

Mercy Pregnancy Emotional Wellbeing Study

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What do we know?



KEY QUESTIONS

•Does antenatal and postnatal depression impact on child outcomes and if so how and how much?

•Does maternal depression increase the risk for child mental illness across lifespan?

•Is maternal antidepressant treatment beneficial or a risk for offspring?





WHAT ARE THE CHALLENGES IN ANSWERING THESE QUESTIONS ?

Confounding Variables

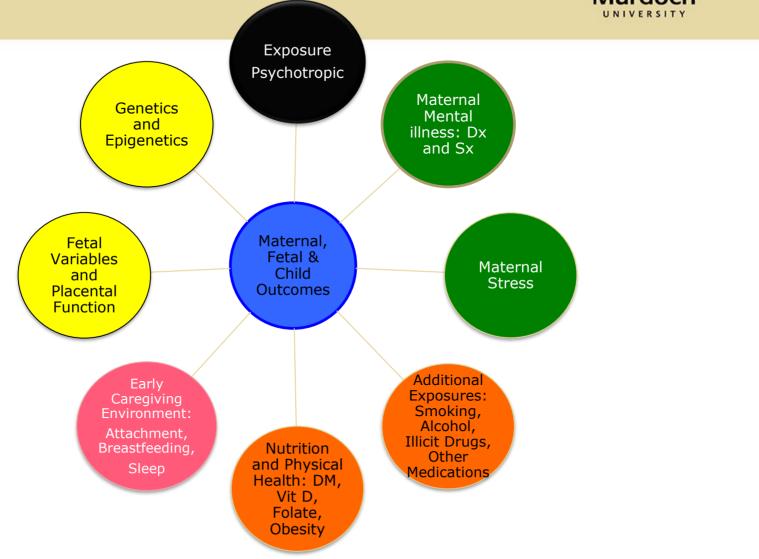












Prenatal or Postnatal? (Galbally & Lewis, Curr Op Psychology, 2017)



Figure 1

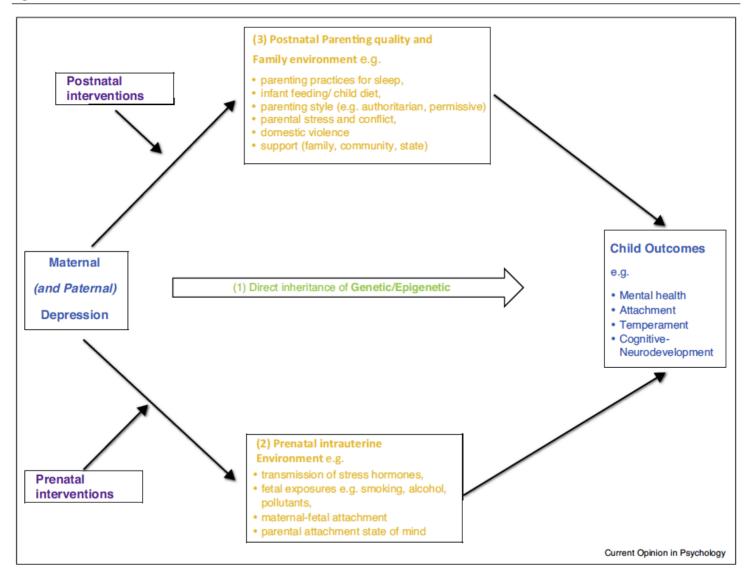


Illustration of potential relationships between parental depression, parenting and child outcomes.



THEN HOW DO WE START TO ANSWER THESE QUESTIONS USING DIFFERENT METHODOLOGY?

Serotonin discontinuation syndrome following in utero exposure to antidepressant medication: prospective controlled study

Megan Galbally, Andrew J. Lewis, Jarrad Lum, Anne Buist

Objectives: The aim of the present study was to examine neonatal symptoms previously reported to be associated with exposure to antidepressant medication in late pregnancy in a group of infants exposed to antidepressants, using a prospective and controlled design. Method: A prospective case-control study recruited 27 pregnant women taking antidepressant medication and 27 matched controls who were not taking antidepressant medication in pregnancy. Of the 27 women taking medication, 25 remained on medication in the third trimester and, of these, 23 women had complete data available. In pregnancy and after delivery women were assessed with the Beck Depression Inventory-II and a purpose-designed questionnaire. After delivery mothers were asked a set of nine questions pertaining to symptoms of discontinuation in the irrewborn and questions about pregnancy and delivery complications.

