Ferrier, I.N., Alda, M., Rouleau, G.A., Turecki, G., Ophoff, R., Pato, C., Anjorin, A., Stahl, E., Leber, M., Czerski, P.M., Edenberg, H.J., Cruceanu, C., Jones, I.R., Posthuma, D., Andlauer, T.F.M., Forstner, A.J., Streit, F., Baune, B.T., Air, T., Sinnamon, G., Wray, N.R., MacIntyre, D.J., Porteous, D., Homuth, G., Rivera, M., Grove, J., Middeldorp, C.M., Hickie, I., Pergadia, M., Mehta, D., Smit, J.H., Jansen, R., de Geus, E., Dunn, E., Li, Q.S., Nauck, M., Schoevers, R.A., Beekman, A.T., Knowles, J.A., Viktorin, A., Arnold, P., Barr, C.L., Bedoya-Berrio, G., Bienvenu, O.J., Brentani, H., Burton, C., Camarena, B., Cappi, C., Cath, D., Cavallini, M., Cusi, D., Darrow, S., Denys, D., Derks, E.M., Dietrich, A., Fernandez, T., Figee, M., Freimer, N., Gerber, G., Grados, M., Greenberg, E., Hanna, G.L., Hartmann, A., Hirschtritt, M.E., Hoekstra, P.J., Huang, A., Huyser, C., Illmann, C., Jenike, M., Kuperman, S., Leventhal, B., Lochner, C., Lyon, G.J., Macciardi, F., Madruga-Garrido, M., Malaty, I.A., Maras, A., McGrath, L., Miguel, E.C., Mir, P., Nestadt, G., Nicolini, H., Okun, M.S., Pakstis, A., Paschou, P., Piacentini, J., Pittenger, C., Plessen, K., Ramensky, V., Ramos, E.M., Reus, V., Richter, M.A., Riddle, M.A., Robertson,

M.M., Roessner, V., Rosário, M., Samuels, J.F., Sandor, P., Stein, D.J., Tsetsos, F., Van Nieuwerburgh, F., Weatherall, S., Wendland, J.R., Wolanczyk, T., Worbe, Y., Zai, G., Goes, F.S., McLaughlin, N., Nestadt, P.S., Grabe, H.-J., Depienne, C., Konkashbaev, A., Lanzagorta, N., Valencia-Duarte, A., Bramer, E., Buccola, N., Cahn, W., Cairns, M., Chong, S.A., Cohen, D., Crespo-Facorro, B., Crowley, J., Davidson, M., DeLisi, L., Dinan, T., Donohoe, G., Drapeau, E., Duan, J., Haan, L., Hougaard, D., Karachanak-Yankova, S., Khrunin, A., Klovins, J., Kučinskas, V., Lee Chee Keong, J., Limborska, S., Loughland, C., Lönnqvist, J., Maher, B., Mattheisen, M., McDonald, C., Murphy, K.C., Murray, R., Nenadic, I., van Os, J., Pantelis, C., Pato, M., Petryshen, T., Quested, D., Roussos, P., Sanders, A.R., Schall, U., Schwab, S.G., Sim, K., So, H.-C., Stögmann, E., Subramaniam, M., Toncheva, D., Waddington, J., Walters, J., Weiser, M., Cheng, W., Cloninger, R., Curtis, D., Gejman, P.V., Henskens, F., Mattingsdal, M., Oh, S.-Y., Scott, R., Webb, B., Breen, G., Churchhouse, C., Bulik, C.M., Daly, M., Dichgans, M., Faraone, S.V., Guerreiro, R., Holmans, P., Kendler, K.S., Koeleman, B., Mathews, C.A., Price, A., Scharf, J., Sklar, P., Williams, J., Wood, N.W., Cotsapas, C., Palotie, A., Smoller, J.W., Sullivan, P., Rosand, J., Corvin, A., Neale, B.M., 2018. Analysis of shared heritability in common disorders of the brain. *Science* 360, eaap8757.

Arns, M., Conners, C.K., Kraemer, H.C., 2013. A decade of EEG Theta/Beta Ratio Research in ADHD: a meta-analysis. *J Atten Disord* 17, 374-383.

Arruda, M.A., Arruda, R., Guidetti, V., Bigal, M.E., 2020. ADHD Is Comorbid to Migraine in Childhood: A Population-Based Study. *J Atten Disord* 24, 990-1001.

Australian ADHD Professionals Association, 2019. The social and economic costs of ADHD in Australia. Deloitte Access Economics.

Banaschewski T, B.M., Bea M, Döpfner M, Gelb M, Grosse KP, Hohmann S, Huss M, Millenet M, Philipsen A, Retz W, Rösler M, Skrodzki K, Spitzcok von Brisinski I, Stollhoff K, Wilken B, 2018. Leitlinien-Detailansicht ADHS bei Kindern, Jugendlichen und Erwachsenen. AWMD online

Barkley, R.A., 2002. International consensus statement on ADHD. January 2002. *Clin Child Fam Psychol Rev* 5, 89-111.

Beaudry, G., Yu, R., Langstrom, N., Fazel, F.S., 2020. Mental Disorders Among Adolescents in Juvenile Detention and Correctional Facilities: An Updated Systematic Review and Metaregression Analysis. *J Am Acad Child Adolesc Psychiatry* [Epub ahead or print] S0890-8567(20)30061-7.

Beheshti, A., Chavanon, M.L., Christiansen, H., 2020. Emotion dysregulation in adults with attention deficit hyperactivity disorder: a meta-analysis. *BMC Psychiatry* 20, 120.

Benedict, F.T., Vivier, P.M., Gjelsvik, A., 2015. Mental health and bullying in the United States among children aged 6 to 17 years. *J Interpers Violence* 30, 782-795.

Bernardi, S., Faraone, S.V., Cortese, S., Kerridge, B.T., Pallanti, S., Wang, S., Blanco, C., 2012. The lifetime impact of attention deficit hyperactivity disorder: results from the National Epidemiologic Survey on Alcohol and Related Conditions (NESARC). *Psychol Med* 42, 875-887.

Bertelsen, E.N., Larsen, J.T., Petersen, L., Christensen, J., Dalsgaard, S., 2016. Childhood Epilepsy, Febrile Seizures, and Subsequent Risk of ADHD. *Pediatrics* 138, e20154654.

Bikic, A., Reichow, B., McCauley, S.A., Ibrahim, K., Sukhodolsky, D.G., 2017. Meta-analysis of organizational skills interventions for children and adolescents with Attention-Deficit/Hyperactivity Disorder. *Clin Psychol Rev* 52, 108-123.

Bjerkeli, P.J., Vicente, R.P., Mulinari, S., Johnell, K., Merlo, J., 2018. Overuse of methylphenidate: an analysis of Swedish pharmacy dispensing data. *Clin Epidemiol* 10, 1657-1665.

Bjorkenstam, E., Bjorkenstam, C., Jablonska, B., Kosidou, K., 2018. Cumulative exposure to childhood adversity, and treated attention deficit/hyperactivity disorder: a cohort study of 543 650 adolescents and young adults in Sweden. *Psychol Med* 48, 498-507.

Bloch, M.H., Qawasmi, A., 2011. Omega-3 fatty acid supplementation for the treatment of children with attention-deficit/hyperactivity disorder symptomatology: systematic review and meta-analysis. *J Am Acad Child Adolesc Psychiatry* 50, 991-1000.

Boedhoe, P.S.W., van Rooij, D., Hoogman, M., Twisk, J.W.R., Schmaal, L., Abe, Y., Alonso, P., Ameis, S.H., Anikin, A., Anticevic, A., Arango, C., Arnold, P.D., Asherson, P., Assogna, F., Auzias, G., Banaschewski, T., Baranov, A., Batistuzzo, M.C., Baumeister, S., Baur-Streubel, R., Behrmann, M., Bellgrove, M.A., Benedetti, F., Beucke, J.C., Biederman, J., Bollettini, I., Bose, A., Bralten, J., Bramati, I.E., Brandeis, D., Brem, S., Brennan, B.P., Busatto, G.F., Calderoni, S., Calvo, A., Calvo, R., Castellanos, F.X., Cercignani, M., Chaim-Avancini, T.M., Chantiluke, K.C., Cheng, Y., Cho, K.I.K., Christakou, A., Coghill, D., Conzelmann, A., Cubillo, A.I., Dale, A.M., Dallaspezia, S., Daly, E., Denys, D., Deruelle, C., Di Martino, A., Dinstein, I., Doyle, A.E., Durston, S., Earl, E.A., Ecker, C., Ehrlich, S., Ely, B.A., Epstein, J.N., Ethofer, T., Fair, D.A., Fallgatter, A.J., Faraone, S.V., Fedor, J., Feng, X., Feusner, J.D., Fitzgerald, J., Fitzgerald, K.D., Fouché, J.P., Freitag, C.M., Fridgeirsson, E.A., Frodl, T., Gabel, M.C., Gallagher, L., Gogberashvili, T., Gori, I., Gruner, P., Gürsel, D.A., Haar, S., Haavik, J., Hall, G.B., Harrison, N.A., Hartman, C.A., Heslenfeld, D.J., Hirano, Y., Hoekstra, P.J., Hoexter, M.Q., Hohmann, S., Høvik, M.F., Hu, H., Huyser, C., Jahanshad, N., Jalbrzikowski, M., James, A., Janssen, J., Jaspers-Fayer, F., Jernigan, T.L., Kapilushniy, D., Kardatzki, B., Karkashadze, G., Kathmann, N., Kaufmann, C., Kelly, C., Khadka, S., King, J.A., Koch, K., Kohls, G., Konrad, K., Kuno, M., Kuntsi, J., Kvale, G., Kwon, J.S., Lázaro, L., Lera-Miguel, S., Lesch, K.P., Hoekstra, L., Liu, Y., Lochner, C., Louza, M.R., Luna, B., Lundervold, A.J., Malpas, C.B., Marques, P., Marsh, R., Martínez-Zalacaín, I., Mataix-Cols, D., Mattos, P., McCarthy, H., McGrath, J., Mehta, M.A., Menchón, J.M., Mennes, M., Martinho, M.M., Moreira, P.S., Morer, A., Morgado, P., Muratori, F., Murphy, C.M., Murphy, D.G.M., Nakagawa, A., Nakamae, T., Nakao, T., Namazova-Baranova, L., Narayanaswamy, J.C., Nicolau, R., Nigg, J.T., Novotny, S.E., Nurmi, E.L., Weiss, E.O., O'Gorman Tuura, R.L., O'Hearn, K., O'Neill, J., Oosterlaan, J., Oranje, B., Paloyelis, Y., Parellada, M., Pauli, P., Perriello, C., Piacentini, J., Piras, F., Piras, F., Plessen, K.J., Puig, O., Ramos-Quiroga, J.A., Reddy, Y.C.J., Reif, A., Reneman, L., Retico, A., Rosa, P.G.P., Rubia, K., Rus, O.G., Sakai, Y., Schranke, A., Schwarz, L., Schweren, L.J.S., Seitz, J., Shaw, P., Shook, D., Silk, T.J., Simpson, H.B., Skokauskas, N., Soliva Vila, J.C., Solovieva, A., Soreni, N., Soriano-Mas, C., Spalletta, G., Stern, E.R., Stevens, M.C., Stewart, S.E., Sudre, G., Szeszko, P.R., Tamm, L., Taylor, M.J., Tolin, D.F., Tosetti, M., Tovar-Moll, F., Tsuchiyagaito, A., van Erp, T.G.M., van Wingen, G.A., Vance, A., Venkatasubramanian, G., Vilarroya, O., Vives-Gilabert, Y., von Polier, G.G., Walitza, S., Wallace, G.L., Wang, Z., Wolfers, T., Yoncheva, Y.N., Yun, J.Y., Zanetti, M.V., Zhou, F., Ziegler, G.C., Zierhut, K.C., Zwiers, M.P., Thompson, P.M., Stein, D.J., Buitelaar, J., Franke, B., van den Heuvel, O.A., 2020. Subcortical Brain Volume, Regional Cortical Thickness, and Cortical Surface Area Across Disorders: Findings From the ENIGMA ADHD, ASD, and OCD Working Groups (Online Ahead of Print). *Am J Psychiatry* 177, 834-843.

- Bolea-Alamanac, B., Nutt, D.J., Adamou, M., Asherson, P., Bazire, S., Coghill, D., Heal, D., Muller, U., Nash, J., Santosh, P., Sayal, K., Sonuga-Barke, E., Young, S.J., British Association for, P., 2014. Evidence-based guidelines for the pharmacological management of attention deficit hyperactivity disorder: update on recommendations from the British Association for Psychopharmacology. *J Psychopharmacol* 28, 179-203.
- Bonvicini, C., Cortese, S., Maj, C., Baune, B.T., Faraone, S.V., Scassellati, C., 2020. DRD4 48 bp multiallelic variants as age-population-specific biomarkers in attention-deficit/hyperactivity disorder. *Transl Psychiatry* 10, 70.
- Bonvicini, C., Faraone, S.V., Scassellati, C., 2016. Attention-deficit hyperactivity disorder in adults: A systematic review and meta-analysis of genetic, pharmacogenetic and biochemical studies. *Mol Psychiatry* 21, 1643.
- Bouchard, M.F., Bellinger, D.C., Wright, R.O., Weisskopf, M.G., 2010. Attention-deficit/hyperactivity disorder and urinary metabolites of organophosphate pesticides. *Pediatrics* 125, e1270-1277.
- Bradley, C., 1937. The behavior of children receiving benzedrine. *American Journal of Psychiatry* 94, 577-585.
- Braun, J.M., Kahn, R.S., Froehlich, T., Auinger, P., Lanphear, B.P., 2006. Exposures to environmental toxicants and attention deficit hyperactivity disorder in U.S. children. *Environ Health Perspect* 114, 1904-1909.
- Breslau, J., Miller, E., Joanie Chung, W.J., Schweitzer, J.B., 2011. Childhood and adolescent onset psychiatric disorders, substance use, and failure to graduate high school on time. *J Psychiatr Res* 45, 295-301.
- Bridgett, D.J., Walker, M.E., 2006. Intellectual functioning in adults with ADHD: a meta-analytic examination of full scale IQ differences between adults with and without ADHD. *Psychol Assess* 18, 1-14.
- Brikell, I., Ghirardi, L., D'Onofrio, B.M., Dunn, D.W., Almqvist, C., Dalsgaard, S., Kuja-Halkola, R., Larsson, H., 2018. Familial Liability to Epilepsy and Attention-Deficit/Hyperactivity Disorder: A Nationwide Cohort Study. *Biol Psychiatry* 83, 173-180.
- Brikell, I., Larsson, H., Lu, Y., Pettersson, E., Chen, Q., Kuja-Halkola, R., Karlsson, R., Lahey, B.B., Lichtenstein, P., Martin, J., 2020. The contribution of common genetic risk variants for ADHD to a general factor of childhood psychopathology. *Mol Psychiatry* 25, 1809-1821.
- Bruxel, E.M., Moreira-Maia, C.R., Akutagawa-Martins, G.C., Quinn, T.P., Klein, M., B., F., Ribasés, M., Rovira, P., Sánchez-Mora, C., Kappel, D.B., Mota, N.R., Grevet, E.H., Bau, C.H.D., Arcos-Burgos, M., Rohde, L.A., Hutz, M.H., 2020. Meta-analysis and systematic review of ADGRL3 (LPHN3) polymorphisms in ADHD susceptibility [Online ahead of print]. *Molecular Psychiatry*.
- Butwicka, A., Lichtenstein, P., Landen, M., Nordenvall, A.S., Nordenstrom, A., Nordenskjold, A., Frisen, L., 2015. Hypospadias and increased risk for neurodevelopmental disorders. *J Child Psychol Psychiatry* 56, 155-161.

Castells, X., Ramos-Quiroga, J.A., Bosch, R., Nogueira, M., Casas, M., 2011. Amphetamines for Attention Deficit Hyperactivity Disorder (ADHD) in adults. *Cochrane Database Syst Rev*, CD007813.

Catala-Lopez, F., Hutton, B., Nunez-Beltran, A., Page, M.J., Ridao, M., Macias Saint-Gerons, D., Catala, M.A., Tabares-Seisdedos, R., Moher, D., 2017. The pharmacological and non-pharmacological treatment of attention deficit hyperactivity disorder in children and adolescents: A systematic review with network meta-analyses of randomised trials. *PLoS One* 12, e0180355.