Results: There was an increased risk of discontinuation symptoms in neonates exposed to antidepressant medication in late pregnancy and an association with higher dose medication. The study group were found to be significantly more likely to display behaviour such as crying, jitteriness, tremor, feeding, reflux and sneezing and sleep for <3 h after a feed. They also had significantly higher rates of jaundice and admissions to the special care nursery.

Conclusions: Exposure to antidepressants in late pregnancy is associated with a range of symptoms in the neonate that are consistent with the effects of exposure to antidepressants in late pregnancy. The clusters of symptoms most highly correlated are the gastrointestinal and central nervous system symptoms. These findings help to identify the common symptoms associated with a neonatal serotonin discontinuation syndrome. **Key words:** antidepressants, discontinuation, pregnancy, selective serotonin re-uptake inhibitor.

Australian a

Research

Child developmental outcomes in preschool children following antidepressant exposure in pregnancy

Developmental outcomes of children exposed to antidepressants in pregnancy



Megan Galbally, Andrew J. Lewis, Anne Buist

Objective: To examine the developmental outcomes in children exposed to antidepressants *in utero* and compare those to children not exposed to these medications Method: A prospective case-controlled study of children exposed to antidepressants in pregnancy assessed 22 exposed and 19 not exposed children using the Bayley Scales of Infant Development, third edition. The control group was measured at a mean age of 23.09 (SD 3.82) months and the medicated group at 28.53 months (SD 6.22). Maternal variables were assessed using a purpose-designed questionnaire and the Beck Depression Inventory (II) in pregnancy and at three assessments in the postpartum.

Results: Children exposed to antidepressant medication in pregnancy scored lower on motor subscales in particular on fine motor scores than non-exposed children with a moderate effect size of Coheris d = 0.47 fine motor and Coheris d = 0.43 for gross motor. Due to lack of power these findings did not reach conventional criteria for statistical significance. There was no association found between maternal depression and neurodevelopment. Conclusions: This finding of a possible effect from antidepressant exposure in pregnancy on children's motor development is similar to the findings from a previous study. Future research is needed which assesses children at an older age using specific assessments

of motor development. Kev words: antidepressants, child development, pregnancy.

Australian & New Zealand Journal of Psychiatry 1–9 DOI: 10.1177/0004867415569800

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Megan Galbally^{1,2}, Andrew J Lewis³ and Anne Buist⁴

Abstract

Objective: To examine child developmental outcomes in preschool-aged children exposed to antidepressant medication in pregnancy and compare their outcomes to children not exposed.

Method: A prospective case-controlled study of 20 children exposed to antidepressants in pregnancy and 21 unexposed controls was available from the Victorian Psychotropic Registry. Child development outcomes at 4 years of age were assessed using the Wechsler Preschool and Primary Scale of Intelligence, third edition; the Movement Assessment Battery for Children; Behaviour Rating Inventory of Executive Functioning–Preschool; and the Child Behavior Checklist (1.5–5 years). Maternal depression was assessed using the Beck Depression Inventory-II in pregnancy and at four time



 Received: 26 August 2016
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 DOI 10.1002/mpr.1558

WILEY

ORIGINAL ARTICLE

Mercy Pregnancy and Emotional Well-being Study (MPEWS): Understanding maternal mental health, fetal programming and child development. Study design and cohort profile

Megan Galbally^{1,2,3} | Marinus van IJzendoorn⁴ | Michael Permezel⁵ | Richard Saffery⁶ | Martha Lappas^{7,8} | Joanne Ryan^{6,9} | Elisabeth van Rossum¹⁰ | Andrew R. Johnson¹¹ | Douglas Teti¹² | Andrew J. Lewis¹



Pregnancy Emotional Wellbeing Study

- Cohort 1: 282 Women now 3-4 years pp Victoria
- Cohort 2: 204 Women now over 12 months pp Victoria
- Cohort 3: WA 200 recruited Metro Perth
- **Cohort 4:** Geraldton, Bunbury and Kalgoorlie
- 3 Groups recruited: Mental Disorders, Antidepressant/Psychotropic Exposed, Control