Caye, A., Petresco, S., de Barros, A.J.D., Bressan, R.A., Gadelha, A., Goncalves, H., Manfro, A.G., Matijasevich, A., Menezes, A.M.B., Miguel, E.C., Munhoz, T.N., Pan, P.M., Salum, G.A., Santos, I.S., Kielling, C., Rohde, L.A., 2020. Relative Age and Attention-Deficit/Hyperactivity Disorder: Data From Three Epidemiological Cohorts and a Meta-Analysis. *J Am Acad Child Adolesc Psychiatry* 59, 990-997.

Cederlof, M., Ohlsson Gotby, A., Larsson, H., Serlachius, E., Boman, M., Langstrom, N., Landen, M., Lichtenstein, P., 2014. Klinefelter syndrome and risk of psychosis, autism and ADHD. *J Psychiatr Res* 48, 128-130.

Cénat, J.M., Blais-Rochette, C., Morse, C., Vandette, M.P., Noorishad, P.G., Kogan, C., Ndengeyingoma, A., Labelle, P.R., 2020. Prevalence and Risk Factors Associated With Attention-Deficit/Hyperactivity Disorder Among US Black Individuals: A Systematic Review and Meta-analysis (Online ahead of print). *JAMA Psychiatry*.

Cepeda, M.S., Fife, D., Berwaerts, J., Yuan, Y., Mastrogiovanni, G., 2014. Shopping behavior for ADHD drugs: results of a cohort study in a pharmacy database. *Drugs R D* 14, 205-211.

Chang, J.P., Su, K.P., Mondelli, V., Pariante, C.M., 2018. Omega-3 Polyunsaturated Fatty Acids in Youths with Attention Deficit Hyperactivity Disorder: a Systematic Review and Meta-Analysis of Clinical Trials and Biological Studies. *Neuropsychopharmacology* 43, 534-545.

Chang, Z., D'Onofrio, B.M., Quinn, P.D., Lichtenstein, P., Larsson, H., 2016. Medication for Attention-Deficit/Hyperactivity Disorder and Risk for Depression: A Nationwide Longitudinal Cohort Study. *Biol Psychiatry* 80, 916-922.

Chang, Z., Lichtenstein, P., D'Onofrio, B.M., Almqvist, C., Kuja-Halkola, R., Sjolander, A., Larsson, H., 2014a. Maternal age at childbirth and risk for ADHD in offspring: a population-based cohort study. *Int J Epidemiol* 43, 1815-1824.

Chang, Z., Lichtenstein, P., D'Onofrio, B.M., Sjolander, A., Larsson, H., 2014b. Serious transport accidents in adults with attention-deficit/hyperactivity disorder and the effect of medication: a population-based study. *JAMA Psychiatry* 71, 319-325.

Chang, Z., Lichtenstein, P., Halldner, L., D'Onofrio, B., Serlachius, E., Fazel, S., Langstrom, N., Larsson, H., 2014c. Stimulant ADHD medication and risk for substance abuse. *J Child Psychol Psychiatry* 55, 878-885.

Chang, Z., Quinn, P.D., Hur, K., Gibbons, R.D., Sjolander, A., Larsson, H., D'Onofrio, B.M., 2017.

Association Between Medication Use for Attention-Deficit/Hyperactivity Disorder and Risk of Motor Vehicle Crashes. *JAMA Psychiatry* 74, 597-603.

- Chen, L., Hu, X., Ouyang, L., He, N., Liao, Y., Liu, Q., Zhou, M., Wu, M., Huang, X., Gong, Q., 2016. A systematic review and meta-analysis of tract-based spatial statistics studies regarding attention-deficit/hyperactivity disorder. *Neurosci Biobehav Rev* 68, 838-847.
- Chen, M.H., Hsu, J.W., Huang, K.L., Bai, Y.M., Ko, N.Y., Su, T.P., Li, C.T., Lin, W.C., Tsai, S.J., Pan, T.L., Chang, W.H., Chen, T.J., 2018a. Sexually Transmitted Infection Among Adolescents and Young Adults With Attention-Deficit/Hyperactivity Disorder: A Nationwide Longitudinal Study. *J Am Acad Child Adolesc Psychiatry* 57, 48-53.
- Chen, M.H., Pan, T.L., Hsu, J.W., Huang, K.L., Su, T.P., Li, C.T., Lin, W.C., Tsai, S.J., Chang, W.H., Chen, T.J., Bai, Y.M., 2018b. Risk of Type 2 Diabetes in Adolescents and Young Adults With Attention-Deficit/Hyperactivity Disorder: A Nationwide Longitudinal Study. *J Clin Psychiatry* 79, 17m11607.
- Chen, M.H., Pan, T.L., Huang, K.L., Hsu, J.W., Bai, Y.M., Su, T.P., Li, C.T., Tsai, S.J., Cheng, C.M., Chen, T.J., 2019a. Coaggregation of Major Psychiatric Disorders in First-Degree Relatives of Individuals With Attention-Deficit/Hyperactivity Disorder: A Nationwide Population-Based Study. *J Clin Psychiatry* 80.
- Chen, M.H., Pan, T.L., Wang, P.W., Hsu, J.W., Huang, K.L., Su, T.P., Li, C.T., Lin, W.C., Tsai, S.J., Chen, T.J., Bai, Y.M., 2019b. Prenatal Exposure to Acetaminophen and the Risk of Attention-Deficit/Hyperactivity Disorder: A Nationwide Study in Taiwan. *J Clin Psychiatry* 80.
- Chen, M.H., Su, T.P., Chen, Y.S., Hsu, J.W., Huang, K.L., Chang, W.H., Chen, T.J., Bai, Y.M., 2017a. Comorbidity of Allergic and Autoimmune Diseases Among Patients With ADHD. *J Atten Disord* 21, 219-227.
- Chen, Q., Hartman, C.A., Haavik, J., Harro, J., Klungsoyr, K., Hegvik, T.A., Wanders, R., Ottosen, C., Dalsgaard, S., Faraone, S.V., Larsson, H., 2018c. Common psychiatric and metabolic comorbidity of adult attention-deficit/hyperactivity disorder: A population-based cross-sectional study. *PLoS One* 13, e0204516.
- Chen, Q., Sjolander, A., Runeson, B., D'Onofrio, B.M., Lichtenstein, P., Larsson, H., 2014. Drug treatment for attention-deficit/hyperactivity disorder and suicidal behaviour: register based study. *BMJ* 348, g3769.
- Chen, V.C., Chan, H.L., Wu, S.I., Lee, M., Lu, M.L., Liang, H.Y., Dewey, M.E., Stewart, R., Lee, C.T., 2019c. Attention-Deficit/Hyperactivity Disorder and Mortality Risk in Taiwan. *JAMA Netw Open* 2, e198714.
- Chen, V.C., Chan, H.L., Wu, S.I., Lu, M.L., Dewey, M.E., Stewart, R., Lee, C.T., 2020a. Methylphenidate and mortality in children with attention-deficit hyperactivity disorder: population-based cohort study. *Br J Psychiatry*, 1-9.
- Chen, V.C., Yang, Y.H., Liao, Y.T., Kuo, T.Y., Liang, H.Y., Huang, K.Y., Huang, Y.C., Lee, Y., McIntyre, R.S., Lin, T.C., 2017b. The association between methylphenidate treatment and the risk for fracture among young ADHD patients: A nationwide population-based study in Taiwan. *PLoS One* 12, e0173762.

Chen, V.C., Yang, Y.H., Yu Kuo, T., Lu, M.L., Tseng, W.T., Hou, T.Y., Yeh, J.Y., Lee, C.T., Chen, Y.L., Lee, M.J., Dewey, M.E., Gossop, M., 2020b. Methylphenidate and the risk of burn injury among children with attention-deficit/hyperactivity disorder. *Epidemiol Psychiatr Sci* 29, e146.

Cheng, C.H., Chan, P.S., Hsieh, Y.W., Chen, K.F., 2016. A meta-analysis of mismatch negativity in children with attention deficit-hyperactivity disorders. *Neurosci Lett* 612, 132-137.

Cheng, J.Y., Chen, R.Y., Ko, J.S., Ng, E.M., 2007. Efficacy and safety of atomoxetine for attention-deficit/hyperactivity disorder in children and adolescents-meta-analysis and meta-regression analysis. *Psychopharmacology (Berl)* 194, 197-209.

Chinese Society of Psychiatry, 2001. Chinese Classification and diagnostic criteria of Mental Disorder, 3rd Edition.). Shandong science and technology press, Jinan, China.

Ching, C., Eslick, G.D., Poulton, A.S., 2019. Evaluation of Methylphenidate Safety and Maximum-Dose Titration Rationale in Attention-Deficit/Hyperactivity Disorder: A Meta-analysis. *JAMA Pediatr* 173, 630-639.

Choi, Y., Shin, J., Cho, K.H., Park, E.C., 2017. Change in household income and risk for attention deficit hyperactivity disorder during childhood: A nationwide population-based cohort study. *J Epidemiol* 27, 56-62.

Chou, I.C., Chang, Y.T., Chin, Z.N., Muo, C.H., Sung, F.C., Kuo, H.T., Tsai, C.H., Kao, C.H., 2013. Correlation between epilepsy and attention deficit hyperactivity disorder: a population-based cohort study. *PLoS One* 8, e57926.

Chou, I.C., Lin, C.C., Sung, F.C., Kao, C.H., 2014. Attention-deficit hyperactivity disorder increases the risk of deliberate self-poisoning: A population-based cohort. *Eur Psychiatry* 29, 523-527.

Christensen, J., Pedersen, L., Sun, Y., Dreier, J.W., Brikell, I., Dalgaard, S., 2019. Association of Prenatal Exposure to Valproate and Other Antiepileptic Drugs With Risk for Attention-Deficit/Hyperactivity Disorder in Offspring. *JAMA Netw Open* 2, e186606.

Christoffersen, M.N., 2019. Violent crime against children with disabilities: A nationwide prospective birth cohort-study. *Child Abuse Negl* 98, 104150.

Christoffersen, M.N., 2020. Sexual Crime Against Schoolchildren With Disabilities: A Nationwide Prospective Birth Cohort Study. *J Interpers Violence*, 886260520934442.

Chudal, R., Joelsson, P., Gyllenberg, D., Lehti, V., Leivonen, S., Hinkka-Yli-Salomaki, S., Gissler, M., Sourander, A., 2015. Parental age and the risk of attention-deficit/hyperactivity disorder: a nationwide, population-based cohort study. *J Am Acad Child Adolesc Psychiatry* 54, 487-494.e481.

Cohen, J., 1988. Statistical Power Analysis for the Behavioral Sciences, Second Edition ed. Erlbaum, Hillsdale, NJ.

Cooper, R.E., Tye, C., Kuntsi, J., Vassos, E., Asherson, P., 2016. The effect of omega-3 polyunsaturated fatty acid supplementation on emotional dysregulation, oppositional behaviour and conduct problems in ADHD: A systematic review and meta-analysis. *J Affect Disord* 190, 474-482.

Cortese, S., Adamo, N., Del Giovane, C., Mohr-Jensen, C., Hayes, A.J., Carucci, S., Atkinson, L.Z., Tessari, L., Banaschewski, T., Coghill, D., Hollis, C., Simonoff, E., Zuddas, A., Barbui, C., Purgato, M., Steinhausen, H.C., Shokraneh, F., Xia, J., Cipriani, A., 2018a. Comparative efficacy and tolerability of medications for attention-deficit hyperactivity disorder in children, adolescents, and adults: a systematic review and network meta-analysis. *Lancet Psychiatry* 5, 727-738.

Cortese, S., Ferrin, M., Brandeis, D., Buitelaar, J., Daley, D., Dittmann, R.W., Holtmann, M., Santosh, P., Stevenson, J., Stringaris, A., Zuddas, A., Sonuga-Barke, E.J., European, A.G.G., 2015. Cognitive training for attention-deficit/hyperactivity disorder: meta-analysis of clinical and neuropsychological outcomes from randomized controlled trials. *J Am Acad Child Adolesc Psychiatry* 54, 164-174.

Cortese, S., Ferrin, M., Brandeis, D., Holtmann, M., Aggensteiner, P., Daley, D., Santosh, P., Simonoff, E., Stevenson, J., Stringaris, A., Sonuga-Barke, E.J., European, A.G.G., 2016a. Neurofeedback for Attention-Deficit/Hyperactivity Disorder: Meta-Analysis of Clinical and Neuropsychological Outcomes From Randomized Controlled Trials. *J Am Acad Child Adolesc Psychiatry* 55, 444-455.

Cortese, S., Moreira-Maia, C.R., St Fleur, D., Morcillo-Penalver, C., Rohde, L.A., Faraone, S.V., 2016b. Association Between ADHD and Obesity: A Systematic Review and Meta-Analysis. *Am J Psychiatry* 173, 34-43.

Cortese, S., Sun, S., Zhang, J., Sharma, E., Chang, Z., Kuja-Halkola, R., Almqvist, C., Larsson, H., Faraone, S.V., 2018b. Association between attention deficit hyperactivity disorder and asthma: a systematic review and meta-analysis and a Swedish population-based study. *Lancet Psychiatry* 5, 717-726.

Coughlin, C.G., Cohen, S.C., Mulqueen, J.M., Ferracioli-Oda, E., Stuckelman, Z.D., Bloch, M.H., 2015. Meta-Analysis: Reduced Risk of Anxiety with Psychostimulant Treatment in Children with Attention-Deficit/Hyperactivity Disorder. *J Child Adolesc Psychopharmacol* 25, 611-617.

Craven, M.P., Groom, M.J., 2015. Computer games for user engagement in Attention Deficit Hyperactivity Disorder (ADHD) monitoring and therapy, 2015 International Conference on Interactive Technologies and Games (iTAG),. IEEE Computer Society conference proceedings, Nottingham, Nottinghamshire, United Kingdom, 22-23, pp. 34-40.

Crunelle, C.L., van den Brink, W., Moggi, F., Konstenius, M., Franck, J., Levin, F.R., van de Glind, G., Demetrovics, Z., Coetsee, C., Luderer, M., Schellekens, A., group, I.c., Matthys, F., 2018. International Consensus Statement on Screening, Diagnosis and Treatment of Substance Use Disorder Patients with Comorbid Attention Deficit/Hyperactivity Disorder. *Eur Addict Res* 24, 43-51.

Cunill, R., Castells, X., Tobias, A., Capella, D., 2013. Atomoxetine for attention deficit hyperactivity disorder in the adulthood: a meta-analysis and meta-regression. *Pharmacoeconom Drug Saf* 22, 961-969.

Curry, A.E., Metzger, K.B., Pfeiffer, M.R., Elliott, M.R., Winston, F.K., Power, T.J., 2017. Motor Vehicle Crash Risk Among Adolescents and Young Adults With Attention-Deficit/Hyperactivity Disorder. *JAMA Pediatr* 171, 756-763.

Daley, D., Jacobsen, R.H., Lange, A.M., Sorensen, A., Walldorf, J., 2019. The economic burden of adult attention deficit hyperactivity disorder: A sibling comparison cost analysis. *Eur Psychiatry* 61, 41-48.

Dalsgaard, S., Kvist, A.P., Leckman, J.F., Nielsen, H.S., Simonsen, M., 2014. Cardiovascular safety of stimulants in children with attention-deficit/hyperactivity disorder: a nationwide prospective cohort study. *J Child Adolesc Psychopharmacol* 24, 302-310.

Dalsgaard, S., Leckman, J.F., Mortensen, P.B., Nielsen, H.S., Simonsen, M., 2015a. Effect of drugs on the risk of injuries in children with attention deficit hyperactivity disorder: a prospective cohort study. *Lancet Psychiatry* 2, 702-709.

Dalsgaard, S., Ostergaard, S.D., Leckman, J.F., Mortensen, P.B., Pedersen, M.G., 2015b. Mortality in children, adolescents, and adults with attention deficit hyperactivity disorder: a nationwide cohort study. *Lancet* 385, 2190-2196.

de Graaf, R., Kessler, R.C., Fayyad, J., ten Have, M., Alonso, J., Angermeyer, M., Borges, G., Demyttenaere, K., Gasquet, I., de Girolamo, G., Haro, J.M., Jin, R., Karam, E.G., Ormel, J., Posada-Villa, J., 2008. The prevalence and effects of adult attention-deficit/hyperactivity disorder (ADHD) on the performance of workers: results from the WHO World Mental Health Survey Initiative. *Occup Environ Med* 65, 835-842.

Dekkers, T.J., Popma, A., Agelink van Rentergem, J.A., Bexkens, A., Huizenga, H.M., 2016. Risky decision making in Attention-Deficit/Hyperactivity Disorder: A meta-regression analysis. *Clin Psychol Rev* 45, 1-16.