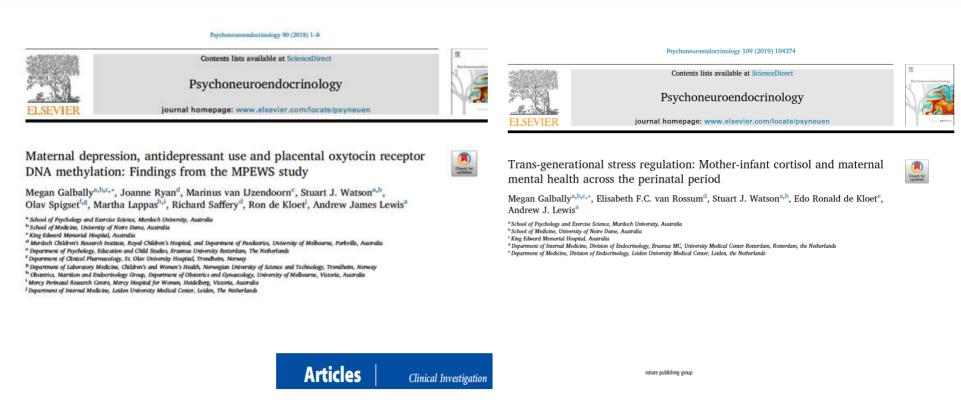




WHAT HAVE THEY SHOWN US SO FAR?

Biological





Neonatal adaptation following intrauterine antidepressant exposure: assessment, drug assay levels, and infant development outcomes

Megan Galbally¹, Olav Spigset², Andrew R. Johnson³, Rolland Kohan⁴, Martha Lappas⁵ and Andrew J. Lewis¹

Psychological



Journal of Affective Disorders 238 (2018) 218-225



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AFFECTIVE DISonder

Research paper

Perinatal maternal depression, antidepressant use and infant sleep outcomes: Exploring cross-lagged associations in a pregnancy cohort study



Megan Galbally^{a,b,c,*}, Stuart J. Watson^{a,b}, Doug Teti^d, Andrew J. Lewis^a

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Check for updates

Original Research

Breastfeeding, Antidepressants, and Depression in the Mercy Pregnancy and Emotional Well-Being Study

Megan Galbally, MBBS, PhD^{1,2,3}, Stuart J. Watson, PhD^{1,2}, Helen Ball, PhD⁴, and Andrew James Lewis, PhD¹ international lactation consultant association®

Journal of Human Lactation I-I0 © The Author(s) 2018 Reprints and permissions: sagepub.com/journals/Permissions.nav DOI: 10.1177/0890334418758558 DOI: 10.1177/0890334418758558 Journals:sagepub.com/home/hl SAGE

Social



Psychological Medicine

cambridge.org/psm

Original Article

Cite this article: Galbally M, Watson SJ, Permezel M, Lewis AJ (2018). Depression across pregnancy and the postpartum. antidepressant use and the association with female sexual function. Psychological Medicine 1-10. https://doi.org/10.1017/ 50033291718002040

Received: 13 April 2018 Revised: 5 July 2018 Accepted: 16 July 2018

Key words: Antidepressants: depression: pregnancy;

Depression across pregnancy and the postpartum, antidepressant use and the association with female sexual function

Megan Galbally^{1,2,3}, Stuart J. Watson^{1,2}, Michael Permezel⁴ and Andrew J. Lewis¹

¹School of Psychology and Exercise Science, Murdoch University, Murdoch, Australia: ²School of Medicine, University of Notre Dame, Perth, Australia; ³King Edward Memorial Hospital, Perth, Australia and ⁴Department of Obstetrics and Gynaecology, University of Melbourne, Parkville, Australia

Abstract

Background. There is an established relationship between depression and sexual functioning in women. However, there is limited research examining the relationship between perinatal depression and sexual functioning

Methods. This study draws on the Mercy Pregnancy and Emotional Wellbeing Study and reports on 211 women recruited in early pregnancy and followed to 12 months postpartum. Women were assessed for depression using the Structured Clinical Interview for the DSM-IV. repeated measurement of depressive experitories using the Edinburgh Postnatal Depression

Original Article

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	Journal of Psychosomatic Research	ferrer Psychosomatic Baranch
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Journal of Psychosomatic Research 111 (2018) 91-99