Demontis, D., Walters, R.K., Martin, J., Mattheisen, M., Als, T.D., Agerbo, E., Baldursson, G., Belliveau, R., Bybjerg-Grauholm, J., Baekvad-Hansen, M., Cerrato, F., Chambert, K., Churchhouse, C., Dumont, A., Eriksson, N., Gandal, M., Goldstein, J.I., Grasby, K.L., Grove, J., Gudmundsson, O.O., Hansen, C.S., Hauberg, M.E., Hollegaard, M.V., Howrigan, D.P., Huang, H., Maller, J.B., Martin, A.R., Martin, N.G., Moran, J., Pallesen, J., Palmer, D.S., Pedersen, C.B., Pedersen, M.G., Poterba, T., Poulsen, J.B., Ripke, S., Robinson, E.B., Satterstrom, F.K., Stefansson, H., Stevens, C., Turley, P., Walters, G.B., Won, H., Wright, M.J., Consortium, A.W.G.o.t.P.G., Early, L., Genetic Epidemiology, C., and Me Research, T., Andreassen, O.A., Asherson, P., Burton, C.L., Boomsma, D.I., Cormand, B., Dalsgaard, S., Franke, B., Gelernter, J., Geschwind, D., Hakonarson, H., Haavik, J., Kranzler, H.R., Kuntsi, J., Langley, K., Lesch, K.P., Middeldorp, C., Reif, A., Rohde, L.A., Roussos, P., Schachar, R., Sklar, P., Sonuga-Barke, E.J.S., Sullivan, P.F., Thapar, A., Tung, J.Y., Waldman, I.D., Medland, S.E., Stefansson, K., Nordentoft, M., Hougaard, D.M., Werge, T., Mors, O., Mortensen, P.B., Daly, M.J., Faraone, S.V., Borglum, A.D., Neale, B.M., 2019. Discovery of the first genome-wide significant risk loci for attention deficit/hyperactivity disorder. *Nat Genet* 51, 63-75.

Dey, M., Paz Castro, R., Haug, S., Schaub, M.P., 2019. Quality of life of parents of mentally-ill children: a systematic review and meta-analysis. *Epidemiol Psychiatr Sci* 28, 563-577.

- Dobrosavljevic, M., Solares, C., Cortese, S., Andershed, H., Larsson, H., 2020. Prevalence of attention-deficit/hyperactivity disorder in older adults: A systematic review and meta-analysis. *Neurosci Biobehav Rev* 118, 282-289.
- Dong, T., Hu, W., Zhou, X., Lin, H., Lan, L., Hang, B., Lv, W., Geng, Q., Xia, Y., 2018. Prenatal Exposure to Maternal Smoking during Pregnancy and Attention-deficit/hyperactivity Disorder in Offspring: A Meta-analysis. *Reproductive Toxicology* 76, 63-70.
- Doshi, J.A., Hodgkins, P., Kahle, J., Sikirica, V., Cangelosi, M.J., Setyawan, J., Erder, M.H., Neumann, P.J., 2012. Economic impact of childhood and adult attention-deficit/hyperactivity disorder in the United States. *J Am Acad Child Adolesc Psychiatry* 51, 990-1002.e1002.
- DosReis, S., Barksdale, C.L., Sherman, A., Maloney, K., Charach, A., 2010. Stigmatizing experiences of parents of children with a new diagnosis of ADHD. *Psychiatr Serv* 61, 811-816.
- Dovis, S., Van der Oord, S., Wiers, R.W., Prins, P.J., 2015. Improving Executive Functioning in Children with ADHD: Training Multiple Executive Functions within the Context of a Computer Game. A Randomized Double-Blind Placebo Controlled Trial. *PLoS One* 10, e0121651.
- Du Rietz, E., Jangmo, A., Kuja-Halkola, R., Chang, Z., D'Onofrio, B.M., Ahnemark, E., Werner-Kiechle, T., Larsson, H., 2020. Trajectories of healthcare utilization and costs of psychiatric and somatic multimorbidity in adults with childhood ADHD: a prospective register-based study [Epub ahead of print]. *J Child Psychol Psychiatry* 61, 959-968.
- Duh-Leong, C., Fuller, A., Brown, N.M., 2020. Associations Between Family and Community Protective Factors and Attention-Deficit/Hyperactivity Disorder Outcomes Among US Children. *J Dev Behav Pediatr* 41, 1-8.
- Ellis, P.D., 2010. *Essential Guide to Effect Sizes*. 41.
- Engel, S.M., Villanger, G.D., Nethery, R.C., Thomsen, C., Sakhi, A.K., Drover, S.S.M., Hoppin, J.A., Zeiner, P., Knudsen, G.P., Reichborn-Kjennerud, T., Herring, A.H., Aase, H., 2018. Prenatal Phthalates, Maternal Thyroid Function, and Risk of Attention-Deficit Hyperactivity Disorder in the Norwegian Mother and Child Cohort. *Environ Health Perspect* 126, 057004.
- Faraone, S.V., 2005. The scientific foundation for understanding attention-deficit/hyperactivity disorder as a valid psychiatric disorder. *Eur Child Adolesc Psychiatry* 14, 1-10.
- Faraone, S.V., Asherson, P., Banaschewski, T., Biederman, J., Buitelaar, J.K., Ramos-Quiroga, J.A., Rohde, L.A., Sonuga-Barke, E.J., Tannock, R., Franke, B., 2015. Attention-deficit/hyperactivity disorder. *Nat Rev Dis Primers* 1, 15020.
- Faraone, S.V., Biederman, J., Mick, E., 2006. The age-dependent decline of attention deficit hyperactivity disorder: a meta-analysis of follow-up studies. *Psychol Med* 36, 159-165.
- Faraone, S.V., Biederman, J., Morley, C.P., Spencer, T.J., 2008. Effect of stimulants on height and weight: a review of the literature. *J Am Acad Child Adolesc Psychiatry* 47, 994-1009.

Faraone, S.V., Biederman, J., Roe, C.M., 2002. Comparative efficacy of adderall and methylphenidate in attention-deficit/hyperactivity disorder: a meta-analysis. *Journal of Clinical Psychopharmacology* 22, 468-473.

Faraone, S.V., Hess, J., Wilens, T., 2019a. Prevalence and Consequences of the Nonmedical Use of Amphetamine Among Persons Calling Poison Control Centers. *J Atten Disord* Vol. 23(11), 1219-1228.

Faraone, S.V., Larsson, H., 2018. Genetics of attention deficit hyperactivity disorder. *Mol Psychiatry* 24, 562-575.

Faraone, S.V., Rostain, A.L., Blader, J., Busch, B., Childress, A.C., Connor, D.F., Newcorn, J.H., 2019b. Practitioner Review: Emotional dysregulation in attention-deficit/hyperactivity disorder - implications for clinical recognition and intervention. *J Child Psychol Psychiatry* 60, 133-150.

Faraone, S.V., Rostain, A.L., Montano, C.B., Mason, O., Antshel, K.M., Newcorn, J.H., 2020. Systematic Review: Nonmedical Use of Prescription Stimulants: Risk Factors, Outcomes, and Risk Reduction Strategies. *J Am Acad Child Adolesc Psychiatry* 59, 100-112.

Faraone, S.V., Spencer, T., Aleardi, M., Pagano, C., Biederman, J., 2004. Meta-analysis of the efficacy of methylphenidate for treating adult attention deficit hyperactivity disorder. *Journal of Clinical Psychopharmacology* 54, 24-29.

Farsad-Naeimi, A., Asjodi, F., Omidian, M., Askari, M., Nouri, M., Pizarro, A.B., Daneshzad, E., 2020. Sugar consumption, sugar sweetened beverages and Attention Deficit Hyperactivity Disorder: A systematic review and meta-analysis. *Complement Ther Med* 53, 102512.

Fayyad, J., Sampson, N.A., Hwang, I., Adamowski, T., Aguilar-Gaxiola, S., Al-Hamzawi, A., Andrade, L.H., Borges, G., de Girolamo, G., Florescu, S., Gureje, O., Haro, J.M., Hu, C., Karam, E.G., Lee, S., Navarro-Mateu, F., O'Neill, S., Pennell, B.E., Piazza, M., Posada-Villa, J., Ten Have, M., Torres, Y., Xavier, M., Zaslavsky, A.M., Kessler, R.C., 2017. The descriptive epidemiology of DSM-IV Adult ADHD in the World Health Organization World Mental Health Surveys. *Atten Defic Hyperact Disord* 9, 47-65.

Feldman, H.M., Reiff, M.I., 2014. Clinical practice. Attention deficit-hyperactivity disorder in children and adolescents. *N Engl J Med* 370, 838-846.

Fitzgerald, C., Dalsgaard, S., Nordentoft, M., Erlangsen, A., 2019. Suicidal behaviour among persons with attention-deficit hyperactivity disorder. *Br J Psychiatry*, 1-6.

Fleming, M., Fitton, C.A., Steiner, M.F.C., McLay, J.S., Clark, D., King, A., Mackay, D.F., Pell, J.P., 2017.

Educational and Health Outcomes of Children Treated for Attention-Deficit/Hyperactivity Disorder. *JAMA Pediatr* 171, e170691.

Fletcher, J.M., 2014. The effects of childhood ADHD on adult labor market outcomes. *Health Econ* 23, 159-181.

Flisher, A.J., Hawkrigde, S., 2013. Attention deficit hyperactivity disorder in children and adolescents. *South African Journal of Psychiatry* 19, 136-140.

Forns, J., Verner, M.A., Iszatt, N., Nowack, N., Bach, C.C., Vrijheid, M., Costa, O., Andiaarena, A., Sovcikova, E., Høyer, B.B., Wittsiepe, J., Lopez-Espinosa, M.J., Ibarluzea, J., Hertz-Picciotto, I., Toft, G., Stigum, H., Guxens, M., Liew, Z., Eggesbø, M., 2020. Early Life Exposure to Perfluoroalkyl Substances (PFAS) and ADHD: A Meta-Analysis of Nine European Population-Based Studies. *Environ Health Perspect* 128, 57002.

Franz, A.P., Bolat, G.U., Bolat, H., Matijasevich, A., Santos, I.S., Silveira, R.C., Procianoy, R.S., Rohde, L.A., Moreira-Maia, C.R., 2018. Attention-Deficit/Hyperactivity Disorder and Very Preterm/Very Low Birth Weight: A Meta-analysis. *Pediatrics* 141, e20171645.

Frazier, T.W., Demaree, H.A., Youngstrom, E.A., 2004. Meta-analysis of intellectual and neuropsychological test performance in attention-deficit/hyperactivity disorder. *Neuropsychology* 18, 543-555.

Froehlich, T.E., Lanphear, B.P., Auinger, P., Hornung, R., Epstein, J.N., Braun, J., Kahn, R.S., 2009. Association of tobacco and lead exposures with attention-deficit/hyperactivity disorder. *Pediatrics* 124, e1054-1063.

Ge, G.M., Leung, M.T.Y., Man, K.K.C., Leung, W.C., Ip, P., Li, G.H.Y., Wong, I.C.K., Kung, A.W.C., Cheung, C.L., 2020. Maternal thyroid dysfunction during pregnancy and the risk of adverse outcomes in the offspring: a systematic review and meta-analysis. *J Clin Endocrinol Metab*.

Ghirardi, L., Brikell, I., Kuja-Halkola, R., Freitag, C.M., Franke, B., Asherson, P., Lichtenstein, P., Larsson, H., 2018. The familial co-aggregation of ASD and ADHD: a register-based cohort study. *Mol Psychiatry* 23, 257-262.

Ghirardi, L., Chen, Q., Chang, Z., Kuja-Halkola, R., Skoglund, C., Quinn, P.D., D'Onofrio, B.M., Larsson, H., 2020. Use of medication for attention-deficit/hyperactivity disorder and risk of unintentional injuries in children and adolescents with co-occurring neurodevelopmental disorders. *J Child Psychol Psychiatry* 61, 140-147.

Goodlad, J.K., Marcus, D.K., Fulton, J.J., 2013. Lead and Attention-Deficit/Hyperactivity Disorder (ADHD) symptoms: A meta-analysis. *Clin Psychol Rev* 33, 417-425.

Graham, J., Banaschewski, T., Buitelaar, J., Coghill, D., Danckaerts, M., Dittmann, R.W., Dopfner, M., Hamilton, R., Hollis, C., Holtmann, M., Hulpke-Wette, M., Lecendreux, M., Rosenthal, E., Rothenberger, A., Santosh, P., Sergeant, J., Simonoff, E., Sonuga-Barke, E., Wong, I.C., Zuddas, A., Steinhausen, H.C., Taylor, E., European Guidelines, G., 2011. European guidelines on managing adverse effects of medication for ADHD. *Eur Child Adolesc Psychiatry* 20, 17-37.

Graziano, P.A., Garcia, A., 2016. Attention-deficit hyperactivity disorder and children's emotion dysregulation: A meta-analysis. *Clin Psychol Rev* 46, 106-123.

Groenman, A.P., Janssen, T.W.P., Oosterlaan, J., 2017. Childhood Psychiatric Disorders as Risk Factor for Subsequent Substance Abuse: A Meta-Analysis. *J Am Acad Child Adolesc Psychiatry* 56, 556-569.

Grunblatt, E., Nemoda, Z., Werling, A.M., Roth, A., Angyal, N., Tarnok, Z., Thomsen, H., Peters, T., Hinney, A., Hebebrand, J., Lesch, K.P., Romanos, M., Walitza, S., 2019a. The involvement of the canonical Wnt-signaling receptor LRP5 and LRP6 gene variants with ADHD and sexual dimorphism: Association study and meta-analysis. *Am J Med Genet B Neuropsychiatr Genet* 180, 365-376.

Grunblatt, E., Werling, A.M., Roth, A., Romanos, M., Walitza, S., 2019b. Association study and a systematic meta-analysis of the VNTR polymorphism in the 3'-UTR of dopamine transporter gene and attention-deficit hyperactivity disorder. *J Neural Transm (Vienna)* 126, 517-529.

Gudjonsson, G.H., Sigurdsson, J.F., Sigfusdottir, I.D., Asgeirsdottir, B.B., Gonzalez, R.A., Young, S., 2016. A national epidemiological study investigating risk factors for police interrogation and false confession among juveniles and young persons. *Soc Psychiatry Psychiatr Epidemiol* 51, 359-367.

Guo, N.W., Lin, C.L., Lin, C.W., Huang, M.T., Chang, W.L., Lu, T.H., Lin, C.J., 2016. Fracture risk and correlating factors of a pediatric population with attention deficit hyperactivity disorder: a nationwide matched study. *J Pediatr Orthop B* 25, 369-374.

Hart, H., Radua, J., Nakao, T., Mataix-Cols, D., Rubia, K., 2013. Meta-analysis of Functional Magnetic Resonance Imaging Studies of Inhibition and Attention in Attention-deficit/Hyperactivity Disorder: Exploring Task-Specific, Stimulant Medication, and Age Effects. *JAMA Psychiatry* 70, 185-198.
Hawkey, E., Nigg, J.T., 2014. Omega-3 fatty acid and ADHD: Blood level analysis and meta-analytic extension of supplementation trials. *Clin Psychol Rev* 34, 496-505.

Hegvik, T.A., Instanes, J.T., Haavik, J., Klungsoyr, K., Engeland, A., 2018. Associations between attention-deficit/hyperactivity disorder and autoimmune diseases are modified by sex: a population-based cross-sectional study. *Eur Child Adolesc Psychiatry* 27, 663-675.

Hilgard, D., Konrad, K., Meusers, M., Bartus, B., Otto, K.P., Lepler, R., Schober, E., Bollow, E., Holl, R.W., 2017. Comorbidity of attention deficit hyperactivity disorder and type 1 diabetes in children and adolescents: Analysis based on the multicentre DPV registry. *Pediatr Diabetes* 18, 706-713.

Ho, J.D., Sheu, J.J., Kao, Y.W., Shia, B.C., Lin, H.C., 2020. Associations between Attention-Deficit/Hyperactivity Disorder and Ocular Abnormalities in Children: A Population-based Study. *Ophthalmic Epidemiol* 27, 194-199.

Hoffmann, H., 1990. *Der Struwwelpeter : oder lustige Geschichten und drollige Bilder für Kinder von 3 bis 6 Jahren.* J.F. Schreiber, Esslingen

Hollis, C., Chen, Q., Chang, Z., Quinn, P.D., Viktorin, A., Lichtenstein, P., D'Onofrio, B., Landén, M., Larsson, H., 2019. Methylphenidate and the risk of psychosis in adolescents and young adults: a population-based cohort study. *The Lancet Psychiatry* 6, 651-658.

Holmskov, M., Storebo, O.J., Moreira-Maia, C.R., Ramstad, E., Magnusson, F.L., Krogh, H.B., Groth, C., Gillies, D., Zwi, M., Skoog, M., Gluud, C., Simonsen, E., 2017. Gastrointestinal adverse events during methylphenidate treatment of children and adolescents with attention deficit hyperactivity disorder: A systematic review with meta-analysis and Trial Sequential Analysis of randomised clinical trials. *PLoS One* 12, e0178187.

Hong, M., Park, B., Lee, S.M., Bahn, G.H., Kim, M.J., Park, S., Oh, I.H., Park, H., 2020. Economic Burden and Disability-Adjusted Life Years (DALYs) of Attention Deficit/Hyperactivity Disorder. *J Atten Disord* 24, 823-829.

Hoogman, M., Bralten, J., Hibar, D.P., Mennes, M., Zwiers, M.P., Schweren, L.S.J., van Hulzen, K.J.E., Medland, S.E., Shumskaya, E., Jahanshad, N., Zeeuw, P., Szekely, E., Sudre, G., Wolfers, T., Onnink, A.M.H., Dammers, J.T., Mostert, J.C., Vives-Gilabert, Y., Kohls, G., Oberwelland, E., Seitz, J., Schulte-Ruther, M., Ambrosino, S., Doyle, A.E., Hovik, M.F., Dramsdahl, M., Tamm, L., van Erp, T.G.M., Dale, A., Schork, A., Conzelmann, A., Zierhut, K., Baur, R., McCarthy, H., Yoncheva, Y.N., Cubillo, A., Chantiluke, K., Mehta, M.A., Paloyelis, Y., Hohmann, S., Baumeister, S., Bramati, I., Mattos, P., Tovar-Moll, F., Douglas, P., Banaschewski, T., Brandeis, D., Kuntsi, J., Asherson, P., Rubia, K., Kelly, C., Martino, A.D., Milham, M.P., Castellanos, F.X., Frodl, T., Zentis, M., Lesch, K.P., Reif, A., Pauli, P., Jernigan, T.L., Haavik, J., Plessen, K.J., Lundervold, A.J., Hugdahl, K., Seidman, L.J., Biederman, J., Rommelse, N., Heslenfeld, D.J., Hartman, C.A., Hoekstra, P.J., Oosterlaan, J., Polier, G.V., Konrad, K., Vilarroya, O., Ramos-Quiroga, J.A., Soliva, J.C., Durston, S., Buitelaar, J.K., Faraone, S.V., Shaw, P., Thompson, P.M., Franke, B., 2017. Subcortical brain volume differences in participants with attention deficit hyperactivity disorder in children and adults: a cross-sectional mega-analysis. *Lancet Psychiatry* 4, 310-319.

Hoogman, M., Muetzel, R., Guimaraes, J.P., Shumskaya, E., Mennes, M., Zwiers, M.P., Jahanshad, N., Sudre, G., Wolfers, T., Earl, E.A., Soliva Vila, J.C., Vives-Gilabert, Y., Khadka, S., Novotny, S.E., Hartman, C.A., Heslenfeld, D.J., Schweren, L.J.S., Ambrosino, S., Oranje, B., de Zeeuw, P., Chaim-Avancini, T.M., Rosa, P.G.P., Zanetti, M.V., Malpas, C.B., Kohls, G., von Polier, G.G., Seitz, J., Biederman, J., Doyle, A.E., Dale, A.M., van Erp, T.G.M., Epstein, J.N., Jernigan, T.L., Baur-Streubel, R., Ziegler, G.C., Zierhut, K.C., Schranke, A., Hovik, M.F., Lundervold, A.J., Kelly, C., McCarthy, H., Skokauskas, N., O'Gorman Tuura, R.L., Calvo, A., Lera-Miguel, S., Nicolau, R., Chantiluke, K.C., Christakou, A., Vance, A., Cercignani, M., Gabel, M.C., Asherson, P., Baumeister, S., Brandeis, D., Hohmann, S., Bramati, I.E., Tovar-Moll, F., Fallgatter, A.J., Kardatzki, B., Schwarz, L., Anikin, A., Baranov, A., Gogberashvili, T., Kapilushniy, D., Solovieva, A., El Marroun, H., White, T., Karkashadze, G., Namazova-Baranova, L., Ethofer, T., Mattos, P., Banaschewski, T., Coghill, D., Plessen, K.J., Kuntsi, J., Mehta, M.A., Paloyelis, Y., Harrison, N.A., Bellgrove, M.A., Silk, T.J., Cubillo, A.I., Rubia, K., Lazaro, L., Brem, S., Walitza, S., Frodl, T., Zentis, M., Castellanos, F.X., Yoncheva, Y.N., Haavik, J., Reneman, L., Conzelmann, A., Lesch, K.P., Pauli, P., Reif, A., Tamm, L., Konrad, K., Oberwelland Weiss, E., Busatto, G.F., Louza, M.R., Durston, S., Hoekstra, P.J., Oosterlaan, J., Stevens, M.C., Ramos-Quiroga, J.A., Vilarroya, O., Fair, D.A., Nigg, J.T., Thompson, P.M., Buitelaar, J.K., Faraone, S.V., Shaw, P., Tiemeier, H., Bralten, J., Franke, B., 2019. Brain Imaging of the Cortex in ADHD: A Coordinated Analysis of Large-Scale Clinical and Population-Based Samples. *Am J Psychiatry* 176, 531-542.

Horton-Salway, M., 2013. Gendering attention deficit hyperactivity disorder: a discursive analysis of UK newspaper stories. *J Health Psychol* 18, 1085-1099.

Hua, M.H., Huang, K.L., Hsu, J.W., Bai, Y.M., Su, T.P., Tsai, S.J., Li, C.T., Lin, W.C., Chen, T.J., Chen, M.H., 2020. Early Pregnancy Risk Among Adolescents With ADHD: A Nationwide Longitudinal Study. *J Atten Disord*, 1087054719900232.

Huang, A., Wu, K., Cai, Z., Lin, Y., Zhang, X., Huang, Y., 2020. Association between postnatal second-hand smoke exposure and ADHD in children: a systematic review and meta-analysis. *Environ Sci Pollut Res Int*.

- Huang, K.L., Wei, H.T., Hsu, J.W., Bai, Y.M., Su, T.P., Li, C.T., Lin, W.C., Tsai, S.J., Chang, W.H., Chen, T.J., Chen, M.H., 2018. Risk of suicide attempts in adolescents and young adults with attention-deficit hyperactivity disorder: a nationwide longitudinal study. *Br J Psychiatry* 212, 234-238.
- Huang, L., Wang, Y., Zhang, L., Zheng, Z., Zhu, T., Qu, Y., Mu, D., 2017. Maternal Smoking and Attention-Deficit/Hyperactivity Disorder in Offspring: A Meta-analysis. *Pediatrics*, e20172465.
- Humphreys, K.L., Eng, T., Lee, S.S., 2013. Stimulant Medication and Substance Use Outcomes: A Meta-analysis. *JAMA Psychiatry*, 1-9.
- Huybrechts, K.F., Broms, G., Christensen, L.B., Einarsdottir, K., Engeland, A., Furu, K., Gissler, M., Hernandez-Diaz, S., Karlsson, P., Karlstad, O., Kieler, H., Lahesmaa-Korpinen, A.M., Mogun, H., Norgaard, M., Reutfors, J., Sorensen, H.T., Zoega, H., Bateman, B.T., 2018. Association Between Methylphenidate and Amphetamine Use in Pregnancy and Risk of Congenital Malformations: A Cohort Study From the International Pregnancy Safety Study Consortium. *JAMA Psychiatry* 75, 167-175.
- Jackson, J.N., MacKillop, J., 2016. Attention-Deficit/Hyperactivity Disorder and Monetary Delay Discounting: A Meta-Analysis of Case-Control Studies. *Biol Psychiatry Cogn Neurosci Neuroimaging* 1, 316-325.
- Jangmo, A., Stalhandske, A., Chang, Z., Chen, Q., Almqvist, C., Feldman, I., Bulik, C.M., Lichtenstein, P., D'Onofrio, B., Kuja-Halkola, R., Larsson, H., 2019. Attention-Deficit/Hyperactivity Disorder, School Performance, and Effect of Medication. *J Am Acad Child Adolesc Psychiatry* 58, 423-432.
- Jenabi, E., Bashirian, S., Khazaei, S., Basiri, Z., 2019. The maternal pre-pregnancy BMI and the risk of ADHD among children and adolescents: A systematic review and meta-Analysis. *Korean J Pediatr*.
- Jennum, P., Hastrup, L.H., Ibsen, R., Kjellberg, J., Simonsen, E., 2020. Welfare consequences for people diagnosed with attention deficit hyperactivity disorder (ADHD): A matched nationwide study in Denmark. *Eur Neuropsychopharmacol* 37, 29-38.
- Ji, J., Chen, T., Sundquist, J., Sundquist, K., 2018. Type 1 Diabetes in Parents and Risk of Attention Deficit/Hyperactivity Disorder in Offspring: A Population-Based Study in Sweden. *Diabetes Care* 41, 770-774.
- Joelsson, P., Chudal, R., Talati, A., Suominen, A., Brown, A.S., Sourander, A., 2016. Prenatal smoking exposure and neuropsychiatric comorbidity of ADHD: a finnish nationwide population-based cohort study. *BMC Psychiatry* 16, 306.
- Kapellen, T.M., Reimann, R., Kiess, W., Kostev, K., 2016. Prevalence of medically treated children with ADHD and type 1 diabetes in Germany - Analysis of two representative databases. *J Pediatr Endocrinol Metab* 29, 1293-1297.
- Katusic, M.Z., Voigt, R.G., Colligan, R.C., Weaver, A.L., Homan, K.J., Barbaresi, W.J., 2011. Attention-deficit hyperactivity disorder in children with high intelligence quotient: results from a population-based study. *J Dev Behav Pediatr* 32, 103-109.

Keilow, M., Holm, A., Fallesen, P., 2018. Medical treatment of Attention Deficit/Hyperactivity Disorder (ADHD) and children's academic performance. *PLoS One* 13, e0207905.

Keilow, M., Wu, C., Obel, C., 2020. Cumulative social disadvantage and risk of attention deficit hyperactivity disorder: Results from a nationwide cohort study. *SSM Popul Health* 10, 100548.

Kennedy, M., Kreppner, J., Knights, N., Kumsta, R., Maughan, B., Golm, D., Rutter, M., Schlotz, W., Sonuga-Barke, E.J., 2016. Early severe institutional deprivation is associated with a persistent variant of adult attention-deficit/hyperactivity disorder: clinical presentation, developmental continuities and life circumstances in the English and Romanian Adoptees study. *J Child Psychol Psychiatry* 57, 1113-1125.

Kidwell, K.M., Van Dyk, T.R., Lundahl, A., Nelson, T.D., 2015. Stimulant Medications and Sleep for Youth With ADHD: A Meta-analysis. *Pediatrics* 136, 1144-1153.

King, S.A., Casavant, M.J., Spiller, H.A., Hodges, N.L., Chounthirath, T., Smith, G.A., 2018. Pediatric ADHD Medication Exposures Reported to US Poison Control Centers. *Pediatrics* 141.

Kirova, A.M., Kelberman, C., Storch, B., DiSalvo, M., Woodworth, K.Y., Faraone, S.V., Biederman, J., 2019. Are subsyndromal manifestations of attention deficit hyperactivity disorder morbid in children? A systematic qualitative review of the literature with meta-analysis. *Psychiatry Res* 274, 75-90.

Knouse, L.E., Teller, J., Brooks, M.A., 2017. Meta-analysis of cognitive-behavioral treatments for adult ADHD. *J Consult Clin Psychol* 85, 737-750.

Kohler-Forsberg, O., Petersen, L., Gasse, C., Mortensen, P.B., Dalsgaard, S., Yolken, R.H., Mors, O., Benros, M.E., 2019. A Nationwide Study in Denmark of the Association Between Treated Infections and the Subsequent Risk of Treated Mental Disorders in Children and Adolescents. *JAMA Psychiatry* 76, 271-279.

Kooij, J.J.S., Bijlenga, D., Salerno, L., Jaeschke, R., Bitter, I., Balazs, J., Thome, J., Dom, G., Kasper, S., Nunes Filipe, C., Stes, S., Mohr, P., Leppamaki, S., Casas, M., Bobes, J., McCarthy, J.M., Richarte, V., Kjems Philipsen, A., Pehlivanidis, A., Niemela, A., Styr, B., Semerci, B., Bolea-Alamanac, B., Edvinsson, D., Baeyens, D., Wynchank, D., Sobanski, E., Philipsen, A., McNicholas, F., Caci, H., Mihailescu, I., Manor, I., Dobrescu, I., Saito, T., Krause, J., Fayyad, J., Ramos-Quiroga, J.A., Foeken, K., Rad, F., Adamou, M., Ohlmeier, M., Fitzgerald, M., Gill, M., Lensing, M., Motavalli Mukaddes, N., Brudkiewicz, P., Gustafsson, P., Tani, P., Oswald, P., Carpentier, P.J., De Rossi, P., Delorme, R., Markovska Simoska, S., Pallanti, S., Young, S., Bejerot, S., Lehtonen, T., Kustow, J., Muller-Sedgwick, U., Hirvikoski, T., Pironti, V., Ginsberg, Y., Felegyhazy, Z., Garcia-Portilla, M.P., Asherson, P., 2019. Updated European Consensus Statement on diagnosis and treatment of adult ADHD. *Eur Psychiatry* 56, 14-34.

Koren, G., Barer, Y., Ornoy, A., 2020. Fetal safety of methylphenidate-A scoping review and meta analysis. *Reprod Toxicol* 93, 230-234.

Korrel, H., Mueller, K.L., Silk, T., Anderson, V., Sciberras, E., 2017. Research Review: Language problems in children with Attention-Deficit Hyperactivity Disorder - a systematic meta-analytic review. *J Child Psychol Psychiatry* 58, 640-654.

Kramer, P.D.F., Pollnow, D.M.e.P.H., 1932. Über eine hyperkinetische Erkrankung im Kindesalter. pp. 21–40. *European Neurology* 82, 21-40.

Lafora, G.R., 1917. *Los Niños Mentalmente Anormales*. Madrid, 1917.

Lange, K.W., Reichl, S., Lange, K.M., Tucha, L., Tucha, O., 2010. The history of attention deficit hyperactivity disorder. *Atten Defic Hyperact Disord* 2, 241-255.

Larsson, H., Chang, Z., D'Onofrio, B.M., Lichtenstein, P., 2014a. The heritability of clinically diagnosed attention deficit hyperactivity disorder across the lifespan. *Psychol Med* 44, 2223-2239.

Larsson, H., Sariaslan, A., Langstrom, N., D'Onofrio, B., Lichtenstein, P., 2014b. Family income in early childhood and subsequent attention deficit/hyperactivity disorder: a quasi-experimental study. *J Child Psychol Psychiatry* 55, 428-435.

Le, H.H., Hodgkins, P., Postma, M.J., Kahle, J., Sikirica, V., Setyawan, J., Erder, M.H., Doshi, J.A., 2014. Economic impact of childhood/adolescent ADHD in a European setting: the Netherlands as a reference case. *Eur Child Adolesc Psychiatry* 23, 587-598.

Lebowitz, M.S., 2016. Stigmatization of ADHD: A Developmental Review. *J Atten Disord* 20, 199-205.
Lebwohl, B., Haggård, L., Emilsson, L., Söderling, J., Roelstraete, B., Butwicka, A., Green, P.H., Ludvigsson, J.F., 2020. Psychiatric disorders in patients with a diagnosis of celiac disease during childhood from 1973 to 2016. *Clin Gastroenterol Hepatol*.