Exercise frequency and maternal mental health: Parallel process modelling across the perinatal period in an Australian pregnancy cohort

Stuart J. Watson^{a,b}, Andrew J. Lewis^a, Philip Boyce^c, Megan Galbally^{a,b,d,*}

a School of Psychology and Exercise Science, Murdoch University, Australia ^b School of Medicine, University of Notre Dame, Australia ^c Westmead Clinical School, Sydney Medical School, University of Sydney, Australia ^d King Edward Memorial Hospital, Australia

HJISIP

The role of trauma and partner support in perinatal depression and parenting stress: An Australian pregnancy cohort study

Megan Galbally^{1,2,3}, Stuart J Watson^{1,2}, Philip Boyce⁴ and Andrew I Lewis¹

International Journal of Social Psychiatry 2019, Vol. 65(3) 225-234 © The Author(s) 2019 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/0020764019838307 journals.sagepub.com/home/isp



Is the Mother-Infant Relationship Relevant?

 "unique without parallel, established unalterably for a whole lifetime as the first and strongest love object and as a prototype for all later love relations" *s.Freud*

Proc. Natl. Acad. Sci. USA Vol. 93, pp. 705–708, January 1996 Medical Sciences

Male fetal progenitor cells persist in maternal blood for as long as 27 years postpartum

(pregnancy/chimerism/CD34/CD38)

Diana W. Bianchi*†‡, Gretchen K. Zickwolf*†, Gary J. Weil*†, Shelley Sylvester*, and Mary Ann DeMaria†

Nature Reviews Neuroscience | AOP, published online 12 December 2008; doi:10.1038/nrn2280



The expectant brain: adapting for motherhood

Paula J. Brunton & John A. Russell

nature neuroscience

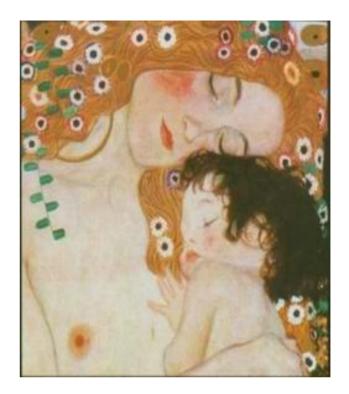
Pregnancy leads to long-lasting changes in human brain structure

ARTICLES

Elseline Hoekzema^{1–3,8}, Erika Barba-Müller^{1,8}, Cristina Pozzobon⁴, Marisol Picado¹, Florencio Lucco⁴, David García-García⁵, Juan Carlos Soliva¹, Adolf Tobeña¹, Manuel Desco⁵, Eveline A Crone^{2,3}, Agustín Ballesteros⁴, Susanna Carmona^{1,5,6,9} & Oscar Vilarroya^{1,7,9}

IF SO, HOW DO WE MEASURE THIS RELATIONSHIP?









Available online at www.sciencedirect.com ScienceDirect



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Depression and parenting: the need for improved intervention models

Megan Galbally^{1,2,3,4} and Andrew J Lewis^{1,4}

The impact of maternal depression on parenting is well established and there is a clear interaction between maternal depression and parenting that is predictive of child outcomes. The research on paternal depression is more limited but suggests the father's mental health may be an independent risk factor for both parenting and child outcomes. There is insufficient evidence that treatment of depression alone – be it through pharmacological or psychological interventions – is able to substantially reduce the impact of depression on child outcomes. The evidence of interventions aimed at parenting and/or child outcomes in the context of depression is limited and the findings that are available are mixed. offspring, second, the effect of maternal depression in pregnancy through fetal programming and, third, the effect of maternal and paternal depression on parenting behavior, the parent-child relationship and general family functioning [2]. Investigating this third point on the role of parenting can itself be divided into three areas of focus:

- 1. Parenting experience such as parenting stress
- 2. Parenting behaviors and practices
- Quality of the parent-child relationship including attachment security.

Parenting experience such as parenting stress
 Parenting behaviors and practices
 Quality of the parent-child relationship including

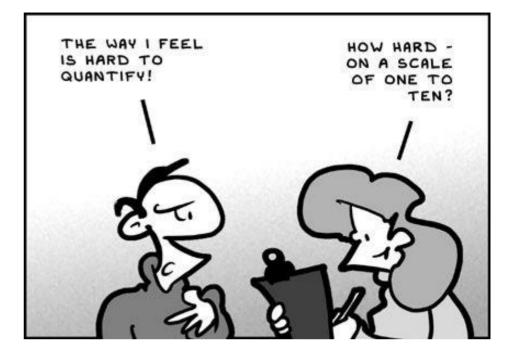
attachment security.