Lee, P.H., Anttila, V., Won, H., Feng, Y.-C.A., Rosenthal, J., Zhu, Z., Tucker-Drob, E.M., Nivard, M.G., Grotzinger, A.D., Posthuma, D., Wang, M.M.J., Yu, D., Stahl, E., Walters, R.K., Anney, R.J.L., Duncan, L.E., Belangero, S., Luykx, J., Kranzler, H., Keski-Rahkonen, A., Cook, E.H., Kirov, G., Coppola, G., Kaprio, J., Zai, C.C., Hoekstra, P.J., Banaschewski, T., Rohde, L.A., Sullivan, P.F., Franke, B., Daly, M.J., Bulik, C.M., Lewis, C.M., McIntosh, A.M., Donovan, M.C., Zheutlin, A., Andreassen, O.A., Borglum, A.D., Breen, G., Edenberg, H.J., Fanous, A.H., Faraone, S.V., Gelernter, J., Mathews, C.A., Mattheisen, M., Mitchell, K., Neale, M.C., Nurnberger, J.I., Ripke, S., Santangelo, S.L., Scharf, J.M., Stein, M.B., Thornton, L.M., Walters, J.T.R., Wray, N.R., Geschwind, D.H., Neale, B., Kendler, K.S., Smoller, J.W., 2019. Genome wide meta-analysis identifies genomic relationships, novel loci, and pleiotropic mechanisms across eight psychiatric disorders. *bioRxiv*, 528117.

Lee PH, A.V., Won H, Feng YA, Rosenthal J, Zhu Z, Tucker-Drob EM, Nivard MG, Grotzinger AD, Posthuma D, Wang MM, Yu D, Stahl EA, Walters RK, Anney RJL, Duncan LE, Ge T, Adolfsson R, Banaschewski T, Belangero S, Cook EH, Coppola G, Derks EM, Hoekstra PJ, Kaprio J, Keski-Rahkonen A, Kirov G, Kranzler HR, Luykx JJ, Rohde LA, Zai CC, Agerbo E, Arranz MJ, Asherson P, Bækvad-Hansen M, Baldursson G, Bellgrove M, Belliveau RA Jr, Buitelaar J, Burton CL, Bybjerg-Grauholm J, Casas M, Cerrato F, Chambert K, Churchhouse C, Cormand B, Crosbie J, Dalsgaard S, Demontis D, Doyle AE, Dumont A, Elia J, Grove J, Gudmundsson OO, Haavik J, Hakonarson H, Hansen CS, Hartman CA, Hawi Z, Hervás A, Hougaard DM, Howrigan DP, Huang H, Kuntsi J, Langley K, Lesch KP, Leung PWL, Loo SK, Martin J, Martin AR, McGough JJ, Medland SE, Moran JL, Mors O, Mortensen PB, Oades RD, Palmer DS, Pedersen CB, Pedersen MG, Peters T, Poterba T, Poulsen JB, Ramos-Quiroga JA, Reif A, Ribasés M, Rothenberger A, Rovira P, Sánchez-Mora C, Satterstrom FK, Schachar R, Artigas MS, Steinberg S, Stefansson H, Turley P, Walters GB, Werge T, Zayats T, Arking DE, Bettella F, Buxbaum JD, Christensen JH, Collins RL, Coon H, De Rubeis S, Delorme R, Grice DE, Hansen TF, Holmans PA, Hope S, Hultman CM, Klei L, Ladd-Acosta C, Magnusson

P, Nærland T, Nyegaard M, Pinto D, Qvist P, Rehnström K, Reichenberg A, Reichert J, Roeder K, Rouleau GA, Saemundsen E, Sanders SJ, Sandin S, St Pourcain B, Stefansson K, Sutcliffe JS, Talkowski ME, Weiss LA, Willsey AJ, Agartz I, Akil H, Albani D, Alda M, Als TD, Anjorin A, Backlund L, Bass N, Bauer M, Baune BT, Bellivier F, Bergen SE, Berrettini WH, Biernacka JM, Blackwood DHR, Bøen E, Budde M, Bunney W, Burmeister M, Byerley W, Byrne EM, Cichon S, Clarke TK, Coleman JRI, Craddock N, Curtis D, Czerski PM, Dale AM, Dalkner N, Dannlowski U, Degenhardt F, Di Florio A, Elvsåshagen T, Etain B, Fischer SB, Forstner AJ, Forty L, Frank J, Frye M, Fullerton JM, Gade K, Gaspar HA, Gershon ES, Gill M, Goes FS, Gordon SD, Gordon-Smith K, Green MJ, Greenwood TA, Grigoriu-Serbanescu M, Guzman-Parra J, Hauser J, Hautzinger M, Heilbronner U, Herms S, Hoffmann P, Holland D, Jamain S, Jones I, Jones LA, Kandaswamy R, Kelsoe JR, Kennedy JL, Joachim OK, Kittel-Schneider S, Kogevinas M, Koller AC, Lavebratt C, Lewis CM, Li QS, Lissowska J, Loohuis LMO, Lucae S, Maaser A, Malt UF, Martin NG, Martinsson L, McElroy SL, McMahon FJ, McQuillin A, Melle I, Metspalu A, Millischer V, Mitchell PB, Montgomery GW, Morken G, Morris DW, Müller-Myhsok B, Mullins N, Myers RM, Nievergelt CM, Nordentoft M, Adolfsson AN, Nöthen MM, Ophoff RA, Owen MJ, Paciga SA, Pato CN, Pato MT, Perlis RH, Perry A, Potash JB, Reinbold CS, Rietschel M, Rivera M, Roberson M, Schalling M, Schofield PR, Schulze TG, Scott LJ, Serretti A, Sigurdsson E, Smeland OB, Stordal E, Streit F, Strohmaier J, Thorgeirsson TE, Treutlein J, Turecki G, Vaaler AE, Vieta E, Vincent JB, Wang Y, Witt SH, Zandi P, Adan RAH, Alfredsson L, Ando T, Aschauer H, Baker JH, Bencko V, Bergen AW, Birgegård A, Perica VB, Brandt H, Burghardt R, Carlberg L, Cassina M, Clementi M, Courtet P, Crawford S, Crow S, Crowley JJ, Danner UN, Davis OSP, Degortes D, DeSocio JE, Dick DM, Dina C, Docampo E, Egberts K, Ehrlich S, Espeseth T, Fernández-Aranda F, Fichter MM, Foretova L, Forzan M, Gambaro G, Giegling I, Gonidakis F, Gorwood P, Mayora MG, Guo Y, Halmi KA, Hatzikotoulas K, Hebebrand J, Helder SG, Herpertz-Dahlmann B, Herzog W, Hinney A, Imgart H, Jiménez-Murcia S, Johnson C, Jordan J, Julià A, Kaminská D, Karhunen L, Karwautz A, Kas MJH, Kaye WH, Kennedy MA, Kim YR, Klareskog L, Klump KL, Knudsen GPS, Landén M, Le Hellard S, Levitan RD, Li D, Lichtenstein P, Maj M, Marsal S, McDevitt S, Mitchell J, Monteleone P, Monteleone AM, Munn-Chernoff MA, Nacmias B, Navratilova M, O'Toole JK, Padyukov L, Pantel J, Papezova H, Rabionet R, Raevuori A, Ramoz N, Reichborn-Kjennerud T, Ricca V, Roberts M, Rujescu D, Rybakowski F, Scherag A, Schmidt U, Seitz J, Slachtova L, Slof-Op't Landt MCT, Slopian A, Sorbi S, Southam L, Strober M, Tortorella A, Tozzi F, Treasure J, Tziouvas K, van Elburg AA, Wade TD, Wagner G, Walton E, Watson HJ, Wichmann HE, Woodside DB, Zeggini E, Zerwas S, Zipfel S, Adams MJ, Andlauer TFM, Berger K, Binder EB, Boomsma DI, Castela E, Colodro-Conde L, Direk N, Docherty AR, Domenici E, Domschke K, Dunn EC, Foo JC, de Geus EJC, Grabe HJ, Hamilton SP, Horn C, Hottenga JJ, Howard D, Ising M, Kloiber S, Levinson DF, Lewis G, Magnusson PKE, Mbarek H, Middeldorp CM, Mostafavi S, Nyholt DR, Penninx BW, Peterson RE, Pistis G, Porteous DJ, Preisig M, Quiroz JA, Schaefer C, Schulte EC, Shi J, Smith DJ, Thomson PA, Tiemeier H, Uher R, van der Auwera S, Weissman MM, Alexander M, Begemann M, Bramon E, Buccola NG, Cairns MJ, Campion D, Carr VJ, Cloninger CR, Cohen D, Collier DA, Corvin A, DeLisi LE, Donohoe G, Dudbridge F, Duan J, Freedman R, Gejman PV, Golimbet V, Godard S, Ehrenreich H, Hartmann AM, Henskens FA, Ikeda M, Iwata N, Jablensky AV, Joa I, Jönsson EG, Kelly BJ, Knight J, Konte B, Laurent-Levinson C, Lee J, Lencz T, Lerer B, Loughland CM, Malhotra AK, Mallet J, McDonald C, Mitjans M, Mowry BJ, Murphy KC, Murray RM, O'Neill FA, Oh SY, Palotie A, Pantelis C, Pulver AE, Petryshen TL, Quested DJ, Riley B, Sanders AR, Schall U, Schwab SG, Scott RJ, Sham PC, Silverman JM, Sim K, Steixner AA, Tooney PA, van Os J, Vawter MP, Walsh D, Weiser M, Wildenauer DB, Williams NM, Wormley BK, Zhang F, Androustos C, Arnold PD, Barr CL, Barta C, Bey K, Bienvenu OJ, Black DW, Brown LW, Budman C, Cath D, Cheon KA, Ciullo V, Coffey BJ, Cusi D, Davis LK, Denys D, Depienne C, Dietrich A, Eapen V, Falkai P, Fernandez TV, Garcia-Delgar B, Geller DA, Gilbert DL, Grados MA, Greenberg E, Grünblatt E, Hagstrøm J, Hanna GL, Hartmann A, Hedderly T, Heiman GA, Heyman I, Hong HJ, Huang A, Huyser C, Ibanez-Gomez L, Khramtsova EA, Kim YK, Kim YS, King RA, Koh YJ, Konstantinidis A, Kook S, Kuperman S, Leventhal BL, Lochner C, Ludolph AG, Madruga-Garrido M, Malaty I, Maras A, McCracken JT, Meijer IA, Mir P, Morer A,

Müller-Vahl KR, Münchau A, Murphy TL, Naarden A, Nagy P, Nestadt G, Nestadt PS, Nicolini H, Nurmi EL, Okun MS, Paschou P, Piras F, Piras F, Pittenger C, Plessen KJ, Richter MA, Rizzo R, Robertson M, Roessner V, Ruhrmann S, Samuels JF, Sandor P, Schlögelhofer M, Shin EY, Singer H, Song DH, Song J, Spalletta G, Stein DJ, Stewart SE, Storch EA, Stranger B, Stuhmann M, Tarnok Z, Tischfield JA, Tübing J, Visscher F, Vulink N, Wagner M, Walitza S, Wanderer S, Woods M, Worbe Y, Zai G, Zinner SH, Sullivan PF, Franke B, Daly MJ, Bulik CM, Lewis CM, McIntosh AM, O'Donovan MC, Zheutlin A, Andreassen OA, Børglum AD, Breen G, Edenberg HJ, Fanous AH, Faraone SV, Gelernter J, Mathews CA, Mattheisen M, Mitchell KS, Neale MC, Nurnberger JI, Ripke S, Santangelo SL, Scharf JM, Stein MB, Thornton LM, Walters JTR, Wray NR, Geschwind DH, Neale BM, Kendler KS, Smoller JW., 2019. Genomic Relationships, Novel Loci, and Pleiotropic Mechanisms across Eight Psychiatric Disorders. *Cell* 179, 1469-1482.e1411.

Lee, S.H., Ripke, S., Neale, B.M., Faraone, S.V., Purcell, S.M., Perlis, R.H., Mowry, B.J., Thapar, A., Goddard, M.E., Witte, J.S., Absher, D., Agartz, I., Akil, H., Amin, F., Andreassen, O.A., Anjorin, A., Anney, R., Anttila, V., Arking, D.E., Asherson, P., Azevedo, M.H., Backlund, L., Badner, J.A., Bailey, A.J., Banaschewski, T., Barchas, J.D., Barnes, M.R., Barrett, T.B., Bass, N., Battaglia, A., Bauer, M., Bayes, M., Bellivier, F., Bergen, S.E., Berrettini, W., Betancur, C., Bettecken, T., Biederman, J., Binder, E.B., Black, D.W., Blackwood, D.H., Bloss, C.S., Boehnke, M., Boomsma, D.I., Breen, G., Breuer, R., Bruggeman, R., Cormican, P., Buccola, N.G., Buitelaar, J.K., Bunney, W.E., Buxbaum, J.D., Byerley, W.F., Byrne, E.M., Caesar, S., Cahn, W., Cantor, R.M., Casas, M., Chakravarti, A., Chambert, K., Choudhury, K., Cichon, S., Cloninger, C.R., Collier, D.A., Cook, E.H., Coon, H., Cormand, B., Corvin, A., Coryell, W.H., Craig, D.W., Craig, I.W., Crosbie, J., Cuccaro, M.L., Curtis, D., Czamara, D., Datta, S., Dawson, G., Day, R., De Geus, E.J., Degenhardt, F., Djurovic, S., Donohoe, G.J., Doyle, A.E., Duan, J., Dudbridge, F., Duketis, E., Ebstein, R.P., Edenberg, H.J., Elia, J., Ennis, S., Etain, B., Fanous, A., Farmer, A.E., Ferrier, I.N., Flickinger, M., Fombonne, E., Foroud, T., Frank, J., Franke, B., Fraser, C., Freedman, R., Freimer, N.B., Freitag, C.M., Friedl, M., Frisen, L., Gallagher, L., Gejman, P.V., Georgieva, L., Gershon, E.S., Geschwind, D.H., Giegling, I., Gill, M., Gordon, S.D., Gordon-Smith, K., Green, E.K., Greenwood, T.A., Grice, D.E., Gross, M., Grozeva, D., Guan, W., Gurling, H., De Haan, L., Haines, J.L., Hakonarson, H., Hallmayer, J., Hamilton, S.P., Hamshire, M.L., Hansen, T.F., Hartmann, A.M., Hautzinger, M., Heath, A.C., Henders, A.K., Herms, S., Hickie, I.B., Hipolito, M., Hoefels, S., Holmans, P.A., Holsboer, F., Hoogendijk, W.J., Hottenga, J.J., Hultman, C.M., Hus, V., Ingason, A., Ising, M., Jamain, S., Jones, E.G., Jones, I., Jones, L., Tzeng, J.Y., Kahler, A.K., Kahn, R.S., Kandaswamy, R., Keller, M.C., Kennedy, J.L., Kenny, E., Kent, L., Kim, Y., Kirov, G.K., Klauck, S.M., Klei, L., Knowles, J.A., Kohli, M.A., Koller, D.L., Konte, B., Korszun, A., Krabbendam, L., Krasucki, R., Kuntsi, J., Kwan, P., Landen, M., Langstrom, N., Lathrop, M., Lawrence, J., Lawson, W.B., Leboyer, M., Ledbetter, D.H., Lee, P.H., Lencz, T., Lesch, K.P., Levinson, D.F., Lewis, C.M., Li, J., Lichtenstein, P., Lieberman, J.A., Lin, D.Y., Linszen, D.H., Liu, C., Lohoff, F.W., Loo, S.K., Lord, C., Lowe, J.K., Lucae, S., MacIntyre, D.J., Madden, P.A., Maestrini, E., Magnusson, P.K., Mahon, P.B., Maier, W., Malhotra, A.K., Mane, S.M., Martin, C.L., Martin, N.G., Mattheisen, M., Matthews, K., Mattingdsdal, M., McCarroll, S.A., McGhee, K.A., McGough, J.J., McGrath, P.J., McGuffin, P., McInnis, M.G., McIntosh, A., McKinney, R., McLean, A.W., McMahon, F.J., McMahon, W.M., McQuillin, A., Medeiros, H., Medland, S.E., Meier, S., Melle, I., Meng, F., Meyer, J., Middeldorp, C.M., Middleton, L., Milanova, V., Miranda, A., Monaco, A.P., Montgomery, G.W., Moran, J.L., Moreno-De-Luca, D., Morken, G., Morris, D.W., Morrow, E.M., Moskvina, V., Muglia, P., Muhleisen, T.W., Muir, W.J., Muller-Myhsok, B., Murtha, M., Myers, R.M., Myin-Germeys, I., Neale, M.C., Nelson, S.F., Nievergelt, C.M., Nikolov, I., Nimgaonkar, V., Nolen, W.A., Nothen, M.M., Nurnberger, J.I., Nwulia, E.A., Nyholt, D.R., O'Dushlaine, C., Oades, R.D., Olincy, A., Oliveira, G., Olsen, L., Ophoff, R.A., Osby, U., Owen, M.J., Palotie, A., Parr, J.R., Paterson, A.D., Pato, C.N., Pato, M.T., Penninx, B.W., Pergadia, M.L., Pericak-Vance, M.A., Pickard, B.S., Pimm, J., Piven, J., Posthuma, D., Potash, J.B., Poustka, F., Propping, P., Puri, V., Quedsted, D.J., Quinn, E.M., Ramos-Quiroga, J.A., Rasmussen, H.B., Raychaudhuri, S., Rehnstrom, K., Reif, A., Ribases, M., Rice, J.P., Rietschel, M.,

Roeder, K., Roeyers, H., Rossin, L., Rothenberger, A., Rouleau, G., Ruderfer, D., Rujescu, D., Sanders, A.R., Sanders, S.J., Santangelo, S.L., Sergeant, J.A., Schachar, R., Schalling, M., Schatzberg, A.F., Scheftner, W.A., Schellenberg, G.D., Scherer, S.W., Schork, N.J., Schulze, T.G., Schumacher, J., Schwarz, M., Scolnick, E., Scott, L.J., Shi, J., Shilling, P.D., Shyn, S.I., Silverman, J.M., Slager, S.L., Smalley, S.L., Smit, J.H., Smith, E.N., Sonuga-Barke, E.J., St Clair, D., State, M., Steffens, M., Steinhausen, H.C., Strauss, J.S., Strohmaier, J., Stroup, T.S., Sutcliffe, J.S., Szatmari, P., Szelinger, S., Thirumalai, S., Thompson, R.C., Todorov, A.A., Tozzi, F., Treutlein, J., Uhr, M., van den Oord, E.J., Van Grootheest, G., Van Os, J., Vicente, A.M., Vieland, V.J., Vincent, J.B., Visscher, P.M., Walsh, C.A., Wassink, T.H., Watson, S.J., Weissman, M.M., Werge, T., Wienker, T.F., Wijsman, E.M., Willemsen, G., Williams, N., Willsey, A.J., Witt, S.H., Xu, W., Young, A.H., Yu, T.W., Zammit, S., Zandi, P.P., Zhang, P., Zitman, F.G., Zollner, S., Devlin, B., Kelsoe, J.R., Sklar, P., Daly, M.J., O'Donovan, M.C., Craddock, N., Sullivan, P.F., Smoller, J.W., Kendler, K.S., Wray, N.R., 2013.

Genetic relationship between five psychiatric disorders estimated from genome-wide SNPs. *Nat Genet* 45, 984-994.

Lee, S.S., Humphreys, K.L., Flory, K., Liu, R., Glass, K., 2011. Prospective association of childhood attention-deficit/hyperactivity disorder (ADHD) and substance use and abuse/dependence: a meta-analytic review. *Clin Psychol Rev* 31, 328-341.

Lee, Y.C., Yang, H.J., Chen, V.C., Lee, W.T., Teng, M.J., Lin, C.H., Gossop, M., 2016. Meta-analysis of quality of life in children and adolescents with ADHD: By both parent proxy-report and child self-report using PedsQL. *Res Dev Disabil* 51-52, 160-172.

Lenzi, F., Cortese, S., Harris, J., Masi, G., 2018. Pharmacotherapy of emotional dysregulation in adults with ADHD: A systematic review and meta-analysis. *Neuroscience & Biobehavioral Reviews* 84, 359-367.

Leucht, S., Hierl, S., Kissling, W., Dold, M., Davis, J.M., 2012. Putting the efficacy of psychiatric and general medicine medication into perspective: review of meta-analyses. *Br J Psychiatry* 200, 97-106.

Li, J., Olsen, J., Vestergaard, M., Obel, C., 2010. Attention-deficit/hyperactivity disorder in the offspring following prenatal maternal bereavement: a nationwide follow-up study in Denmark. *Eur Child Adolesc Psychiatry* 19, 747-753.

Li, J.J., 2019. The positive end of the polygenic score distribution for ADHD: a low risk or a protective factor? *Psychol Med*, 1-10.

Li, L., Taylor, M.J., Bälter, K., Kuja-Halkola, R., Chen, Q., Hegvik, T.A., Tate, A.E., Chang, Z., Arias-Vásquez, A., Hartman, C.A., Larsson, H., 2020. Attention-deficit/hyperactivity disorder symptoms and dietary habits in adulthood: A large population-based twin study in Sweden. *Am J Med Genet B Neuropsychiatr Genet* 183, 475-485.

Li, X., Sjöstedt, C., Sundquist, J., Zoller, B., Sundquist, K., 2019. Familial association of attention-deficit hyperactivity disorder with autoimmune diseases in the population of Sweden. *Psychiatr Genet* 29, 37-43.

Liang, E.F., Lim, S.Z., Tam, W.W., Ho, C.S., Zhang, M.W., McIntyre, R.S., Ho, R.C., 2018a. The Effect of Methylphenidate and Atomoxetine on Heart Rate and Systolic Blood Pressure in Young People and Adults with Attention-Deficit Hyperactivity Disorder (ADHD): Systematic Review, Meta-Analysis, and Meta-Regression. *Int J Environ Res Public Health* 15, 1789.

- Liang, S.H., Yang, Y.H., Kuo, T.Y., Liao, Y.T., Lin, T.C., Lee, Y., McIntyre, R.S., Kelsen, B.A., Wang, T.N., Chen, V.C., 2018b. Suicide risk reduction in youths with attention-deficit/hyperactivity disorder prescribed methylphenidate: A Taiwan nationwide population-based cohort study. *Res Dev Disabil* 72, 96-105.
- Liao, Y.T., Yang, Y.H., Kuo, T.Y., Liang, H.Y., Huang, K.Y., Wang, T.N., Lee, Y., McIntyre, R.S., Chen, V.C., 2018. Dosage of methylphenidate and traumatic brain injury in ADHD: a population-based study in Taiwan. *Eur Child Adolesc Psychiatry* 27, 279-288.
- Libutzki, B., Ludwig, S., May, M., Jacobsen, R.H., Reif, A., Hartman, C.A., 2019. Direct medical costs of ADHD and its comorbid conditions on basis of a claims data analysis. *Eur Psychiatry* 58, 38-44.
- Libutzki, B., May, M., Gleitz, M., Karus, M., Neukirch, B., Hartman, C.A., Reif, A., 2020. Disease burden and direct medical costs of incident adult ADHD: A retrospective longitudinal analysis based on German statutory health insurance claims data. *Eur Psychiatry* 63, e86.
- Lichtenstein, P., Halldner, L., Zetterqvist, J., Sjolander, A., Serlachius, E., Fazel, S., Langstrom, N., Larsson, H., 2012. Medication for attention deficit-hyperactivity disorder and criminality. *N Engl J Med* 367, 2006-2014.
- Lindstrom, K., Lindblad, F., Hjern, A., 2011. Preterm birth and attention-deficit/hyperactivity disorder in schoolchildren. *Pediatrics* 127, 858-865.
- Liu, H., Feng, W., Zhang, D., 2019a. Association of ADHD medications with the risk of cardiovascular diseases: a meta-analysis. *Eur Child Adolesc Psychiatry* 28, 1283-1293.
- Liu, Q., Zhang, H., Fang, Q., Qin, L., 2017a. Comparative efficacy and safety of methylphenidate and atomoxetine for attention-deficit hyperactivity disorder in children and adolescents: Meta-analysis based on head-to-head trials. *J Clin Exp Neuropsychol* 39, 854-865.
- Liu, X., Dalsgaard, S., Munk-Olsen, T., Li, J., Wright, R.J., Momen, N.C., 2019b. Parental asthma occurrence, exacerbations and risk of attention-deficit/hyperactivity disorder. *Brain Behav Immun* 82, 302-308.
- Liu, Y.S., Dai, X., Wu, W., Yuan, F.F., Gu, X., Chen, J.G., Zhu, L.Q., Wu, J., 2017b. The Association of SNAP25 Gene Polymorphisms in Attention Deficit/Hyperactivity Disorder: a Systematic Review and Meta-Analysis. *Mol Neurobiol* 54, 2189-2200.
- Loyer Carbonneau, M., Demers, M., Bigras, M., Guay, M.C., 2020. Meta-Analysis of Sex Differences in ADHD Symptoms and Associated Cognitive Deficits. *J Atten Disord*, 1087054720923736.
- Lu, Y., Sjölander, A., Cederlöf, M., et al., 2017. Association between medication use and performance on higher education entrance tests in individuals with attention-deficit/hyperactivity disorder. *JAMA Psychiatry* 74, 815-822.

- Lugo, J., Fadeuilhe, C., Gisbert, L., Setien, I., Delgado, M., Corrales, M., Richarte, V., Ramos-Quiroga, J.A., 2020. Sleep in adults with autism spectrum disorder and attention deficit/hyperactivity disorder: A systematic review and meta-analysis *Eur Neuropsychopharmacol* 38, 1-24.
- Lukito, S., Norman, L., Carlisi, C., Radua, J., Hart, H., Simonoff, E., Rubia, K., 2020. Comparative meta-analyses of brain structural and functional abnormalities during cognitive control in attention-deficit/hyperactivity disorder and autism spectrum disorder. *Psychol Med* 50, 894-919.
- Maher, G.M., Dalman, C., O'Keeffe, G.W., Kearney, P.M., McCarthy, F.P., Kenny, L.C., Khashan, A.S., 2020. Association between preeclampsia and attention-deficit hyperactivity disorder: a population-based and sibling-matched cohort study. *Acta Psychiatr Scand*.
- Maher, G.M., O'Keeffe, G.W., Kearney, P.M., Kenny, L.C., Dinan, T.G., Mattsson, M., Khashan, A.S., 2018. Association of Hypertensive Disorders of Pregnancy With Risk of Neurodevelopmental Disorders in Offspring: A Systematic Review and Meta-analysis. *JAMA Psychiatry* 75, 809-819.
- Man, K.K., Chan, E.W., Coghill, D., Douglas, I., Ip, P., Leung, L.P., Tsui, M.S., Wong, W.H., Wong, I.C., 2015. Methylphenidate and the risk of trauma. *Pediatrics* 135, 40-48.
- Man, K.K., Coghill, D., Chan, E.W., Lau, W.C., Hollis, C., Liddle, E., Banaschewski, T., McCarthy, S., Neubert, A., Sayal, K., Ip, P., Wong, I.C., 2016. Methylphenidate and the risk of psychotic disorders and hallucinations in children and adolescents in a large health system. *Transl Psychiatry* 6, e956.
- Man, K.K.C., Coghill, D., Chan, E.W., Lau, W.C.Y., Hollis, C., Liddle, E., Banaschewski, T., McCarthy, S., Neubert, A., Sayal, K., Ip, P., Schuemie, M.J., Sturkenboom, M., Sonuga-Barke, E., Buitelaar, J., Carucci, S., Zuddas, A., Kovshoff, H., Garas, P., Nagy, P., Inglis, S.K., Konrad, K., Hage, A., Rosenthal, E., Wong, I.C.K., 2017. Association of Risk of Suicide Attempts With Methylphenidate Treatment. *JAMA Psychiatry* 74, 1048-1055.
- Maneeton, N., Maneeton, B., Woottitluk, P., Suttajit, S., Likhitsathian, S., Charmsil, C., Srisurapanont, M., 2015. Comparative efficacy, acceptability, and tolerability of dexamethylphenidate versus placebo in child and adolescent ADHD: a meta-analysis of randomized controlled trials. *Neuropsychiatr Dis Treat* 11, 2943-2952.
- Martin, J., Taylor, M.J., Rydell, M., Riglin, L., Eyre, O., Lu, Y., Lundstrom, S., Larsson, H., Thapar, A., Lichtenstein, P., 2018. Sex-specific manifestation of genetic risk for attention deficit hyperactivity disorder in the general population. *J Child Psychol Psychiatry* 59, 908-916.
- Martinez-Badia, J., Martinez-Raga, J., 2015. Who says this is a modern disorder? The early history of attention deficit hyperactivity disorder. *World J Psychiatry* 5, 379-386.
- Marx, I., Hacker, T., Yu, X., Cortese, S., Sonuga-Barke, E., 2018. ADHD and the Choice of Small Immediate Over Larger Delayed Rewards: A Comparative Meta-Analysis of Performance on Simple Choice-Delay and Temporal Discounting Paradigms. *J Atten Disord*, 1087054718772138.
- McCabe, S.E., Veliz, P., Wilens, T.E., Schulenberg, J.E., 2017. Adolescents' Prescription Stimulant Use and Adult Functional Outcomes: A National Prospective Study. *J Am Acad Child Adolesc Psychiatry* 56, 226-233.e224.

McCarthy, S., Neubert, A., Man, K.K.C., Banaschewski, T., Buitelaar, J., Carucci, S., Coghill, D., Danckaerts, M., Falissard, B., Garas, P., Hage, A., Hollis, C., Inglis, S., Kovshoff, H., Liddle, E., Mechler, K., Nagy, P., Rosenthal, E., Schlack, R., Sonuga-Barke, E., Zuddas, A., Wong, I.C.K., 2018. Effects of long-term methylphenidate use on growth and blood pressure: results of the German Health Interview and Examination Survey for Children and Adolescents (KiGGS). *BMC Psychiatry* 18, 327.

McCauley, H.L., Breslau, J.A., Saito, N., Miller, E., 2015. Psychiatric disorders prior to dating initiation and physical dating violence before age 21: findings from the National Comorbidity Survey Replication (NCS-R). *Soc Psychiatry Psychiatr Epidemiol* 50, 1357-1365.

McGough, J.J., Sturm, A., Cowen, J., Tung, K., Salgari, G.C., Leuchter, A.F., Cook, I.A., Sugar, C.A., Loo, S.K., 2019. Double-Blind, Sham-Controlled, Pilot Study of Trigeminal Nerve Stimulation for Attention-Deficit/Hyperactivity Disorder. *J Am Acad Child Adolesc Psychiatry* 58, 403-411.e403.

McLeod, J.D., Fettes, D.L., Jensen, P.S., Pescosolido, B.A., Martin, J.K., 2007. Public knowledge, beliefs, and treatment preferences concerning attention-deficit hyperactivity disorder. *Psychiatr Serv* 58, 626-631.

Melby-Lervag, M., Hulme, C., 2013. Is working memory training effective? A meta-analytic review. *Dev Psychol* 49, 270-291.

Micoulaud-Franchi, J.A., Geoffroy, P.A., Fond, G., Lopez, R., Bioulac, S., Philip, P., 2014. EEG neurofeedback treatments in children with ADHD: an updated meta-analysis of randomized controlled trials. *Front Hum Neurosci* 8, 906.

Mohr-Jensen, C., Muller Bisgaard, C., Boldsen, S.K., Steinhausen, H.C., 2019. Attention-Deficit/Hyperactivity Disorder in Childhood and Adolescence and the Risk of Crime in Young Adulthood in a Danish Nationwide Study. *J Am Acad Child Adolesc Psychiatry* 58, 443-452.

Momany, A.M., Kamradt, J.M., Nikolas, M.A., 2018. A Meta-Analysis of the Association Between Birth Weight and Attention Deficit Hyperactivity Disorder. *J Abnorm Child Psychol* 46, 1409-1426.

Montes, G., Halterman, J.S., 2007. Bullying among children with autism and the influence of comorbidity with ADHD: a population-based study. *Ambul Pediatr* 7, 253-257.

Morris, H.H., Escoll, P.J., Wexler, R., 1956. Aggressive Behavior Disorders of Childhood: A Follow-Up Study. *Am J Psychiatry* 112, 991-997.

Mueller, A.K., Fuermaier, A.B., Koerts, J., Tucha, L., 2012. Stigma in attention deficit hyperactivity disorder. *Atten Defic Hyperact Disord* 4, 101-114.

National Collaborating Centre for Mental Health, 2018. Attention Deficit Hyperactivity Disorder: Diagnosis and Management of ADHD in Children, Young People and Adults. British Psychological Society

Copyright (c) National Institute for Health and Care Excellence 2018., Leicester (UK).

National Institute for Health Care and Excellence, 2018. Attention deficit hyperactivity disorder: diagnosis and management, March 14, 2018 ed. National Institute for Health Care and Excellence, United Kingdom.

Nazar, B.P., Bernardes, C., Peachey, G., Sergeant, J., Mattos, P., Treasure, J., 2016. The risk of eating disorders comorbid with attention-deficit/hyperactivity disorder: A systematic review and meta-analysis. *Int J Eat Disord* 49, 1045-1057.

Nelson, L.D., Guskiewicz, K.M., Marshall, S.W., Hammeke, T., Barr, W., Randolph, C., McCrea, M.A., 2016. Multiple Self-Reported Concussions Are More Prevalent in Athletes With ADHD and Learning Disability. *Clin J Sport Med* 26, 120-127.