Measuring Mother-Infant Relationship In MPEWS



 Maternal Self-Report Attachment, Bonding, Support, Parenting Stress

 6 month video tapes



• 12 month SSP

Reference: Leunig



lenig

INTERVIEW WITH A BABY





let me put it differently. Do babies have particular, inalienable, human rights?



I seem to have hit some sont of glass ceiling here.





HOW CENTRAL IS THE MOTHER-INFANT RELATIONSHIP IN TREATING WOMEN WITH PERINATAL MENTAL **HEALTH DISORDERS?**

How to assess the mother-infant relationship



Self report measures:

- Cost effective and timely.
- How well placed is a mother to provide accurate information about the relationship with her infant?
- What about women with mental health problems?



What other options are available to assess the mother-infant relationship?

Observational measure: Emotional Availability Scales





Emotional availability provides the framework for assessing the quality of a mother-infant interaction.

The Emotional Availability Scales (Biringen, 2008) is a widely applied observational measure.

- Global assessment.
- Emotional connection.
- Relational assessment.
- Space for positive and negative emotions.

Application of the Emotional Availability Scales



- Recommended length of interaction:
 - **30 40 minutes.**
 - 20 minutes for valid and reliable sample.
- However, what you can see in the EA literature:
 - Interactions 1 5 minutes: almost 10%.
 - $_{\odot}$ Interactions 6 10 minutes: 20%.
 - ∘ Interactions 11 19 minutes: 22%.



Think about the mothers and infants that you have worked with – how has your assessment of the relationship changed over time?

Let's watch a mother and her infant together





Let's watch a mother and her infant together







WHAT INFORMATION COULD YOU EXTRACT FROM THAT INTERACTION AND WHAT MIGHT YOU MISS?

Coding challenges to consider



- Length of the interaction sample.
- Video recording versus naturalistic observation.
- Setting:
 - Laboratory/hospital versus home visit.
 - o Why is the context relevant?
- Reliability of the coder.
- Mother-infant activity observed:
 - Face to face with no props versus free play.

How might these details impact on the mother-infant assessment?



WHAT ABOUT THE POSSIBILITY OF NEW DISCOVERIES IN MOTHER-INFANT BEHAVIOUR?

Pacifier use in the mother-infant interaction



What we observed:

- Maternal use of the pacifier during a mother-infant interaction for non-sleep settling purposes.
- What we investigated:
- Maternal psychosocial predictors of pacifier use.
 - Maternal trauma, depression & emotional availability.





WHAT HAPPENS WHEN WE DO NOT FIND IN OUR RESEARCH WHAT WE WERE EXPECTING TO SEE?

Maternal depression and the mother-infant relationship



Literature cites the negative effect of maternal depression on mother-infant interactions:

- Increased negative affect;
- Reduced maternal responsiveness;
- Reduced verbal involvement of infant and mother;
- Reduced shared gaze activity;
- Reduced touch.





What we saw when we looked more closely at the literature

- The variety of ways the quality of the interaction was measured.
- Who comprised the `depressed' women?
 - $_{\odot}$ $\,$ How is depression conceptualized:
 - Symptomatically;
 - Diagnostically.
- Timing of measurement:
 - e.g., 8 weeks postpartum versus
 6 months postpartum.
- Benefit of longitudinal measurement:
 - Importance of the antenatal <u>and</u> postnatal period.







IS OUR FINDING THAT MATERNAL DEPRESSION WAS NOT A STRONG PREDICTOR OF MATERNAL EMOTIONAL AVAILABILITY SURPRISING?

Treatment of the mother and the mother-infant dyad



- Evidence of limited improvement in the mother-infant relationship when the mother's mental health is treated in isolation.
- Consider triggers for women's mental health during the perinatal period:
 - How can these be managed without working <u>with</u> the dyad?
 - Long-term outcomes for the mother and the infant.



Dual treatment model.



HOW DO YOU ASSESS THE MOTHER-INFANT DYAD IN CLINICAL PRACTICE?

Translation of a research tool to clinical practice



- Emotional Availability Scales:
 - Time and financial cost.
 - $_{\odot}$ Training required for reliability.
 - $_{\odot}\,$ Information the scale provides.
 - Application to clinical intervention.
- What is the alternative?
 - Self report.
 - \circ Another observational measure.
 - $_{\odot}$ Working with the mother alone?



Dual treatment model



- Assessment of:
 - The mother's mental health.
 - What is observed between the dyad e.g., using the Emotional Availability Scales.
 - The maternal representations of the relationship. For example – observed maternal intrusiveness:
 - May represent a mother's desire for her child to achieve and be independent; or
 - May be driven by a need of the mother to prove that she is wanted by her child and show the child's dependence.



Very different implications for clinical treatment depending on the information that we have.



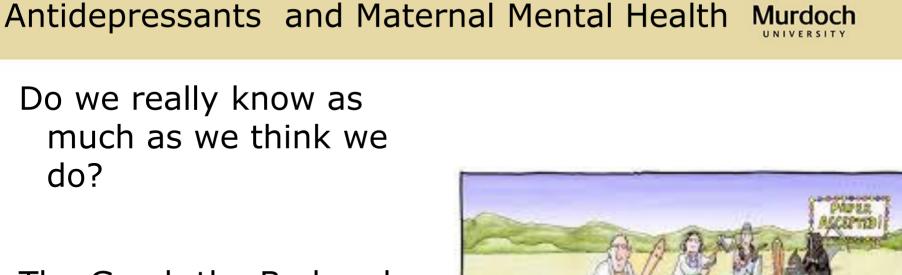
So let's return to our first question: HOW CENTRAL IS THE MOTHER-INFANT RELATIONSHIP IN THE TREATMENT OF PERINATAL MENTAL HEALTH DISORDERS?

Do we really know as much as we think we do?

Child Development, Mental Disorders,

- The Good, the Bad and the Ugly in this Research field
- (a.k.a. what we learnt doing systematic reviews and then undertaking our own research in this area)

Most scientists regarded the new streamlined peer-review processas "quite an improvement.







WHAT DO WOMEN WITH PERINATAL DEPRESSION WANT TO KNOW?



Child Outcomes

Perinatal depression associated with adverse child outcomes across domains

- Complex, evolving information
- Research methodology crucial for interpreting results
- In general, effect sizes are small
- Not inevitable

What is a helpful approach for parents?



WHAT MAKES A USEFUL OUTCOME MEASURE?

Child outcomes



Measurable and meaningful

- What is meaningful in terms of child outcomes?
 To parents, to governments, to schools, to researchers
 The role of a diagnosis
- When is the ideal time to identify a child who is vulnerable? As early as possible for intervention Risk of misidentification of cases
- When is the ideal time to measure an outcome?



WHAT MIGHT INDICATE THAT A CHILD IS ON A VULNERABLE DEVELOPMENTAL TRAJECTORY?



- How can these be measured?
 - Parent report
 - Objective assessment
 - Validated tool

At what age?

Is there an effective intervention?





What is it?

An evolving concept in early childhood

 Overlap with temperament research

Suite of cognitive functions that regulate behaviour

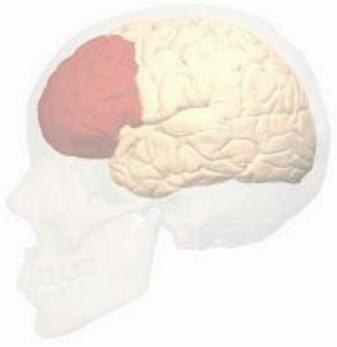
- Inhibit impulses
- Delay gratification
- Control attention
- Hold and manipulate relevant information