Neumarker, K.J., 2005. The Kramer-Pollnow syndrome: a contribution on the life and work of Franz Kramer and Hans Pollnow. *Hist Psychiatry* 16, 435-451.

Nielsen, P.R., Benros, M.E., Dalsgaard, S., 2017. Associations Between Autoimmune Diseases and Attention-Deficit/Hyperactivity Disorder: A Nationwide Study. *J Am Acad Child Adolesc Psychiatry* 56, 234-240.e231.

Nigg, J.T., Johnstone, J.M., Musser, E.D., Long, H.G., Willoughby, M.T., Shannon, J., 2016. Attention-deficit/hyperactivity disorder (ADHD) and being overweight/obesity: New data and meta-analysis. *Clin Psychol Rev* 43, 67-79.

Nigg, J.T., Lewis, K., Edinger, T., Falk, M., 2012. Meta-analysis of attention-deficit/hyperactivity disorder or attention-deficit/hyperactivity disorder symptoms, restriction diet, and synthetic food color additives. *J Am Acad Child Adolesc Psychiatry* 51, 86-97 e88.

Nilsen, F.M., Tulve, N.S., 2020. A systematic review and meta-analysis examining the interrelationships between chemical and non-chemical stressors and inherent characteristics in children with ADHD. *Environ Res* 180, 108884.

Norman, L.J., Carlisi, C., Lukito, S., Hart, H., Mataix-Cols, D., Radua, J., Rubia, K., 2016. Structural and Functional Brain Abnormalities in Attention-Deficit/Hyperactivity Disorder and Obsessive-Compulsive Disorder: A Comparative Meta-analysis. *JAMA Psychiatry* 73, 815-825.

O'Neal, P., Robins, L.N., 1958. Childhood patterns predictive of adult schizophrenia: a 30-year follow-up study. *Am J Psychiatry* 115, 385-391.

Obel, C., Zhu, J.L., Olsen, J., Breining, S., Li, J., Gronborg, T.K., Gissler, M., Rutter, M., 2016. The risk of attention deficit hyperactivity disorder in children exposed to maternal smoking during pregnancy - a re-examination using a sibling design. *J Child Psychol Psychiatry* 57, 532-537.

Ostergaard, S.D., Dalsgaard, S., Faraone, S.V., Munk-Olsen, T., Laursen, T.M., 2017. Teenage Parenthood and Birth Rates for Individuals With and Without Attention-Deficit/Hyperactivity Disorder: A Nationwide Cohort Study. *J Am Acad Child Adolesc Psychiatry* 56, 578-584 e573.

Ostergaard, S.D., Larsen, J.T., Dalsgaard, S., Wilens, T.E., Mortensen, P.B., Agerbo, E., Mors, O., Petersen, L., 2016. Predicting ADHD by Assessment of Rutter's Indicators of Adversity in Infancy. *PLoS One* 11, e0157352.

Ouyang, L., Fang, X., Mercy, J., Perou, R., Grosse, S.D., 2008. Attention-deficit/hyperactivity disorder symptoms and child maltreatment: a population-based study. *J Pediatr* 153, 851-856.

Palmer, E.D., Finger, S., 2001. An Early Description of ADHD (Inattentive Subtype): Dr Alexander Crichton and Mental Restlessness (1798). *Child Psychology and Psychiatry Review* 6, 66-73.

Pan, Y.Q., Qiao, L., Xue, X.D., Fu, J.H., 2015. Association between ANKK1 (rs1800497) polymorphism of DRD2 gene and attention deficit hyperactivity disorder: A meta-analysis. *Neurosci Lett* 590, 101-105.

Park, J., Sohn, J.H., Cho, S.J., Seo, H.Y., Hwang, I.U., Hong, Y.C., Kim, K.N., 2020. Association between short-term air pollution exposure and attention-deficit/hyperactivity disorder-related hospital admissions among adolescents: A nationwide time-series study. *Environ Pollut* 266, 115369.

Patros, C.H., Alderson, R.M., Kasper, L.J., Tarle, S.J., Lea, S.E., Hudec, K.L., 2016. Choice-impulsivity in children and adolescents with attention-deficit/hyperactivity disorder (ADHD): A meta-analytic review. *Clin Psychol Rev* 43, 162-174.

Patros, C.H.G., Tarle, S.J., Alderson, R.M., Lea, S.E., Arrington, E.F., 2019. Planning deficits in children with attention-deficit/hyperactivity disorder (ADHD): A meta-analytic review of tower task performance. *Neuropsychology* 33, 425-444.

Pauli-Pott, U., Mann, C., Becker, K., 2020. Do cognitive interventions for preschoolers improve executive functions and reduce ADHD and externalizing symptoms? A meta-analysis of randomized controlled trials. *Eur Child Adolesc Psychiatry*.

Pearl, P.L., Weiss, R.E., Stein, M.A., 2001. Medical mimics. Medical and neurological conditions simulating ADHD. *Ann N Y Acad Sci* 931, 97-112.

Pettersson, E., Lichtenstein, P., Larsson, H., Song, J., Agrawal, A., Borglum, A.D., Bulik, C.M., Daly, M.J., Davis, L.K., Demontis, D., Edenberg, H.J., Grove, J., Gelernter, J., Neale, B.M., Pardinas, A.F., Stahl, E., Walters, J.T.R., Walters, R., Sullivan, P.F., Posthuma, D., Polderman, T.J.C., 2019. Genetic influences on eight psychiatric disorders based on family data of 4 408 646 full and half-siblings, and genetic data of 333 748 cases and controls. *Psychol Med* 49, 1166-1173.

Pievsky, M.A., McGrath, R.E., 2018. The Neurocognitive Profile of Attention-Deficit/Hyperactivity Disorder: A Review of Meta-Analyses. *Arch Clin Neuropsychol* 33, 143-157.

Pliszka, S., 2007. Practice parameter for the assessment and treatment of children and adolescents with attention-deficit/hyperactivity disorder. *J Am Acad Child Adolesc Psychiatry* 46, 894-921.

Pohlabeln, H., Rach, S., De Henauw, S., Eiben, G., Gwozdz, W., Hadjigeorgiou, C., Molnar, D., Moreno, L.A., Russo, P., Veidebaum, T., Pigeot, I., 2017. Further evidence for the role of pregnancy-induced hypertension and other early life influences in the development of ADHD: results from the IDEFICS study. *Eur Child Adolesc Psychiatry* 26, 957-967.

- Polanczyk, G.V., Willcutt, E.G., Salum, G.A., Kieling, C., Rohde, L.A., 2014. ADHD prevalence estimates across three decades: an updated systematic review and meta-regression analysis. *Int J Epidemiol* 43, 434-442.
- Pozzi, M., Carnovale, C., Peeters, G., Gentili, M., Antoniazzi, S., Radice, S., Clementi, E., Nobile, M., 2018. Adverse drug events related to mood and emotion in paediatric patients treated for ADHD: A meta-analysis. *J Affect Disord* 238, 161-178.
- Pringsheim, T., Hirsch, L., Gardner, D., Gorman, D.A., 2015. The pharmacological management of oppositional behaviour, conduct problems, and aggression in children and adolescents with attention-deficit hyperactivity disorder, oppositional defiant disorder, and conduct disorder: a systematic review and meta-analysis. Part 1: psychostimulants, alpha-2 agonists, and atomoxetine. *Can J Psychiatry* 60, 42-51.
- Puri, B.K., Martins, J.G., 2014. Which polyunsaturated fatty acids are active in children with attention-deficit hyperactivity disorder receiving PUFA supplementation? A fatty acid validated meta-regression analysis of randomized controlled trials. *Prostaglandins Leukot Essent Fatty Acids* 90, 179-189.
- Ramos, A.A., Hamdan, A.C., Machado, L., 2020. A meta-analysis on verbal working memory in children and adolescents with ADHD. *Clin Neuropsychol* 34, 873-898.
- Rimestad, M.L., Lambek, R., Zacher Christiansen, H., Hougaard, E., 2019. Short- and Long-Term Effects of Parent Training for Preschool Children With or at Risk of ADHD: A Systematic Review and Meta-Analysis. *J Atten Disord* 23, 423-434.
- Robins, E., Guze, S.B., 1970. Establishment of diagnostic validity in psychiatric illness: its application to schizophrenia. *Am J Psychiatry* 126, 983-987.
- Rommelse, N., Antshel, K., Smeets, S., Greven, C., Hoogveen, L., Faraone, S.V., Hartman, C.A., 2017. High intelligence and the risk of ADHD and other psychopathology. *Br J Psychiatry* 211, 359-364.
- Ros, R., Graziano, P.A., 2018. Social Functioning in Children With or At Risk for Attention Deficit/Hyperactivity Disorder: A Meta-Analytic Review. *J Clin Child Adolesc Psychol* 47, 213-235.
- Rosenthal, R., Rosnow, R.L., 1984. *Essentials of Behavioral Research: Methods and Data Analysis*. 361.
- Rubia, K., Alegria, A.A., Cubillo, A.I., Smith, A.B., Brammer, M.J., Radua, J., 2014. Effects of stimulants on brain function in attention-deficit/hyperactivity disorder: a systematic review and meta-analysis. *Biol Psychiatry* 76, 616-628.
- Ruiz-Goikoetxea, M., Cortese, S., Aznarez-Sanado, M., Magallon, S., Alvarez Zallo, N., Luis, E.O., de Castro-Manglano, P., Soutullo, C., Arrondo, G., 2018a. Risk of unintentional injuries in children and adolescents with ADHD and the impact of ADHD medications: A systematic review and meta-analysis. *Neurosci Biobehav Rev* 84, 63-71.

Ruiz-Goikoetxea, M., Cortese, S., Magallon, S., Aznarez-Sanado, M., Alvarez Zallo, N., Luis, E.O., de Castro-Manglano, P., Soutullo, C., Arrondo, G., 2018b. Risk of poisoning in children and adolescents with ADHD: a systematic review and meta-analysis. *Sci Rep* 8, 7584.

Rydell, M., Lundstrom, S., Gillberg, C., Lichtenstein, P., Larsson, H., 2018. Has the attention deficit hyperactivity disorder phenotype become more common in children between 2004 and 2014? Trends over 10 years from a Swedish general population sample. *J Child Psychol Psychiatry* 59, 863-871.

Samea, F., Soluki, S., Nejati, V., Zarei, M., Cortese, S., Eickhoff, S.B., Tahmasian, M., Eickhoff, C.R., 2019. Brain alterations in children/adolescents with ADHD revisited: a neuroimaging meta-analysis of 96 structural and functional studies. *Neurosci Biobehav Rev*.

Sanchez, C., Barry, C., Sabhlok, A., Russell, K., Majors, A., Kollins, S., Fuemmeler, B., 2018. Maternal pre-pregnancy obesity and child neurodevelopmental outcomes: a meta-analysis. *Obesity Reviews* 19, 464-484.

Satterstrom, F.K., Walters, R.K., Singh, T., Wigdor, E.M., Lescai, F., Demontis, D., Kosmicki, J.A., Grove, J., Stevens, C., Bybjerg-Grauholm, J., Baekvad-Hansen, M., Palmer, D.S., Maller, J.B., Nordentoft, M., Mors, O., Robinson, E.B., Hougaard, D.M., Werge, T.M., Bo Mortensen, P., Neale, B.M., Borglum, A.D., Daly, M.J., 2019. Autism spectrum disorder and attention deficit hyperactivity disorder have a similar burden of rare protein-truncating variants. *Nat Neurosci* 22, 1961-1965.

Schab, D.W., Trinh, N.H., 2004. Do artificial food colors promote hyperactivity in children with hyperactive syndromes? A meta-analysis of double-blind placebo-controlled trials. *J Dev Behav Pediatr* 25, 423-434.

Schoechlin, C., Engel, R.R., 2005. Neuropsychological performance in adult attention-deficit hyperactivity disorder: meta-analysis of empirical data. *Arch Clin Neuropsychol* 20, 727-744.

Schoeman, R., Liebenberg, R., 2017. The South African Society of Psychiatrists/Psychiatry Management Group management guidelines for adult attention-deficit/hyperactivity disorder. *The South African journal of psychiatry : SAJP : the journal of the Society of Psychiatrists of South Africa* 23, 1060-1060.

Schoenfelder, E.N., Faraone, S.V., Kollins, S.H., 2014. Stimulant treatment of ADHD and cigarette smoking: a meta-analysis. *Pediatrics* 133, 1070-1080.

Schwartz, S., Correll, C.U., 2014. Efficacy and safety of atomoxetine in children and adolescents with attention-deficit/hyperactivity disorder: results from a comprehensive meta-analysis and metaregression. *J Am Acad Child Adolesc Psychiatry* 53, 174-187.

Scionti, N., Cavallero, M., Zogmaister, C., Marzocchi, G.M., 2019. Is Cognitive Training Effective for Improving Executive Functions in Preschoolers? A Systematic Review and Meta-Analysis. *Front Psychol* 10, 2812.

Sedky, K., Bennett, D.S., Carvalho, K.S., 2014. Attention deficit hyperactivity disorder and sleep disordered breathing in pediatric populations: A meta-analysis. *Sleep Med Rev* 18, 349-356.

Seixas, M., Weiss, M., Muller, U., 2012. Systematic review of national and international guidelines on attention-deficit hyperactivity disorder. *J Psychopharmacol* 26, 753-765.

Septier, M., Stordeur, C., Zhang, J., Delorme, R., Cortese, S., 2019. Association between suicidal spectrum behaviors and Attention-Deficit/Hyperactivity Disorder: A systematic review and meta-analysis. *Neurosci Biobehav Rev* 103, 109-118.

Shih, P., Huang, C.C., Pan, S.C., Chiang, T.L., Guo, Y.L., 2020. Hyperactivity disorder in children related to traffic-based air pollution during pregnancy. *Environ Res* 188, 109588.

Simon, V., Czobor, P., Balint, S., Meszaros, A., Bitter, I., 2009. Prevalence and correlates of adult attention-deficit hyperactivity disorder: meta-analysis. *Br J Psychiatry* 194, 204-211.

Skoglund, C., Chen, Q., D'Onofrio, B.M., Lichtenstein, P., Larsson, H., 2014. Familial confounding of the association between maternal smoking during pregnancy and ADHD in offspring. *J Child Psychol Psychiatry* 55, 61-68.

Skoglund, C., Kopp Kallner, H., Skalkidou, A., Wikstrom, A.K., Lundin, C., Hesselman, S., Wikman, A., Sundstrom Poromaa, I., 2019. Association of Attention-Deficit/Hyperactivity Disorder With Teenage Birth Among Women and Girls in Sweden. *JAMA Netw Open* 2, e1912463.

Solberg, B.S., Halmoy, A., Engeland, A., Igland, J., Haavik, J., Klungsoyr, K., 2018. Gender differences in psychiatric comorbidity: a population-based study of 40 000 adults with attention deficit hyperactivity disorder. *Acta Psychiatr Scand* 137, 176-186.

Solmi, M., Fornaro, M., Ostinelli, E.G., Zangani, C., Croatto, G., Monaco, F., Krinitski, D., Fusar-Poli, P., Correll, C.U., 2020. Safety of 80 antidepressants, antipsychotics, anti-attention-deficit/hyperactivity medications and mood stabilizers in children and adolescents with psychiatric disorders: a large scale systematic meta-review of 78 adverse effects. *World Psychiatry* 19, 214-232.

Song, M., Dieckmann, N.F., Nigg, J.T., 2019. Addressing Discrepancies Between ADHD Prevalence and Case Identification Estimates Among U.S. Children Utilizing NSCH 2007-2012. *J Atten Disord* 23, 1691-1702.

Spencer, T.J., Brown, A., Seidman, L.J., Valera, E.M., Makris, N., Lomedico, A., Faraone, S.V., Biederman, J., 2013. Effect of psychostimulants on brain structure and function in ADHD: a qualitative literature review of magnetic resonance imaging-based neuroimaging studies. *J Clin Psychiatry* 74, 902-917.

Stein, M.A., 2008. Medical mimics and differential diagnosis in adult ADHD. *CNS Spectr* 13, 14-16.

Still, G., 1902a. The Goulstonian lectures on some abnormal physical conditions in children. Lecture 1. *Lancet*, 1008-10102, 1077-1082, 1163-1168.

Still, G., 1902b. The Goulstonian lectures on some abnormal psychical conditions in children. Lecture II. *Lancet* 1, 1077-1082.

Still, G., 1902c. The Goulstonian lectures on some abnormal psychical conditions in children. Lecture III. *Lancet* 1, 1163-1168.