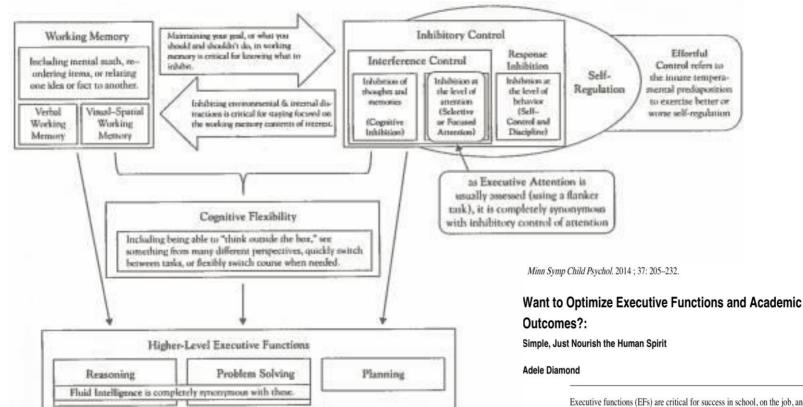


In adults

- Unity and diversity model (Miyake, 2000) most widely, though not universally, accepted
- Core functions:
 - Working memory Cognitive flexibility (shifting) Inhibition
 - Deficits in a range of disorders







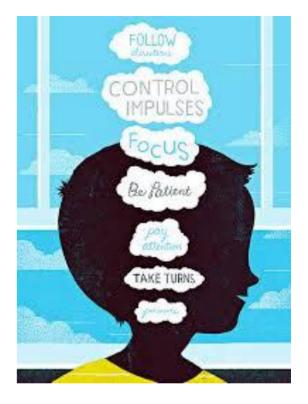
Executive functions (EFs) are critical for success in school, on the job, and in life. EFs suffer if you are lonely, sad, stressed, or not physically fit. Therefore, if we care about academic outcomes, we should care that students feel they are in a supportive community they can count on, that they are happy (even joyful), and that their bodies are strong and healthy. A school curriculum that ignores children's emotional, social, or physical needs is likely to find that those unmet needs will work against achieving the academic goals.



May be different during early development

Infancy and Preschool

- Attentional control observed in infancy
- Unitary EF factor found in early childhood
- At some stage of development, around age 3, separable functions can be measured





IS EXECUTIVE A GOOD CANDIDATE OUTCOME MEASURE?



In early development

- Thought until relatively recently that executive function didn't emerge until later in development
 - Now understood that EF emerges relatively early
 - Matures over a prolonged period
 - First 5 years may be particularly important (Garon et al. 2008)
- Stable individual differences, yet EF development is malleable
 - Develops as the underlying brain processes adapt to the environment
 - Concept of neuroplasticity, modifiable through experience
 - Early childhood is a period of relative plasticity
 - Very prolonged development
- We grow our brains by using them



Is this a meaningful measure?

- Predicts key developmental outcomes
 Self and social understanding
 School readiness (early mathematics and reading)(Blair &
 - Razza, 2012)
- Executive function in childhood predicts outcomes at age 32

Physical health, drug dependence , socioeconomic status, criminal convictions when controlling for socioeconomic status and IQ in childhood (Moffitt et al. 2011)

• Associated with child psychopathology

Conduct Disorder, Autism Spectrum Disorder, ADHD Problem behaviours such as aggression



How does it relate to behaviour in children?



Behavioural control

Inhibition Consider consequences Shifting tasks Stopping a preferred activity



Emotional control

Reactivity to stimulation Regulation of reaction



Cognitive control

Working memory Planning tasks Organisation Intitiation of tasks

Executive Fucntion Deficit



What might this look like in children?

- Emotional dysregulation
- Difficulty adjusting to more structured environments
- Behavioural disturbances
- Diagnosis of ADHD, ODD

Executive Function: Potentially Modifiable?



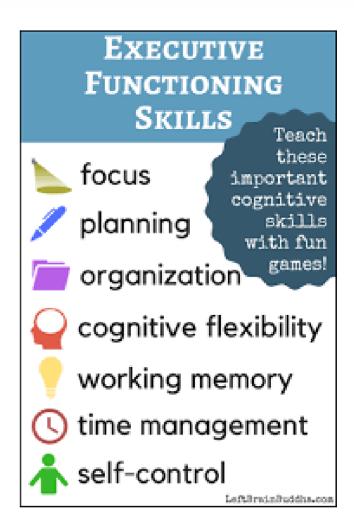
Research supports this (Diamond, 2012)

- School-based programs
- Parent training

Improvements with nonspecific interventions

• Yoga, mindfulness

Those most disadvantaged appear to benefit most





HOW IS EXECUTIVE FUNCTION MEASURED IN CHILDREN?