Stojanovski, S., Felsky, D., Viviano, J.D., Shahab, S., Bangali, R., Burton, C.L., Devenyi, G.A., O'Donnell, L.J., Szatmari, P., Chakravarty, M.M., Ameis, S., Schachar, R., Voineskos, A.N., Wheeler, A.L., 2019. Polygenic Risk and Neural Substrates of Attention-Deficit/Hyperactivity Disorder Symptoms in Youths With a History of Mild Traumatic Brain Injury. *Biol Psychiatry* 85, 408-416.

Storebo, O.J., Elmoose Andersen, M., Skoog, M., Joost Hansen, S., Simonsen, E., Pedersen, N., Tendal, B., Callesen, H.E., Faltinsen, E., Gluud, C., 2019. Social skills training for attention deficit hyperactivity disorder (ADHD) in children aged 5 to 18 years. *Cochrane Database Syst Rev* 6, Cd008223.

Storebø, O.J., Ramstad, E., Krogh, H.B., Nilausen, T.D., Skoog, M., Holmskov, M., Rosendal, S., Groth, C., Magnusson, F.L., Moreira-Maia, C.R., Gillies, D., Buch Rasmussen, K., Gauci, D., Zwi, M., Kirubakaran, R., Forsbøl, B., Simonsen, E., Gluud, C., 2015. Methylphenidate for children and adolescents with attention deficit hyperactivity disorder (ADHD). *Cochrane Database Syst Rev*, Cd009885.

Strine, T.W., Lesesne, C.A., Okoro, C.A., McGuire, L.C., Chapman, D.P., Balluz, L.S., Mokdad, A.H., 2006. Emotional and behavioral difficulties and impairments in everyday functioning among children with a history of attention-deficit/hyperactivity disorder. *Prev Chronic Dis* 3, A52.

Su, C.C., Tsai, C.Y., Tsai, T.H., Tsai, I.J., 2019. Incidence and risk of attention-deficit hyperactivity disorder in children with amblyopia: A nationwide cohort study. *Clin Exp Ophthalmol* 47, 259-264.

Sucksdorff, M., Brown, A.S., Chudal, R., Surcel, H.M., Hinkka-Yli-Salomaki, S., Cheslack-Postava, K., Gyllenberg, D., Sourander, A., 2019. Maternal Vitamin D Levels and the Risk of Offspring Attention-Deficit/Hyperactivity Disorder. *J Am Acad Child Adolesc Psychiatry* S0890-8567(19)32232-4.

Sucksdorff, M., Lehtonen, L., Chudal, R., Suominen, A., Joelsson, P., Gissler, M., Sourander, A., 2015. Preterm Birth and Poor Fetal Growth as Risk Factors of Attention-Deficit/ Hyperactivity Disorder. *Pediatrics* 136, e599-608.

Sun, C.K., Tseng, P.T., Wu, C.K., Li, D.J., Chen, T.Y., Stubbs, B., Carvalho, A.F., Chen, Y.W., Lin, P.Y., Cheng, Y.S., Wu, M.K., 2019a. Therapeutic effects of methylphenidate for attention-deficit/hyperactivity disorder in children with borderline intellectual functioning or intellectual disability: A systematic review and meta-analysis. *Sci Rep* 9, 15908.

Sun, S., Kuja-Halkola, R., Faraone, S.V., D'Onofrio, B.M., Dalsgaard, S., Chang, Z., Larsson, H., 2019b. Association of Psychiatric Comorbidity With the Risk of Premature Death Among Children and Adults With Attention-Deficit/Hyperactivity Disorder. *JAMA Psychiatry* 76, 1141-1149.

Sundquist, J., Ohlsson, H., Sundquist, K., Kendler, K.S., 2015. Attention-deficit/hyperactivity disorder and risk for drug use disorder: a population-based follow-up and co-relative study. *Psychol Med* 45, 977-983.

Sweeney, C.T., Sembower, M.A., Ertischek, M.D., Shiffman, S., Schnoll, S.H., 2013. Nonmedical use of prescription ADHD stimulants and preexisting patterns of drug abuse. *J Addict Dis* 32, 1-10.

Swensen, A.R., Birnbaum, H.G., Secnik, K., Marynchenko, M., Greenberg, P., Claxton, A., 2003. Attention-deficit/hyperactivity disorder: increased costs for patients and their families. *J Am Acad Child Adolesc Psychiatry* 42, 1415-1423.

- Tamminga, H.G., Reneman, L., Huizenga, H.M., Geurts, H.M., 2016. Effects of methylphenidate on executive functioning in attention-deficit/hyperactivity disorder across the lifespan: a meta-regression analysis. *Psychol Med* 46, 1791-1807.
- Taylor, E., 2011. Antecedents of ADHD: a historical account of diagnostic concepts. *Atten Defic Hyperact Disord* 3, 69-75.
- Taylor, E., Dopfner, M., Sergeant, J., Asherson, P., Banaschewski, T., Buitelaar, J., Coghill, D., Danckaerts, M., Rothenberger, A., Sonuga-Barke, E., Steinhausen, H.C., Zuddas, A., 2004. European clinical guidelines for hyperkinetic disorder-first upgrade. *Eur Child Adolesc Psychiatry* 13, i7-i30.
- Taylor, M.J., Martin, J., Lu, Y., Brikell, I., Lundstrom, S., Larsson, H., Lichtenstein, P., 2019. Association of Genetic Risk Factors for Psychiatric Disorders and Traits of These Disorders in a Swedish Population Twin Sample. *JAMA Psychiatry* 76, 280-289.
- Thome, J., Ehli, A.C., Fallgatter, A.J., Krauel, K., Lange, K.W., Riederer, P., Romanos, M., Taurines, R., Tucha, O., Uzbekov, M., Gerlach, M., 2012. Biomarkers for attention-deficit/hyperactivity disorder (ADHD). A consensus report of the WFSBP task force on biological markers and the World Federation of ADHD. *World J Biol Psychiatry* 13, 379-400.
- Tseng, J.J., Lin, C.H., Lin, M.C., 2020. Long-Term Outcomes of Pediatric Enterovirus Infection in Taiwan: A Population-Based Cohort Study. *Front Pediatr* 8, 285.
- Tseng, P.T., Cheng, Y.S., Yen, C.F., Chen, Y.W., Stubbs, B., Whiteley, P., Carvalho, A.F., Li, D.J., Chen, T.Y., Yang, W.C., Tang, C.H., Chu, C.S., Yang, W.C., Liang, H.Y., Wu, C.K., Lin, P.Y., 2018. Peripheral iron levels in children with attention-deficit hyperactivity disorder: a systematic review and meta-analysis. *Sci Rep* 8, 788.
- Tsuji, N., Okada, T., Usami, M., Kuwabara, H., Fujita, J., Negoro, H., Kawamura, M., Iida, J., Saito, T., 2020. Effect of Continuing and Discontinuing Medications on Quality of Life After Symptomatic Remission in Attention-Deficit/Hyperactivity Disorder: A Systematic Review and Meta-Analysis. *J Clin Psychiatry* 81, 19r13015.
- Tung, I., Li, J.J., Meza, J.I., Jezior, K.L., Kianmahd, J.S., Hentschel, P.G., O'Neil, P.M., Lee, S.S., 2016. Patterns of Comorbidity Among Girls With ADHD: A Meta-analysis. *Pediatrics* 138, e20160430.
- Tylee, D.S., Sun, J., Hess, J.L., Tahir, M.A., Sharma, E., Malik, R., Worrall, B.B., Levine, A.J., Martinson, J.J., Nejentsev, S., Speed, D., Fischer, A., Mick, E., Walker, B.R., Crawford, A., Grant, S.F.A., Polychronakos, C., Bradfield, J.P., Sleiman, P.M.A., Hakonarson, H., Ellinghaus, E., Elder, J.T., Tsoi, L.C., Trembath, R.C., Barker, J.N., Franke, A., Dehghan, A., Team, a.M.R., Consortium, I.W.G.o.t.C., Consortium, M.C.o.t.I.S.G., Registry, N.T., Group, n.W., Consortium, O.C.a.T.S.W.G.o.t.P.G., Faraone, S.V., Glatt, S.J., 2018. Genetic correlations among psychiatric and immune-related phenotypes based on genome-wide association data. *Am J Med Genet B Neuropsychiatr Genet* 177, 641-657.
- Tzeng, N.S., Chung, C.H., Lin, F.H., Yeh, C.B., Huang, S.Y., Lu, R.B., Chang, H.A., Kao, Y.C., Yeh, H.W., Chiang, W.S., Chou, Y.C., Tsao, C.H., Wu, Y.F., Chien, W.C., 2019. Risk of Dementia in Adults With ADHD: A Nationwide, Population-Based Cohort Study in Taiwan. *J Atten Disord* 23, 995-1006.

Vaa, T., 2014. ADHD and relative risk of accidents in road traffic: a meta-analysis. *Accid Anal Prev* 62, 415-425.

van der Schans, J., Aikman, B., de Vries, T.W., Hoekstra, P.J., Hak, E., 2017. Association Between Attention-Deficit/Hyperactivity Disorder and Asthma Among Adults: A Case-Control Study. *Chest* 151, 1406-1407.

Van Doren, J., Arns, M., Heinrich, H., Vollebregt, M.A., Strehl, U., S, K.L., 2019. Sustained effects of neurofeedback in ADHD: a systematic review and meta-analysis. *Eur Child Adolesc Psychiatry* 28, 293-305.

van Hulzen, K.J.E., Scholz, C.J., Franke, B., Ripke, S., Klein, M., McQuillin, A., Sonuga-Barke, E.J., Group, P.A.W., Kelsoe, J.R., Landen, M., Andreassen, O.A., Group, P.G.C.B.D.W., Lesch, K.P., Weber, H., Faraone, S.V., Arias-Vasquez, A., Reif, A., 2017. Genetic Overlap Between Attention-Deficit/Hyperactivity Disorder and Bipolar Disorder: Evidence From Genome-wide Association Study Meta-analysis. *Biol Psychiatry* 82, 634-641.

Vidal Perera, A., 1907. *Compendio de psiquiatría infantil* 1st ed. Librería del Magisterio, Barcelona.

Vink, J.M., Schellekens, A., 2018. Relating addiction and psychiatric disorders. *Science* 361, 1323-1324.

Vysniauske, R., Verburgh, L., Oosterlaan, J., Molendijk, M.L., 2020. The Effects of Physical Exercise on Functional Outcomes in the Treatment of ADHD: A Meta-Analysis. *J Atten Disord* 24, 644-654.

Wang, H., Li, F., Miao, M., Yu, Y., Ji, H., Liu, H., Huang, R., Obel, C., Zhang, J., Li, J., 2020. Maternal spontaneous abortion and the risk of attention-deficit/hyperactivity disorder in offspring: a population-based cohort study. *Hum Reprod* 35, 1211-1221.

Wang, L.J., Lee, S.Y., Chou, W.J., Lee, M.J., Tsai, C.S., Lee, T.L., Yang, C.J., Yang, K.C., Chen, C.K., Shyu, Y.C., 2019. Testicular Function After Long-Term Methylphenidate Treatment in Boys with Attention-Deficit/Hyperactivity Disorder. *J Child Adolesc Psychopharmacol* 29, 433-438.

Wang, Y., Huang, L., Zhang, L., Qu, Y., Mu, D., 2017. Iron Status in Attention-Deficit/Hyperactivity Disorder: A Systematic Review and Meta-Analysis. *PLoS One* 12, e0169145.

Weikard, M.A., 1799. *Der philosophische Arzt. 3 Philosophische Arzeneykunst oder von Gebrechen der Sensationen, des Verstandes, und des Willens / von M.A. Weikard. in der Andreäischen Buchhandlung, Frankfurt am Main.*

Willcutt, E.G., 2012. The Prevalence of DSM-IV Attention-Deficit/Hyperactivity Disorder: A Meta-Analytic Review. *Neurotherapeutics* 9, 490-499.

Willcutt, E.G., Nigg, J.T., Pennington, B.F., Solanto, M.V., Rohde, L.A., Tannock, R., Loo, S.K., Carlson, C.L., McBurnett, K., Lahey, B.B., 2012. Validity of DSM-IV attention deficit/hyperactivity disorder symptom dimensions and subtypes. *J Abnorm Psychol* 121, 991-1010.

Wolraich, M., Brown, L., Brown, R.T., DuPaul, G., Earls, M., Feldman, H.M., Ganiats, T.G., Kaplanek, B., Meyer, B., Perrin, J., Pierce, K., Reiff, M., Stein, M.T., Visser, S., 2011. ADHD: clinical practice guideline

for the diagnosis, evaluation, and treatment of attention-deficit/hyperactivity disorder in children and adolescents. *Pediatrics* 128, 1007-1022.

World Health Organization, 2018. International statistical classification of diseases and related health problems (11th Revision).

Xu, G., Strathearn, L., Liu, B., Yang, B., Bao, W., 2018. Twenty-Year Trends in Diagnosed Attention-Deficit/Hyperactivity Disorder Among US Children and Adolescents, 1997-2016. *JAMA Netw Open* 1, e181471.

Yao, S., Kuja-Halkola, R., Martin, J., Lu, Y., Lichtenstein, P., Norring, C., Birgegard, A., Yilmaz, Z., Hubel, C., Watson, H., Baker, J., Almqvist, C., Thornton, L.M., Magnusson, P.K., Bulik, C.M., Larsson, H., 2019. Associations Between Attention-Deficit/Hyperactivity Disorder and Various Eating Disorders: A Swedish Nationwide Population Study Using Multiple Genetically Informative Approaches. *Biol Psychiatry* 86, 577-586.

Yeh, J.Y., Hou, T.Y., Tseng, W.T., Chen, V.C., Yang, Y.H., Kuo, T.Y., Weng, J.C., Lee, C.T., Chen, Y.L., Lee, M.J., 2020. Association Between Attention Deficit Hyperactivity Disorder and Risk of Burn Injury: A Propensity-Matched Cohort Study. *Neuropsychiatr Dis Treat* 16, 1249-1255.

Yi, Z., Jing, L., 2015. Prevention and treatment guidelines for attention deficit hyperactivity disorder (the 2nd edition). Peking University Medical Press, Beijing.

Young, S., Moss, D., Sedgwick, O., Fridman, M., Hodgkins, P., 2015. A meta-analysis of the prevalence of attention deficit hyperactivity disorder in incarcerated populations. *Psychol Med* 45, 247-258.

Young, Z., Moghaddam, N., Tickle, A., 2020. The Efficacy of Cognitive Behavioral Therapy for Adults With ADHD: A Systematic Review and Meta-Analysis of Randomized Controlled Trials. *J Atten Disord* 24, 875-888.

Ystrom, E., Gustavson, K., Brandlistuen, R.E., Knudsen, G.P., Magnus, P., Susser, E., Davey Smith, G., Stoltenberg, C., Suren, P., Haberg, S.E., Hornig, M., Lipkin, W.I., Nordeng, H., Reichborn-Kjennerud, T., 2017. Prenatal Exposure to Acetaminophen and Risk of ADHD. *Pediatrics* 140, e20163840.

Zang, Y., 2019. Impact of physical exercise on children with attention deficit hyperactivity disorders: Evidence through a meta-analysis. *Medicine (Baltimore)* 98, e17980.

Zeng, Y., Tang, Y., Yue, Y., Li, W., Qiu, X., Hu, P., Tang, J., Wang, H., Yang, X., Qu, Y., Mu, D., 2019. Cumulative evidence for association of parental diabetes mellitus and attention-deficit/hyperactivity disorder. *Neurosci Biobehav Rev* S0149-7634(19)30721-3.

Zhang, J., Diaz-Roman, A., Cortese, S., 2018. Meditation-based therapies for attention-deficit/hyperactivity disorder in children, adolescents and adults: a systematic review and meta-analysis. *Evid Based Ment Health* 21, 87-94.

Zhang, L., Reif, A., Du Rietz, E., Lagerberg, T., Butwicka, A., D'Onofrio, B.M., Johnell, K., Pedersen, N.L., Larsson, H., Chang, Z., 2020a. Comedication and Polypharmacy With ADHD Medications in Adults: A Swedish Nationwide Study. *J Atten Disord*, 1087054720923725.

Zhang, M., Wang, C., Zhang, X., Song, H., Li, Y., 2020b. Association between exposure to air pollutants and attention-deficit hyperactivity disorder (ADHD) in children: a systematic review and meta-analysis. *Int J Environ Health Res*, 1-13.

Journal Pre-proof