Measurement of Executive Function

Self-report:

- Parents
- Teachers
- BRIEF-P (Behaviour Rating Inventory of Executive Function – Parent version)

Objective measures:

- Many used in this age group
- Few have established reliability
- NEPSY-II
- Shape School

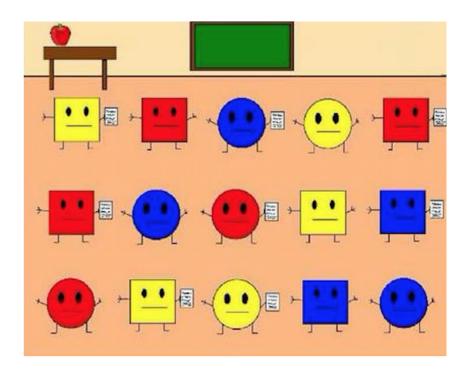
Child Executive Function



Shape School

(Espy, 1997)

- Test of inhibition and shifting
- Validated for use in children from age 3.5 years
- Four tasks of increasing complexity
- Time to complete tasks and errors recorded, although,
- Validated against other neuropsychological tests of executive function
- Performance has been associated with future academic performance



NEPSY-II



Developmental Neuropsychological Assessment-II

(Korkman et al. 2007)

Set of normed neuropsychological tests

Validated in clinical populations Normed in 1200 children at all age brackets

Children aged 3 to 16 years

Six domains

- Attention and executive function
- Language
- Sensorimotor
- Visuospatial
- Learning and memory
- Social perception

Executive function measures

Statue task

- Test of executive function under 5 years
- Memory for designs task
- Measurement of working memory

Control measures Speeded naming



WHAT ARE THE POTENTIAL ISSUES WITH MEASURING EXECUTIVE FUNCTION IN THIS AGE GROUP?

Issues with Measurement



Conceptual problems

Executive function in young children is a relatively new concept

Timing of measurement

Accounting for normative developmental trajectories

- Highly dependent on other cognitive processes Language, sensory processing
- Testing in this age group is highly dependent on environmental factors

Fatigue

Hunger

Distress

Separation issues



DOES THE LITERATURE REPORT AN ASSOCIATION BETWEEN PERINATAL DEPRESSION AND CHILD EXECUTIVE FUNCTION?



Review of the literature

22 studies examined executive function outcomes following measurement of maternal depressive symptoms across the perinatal period

- Great variation in measurement of exposure, outcome and covariates
- Many drew from the same pools of longitudinal studies
- Mixed findings
- Insufficient to determine with certainty
- Greater consistency with use of reliable measures required



HOW COULD PERINATAL DEPRESSION AFFECT DEVELOPMENT OF EXECUTIVE FUNCTION?



Effect on maternal behaviour

- Reading time?
- Screen time?
- Language exposure?
- Social engagement and enrichment?

Effect on mother-child relationship

Access to social supports

Vygotsky

 Zone of proximal development



Maternal Depression



What might be most important about depression?

• Timing of exposure to depression?

At what point in the lifespan Definition of perinatal period Antenatal or postnatal exposure, or both? Different mediating pathways are involved

• Trajectory of depression?

Acute, single-episode, chronic, relapsing? These may have different effects Many studies are cross-sectional

Mothers in Cohort 2 of MPEWS

- A past history of depression was associated with reduced number of days of at least 15 minutes reading time at 6 months postpartum
- Current depression showed lower average reading time, but not statistically significant
- Differences in screen time not significant
- Objective measure of depression
- Self-report of screen time and reading time
- Data analysed by Dr Grace Cowderoy

Observation of mother-infant interaction

• EAS?

The Overall MPEWS Team...



Melbourne

Key Co- Investigators

Andrew Lewis, Murdoch University Marinus van Ijzendoorn, Leiden and Erasmus Ron de Kloet, Leiden University Elisabeth van Rossum, Erasmus Martha Lappas, University of Melbourne Richard Saffery, MCRI Philip Boyce, University of Sydney Olav Spigset, Norway Douglas Teti Penn State University Helen Ball, Durham University, UK And many more...

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Students: Murdoch, UNDA

Funding: Beyond Blue/NHMRC/Spinnaker Foundation/Telethon/WACHS



Perth





Thanks to MPEWS team who are not here...

And the Mothers and Children who participate in our study

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