



Fás Aníos in Éirinn  
Growing Up in Ireland

# Pregnancy Complications and Child Mental Health

**Emma Butler (RCSI), Michelle Spirtos (TCD), Linda O Keeffe (UCC), Mary Clarke (RCSI) and Niamh Dooley (RCSI).**

08.11.2023



**RCSI**

UNIVERSITY  
OF MEDICINE  
AND HEALTH  
SCIENCES



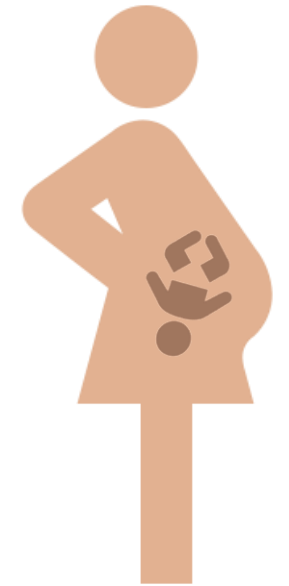
**SPHeRE** <sup>HRB</sup>


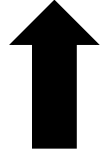
Structured Population health, Policy  
and Health-services Research Education

Specific pregnancy complications have been found to associate with increased odds of mental health difficulties in the offspring





## Cumulative pregnancy/obstetric complications?



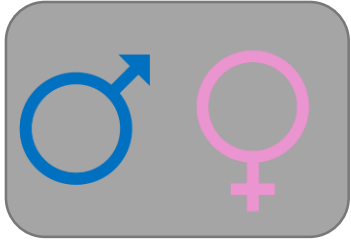
  $\neq$   ADHD or conduct disorder

  $=$  ADHD diagnosis

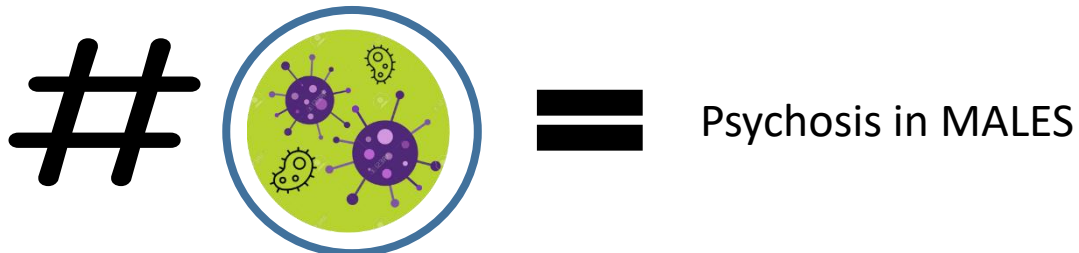
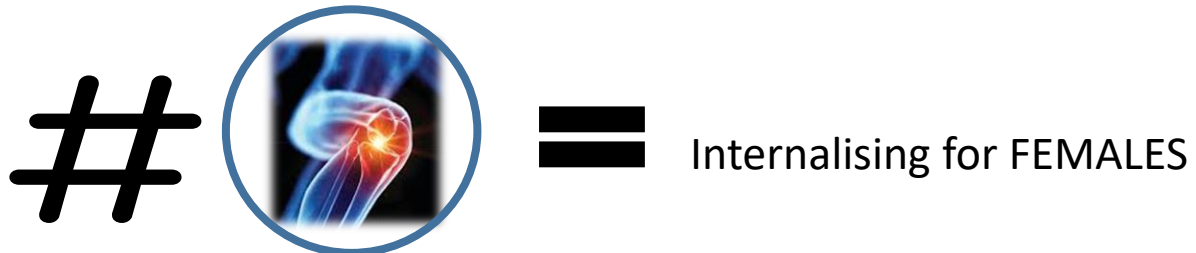
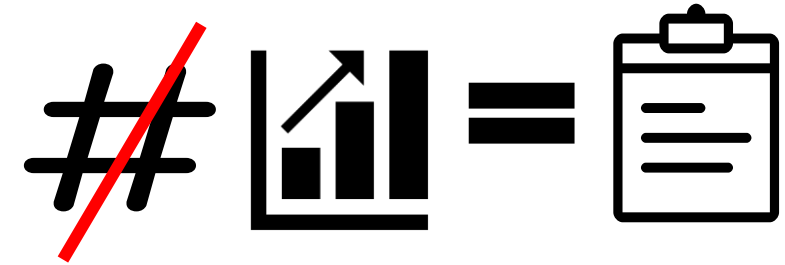
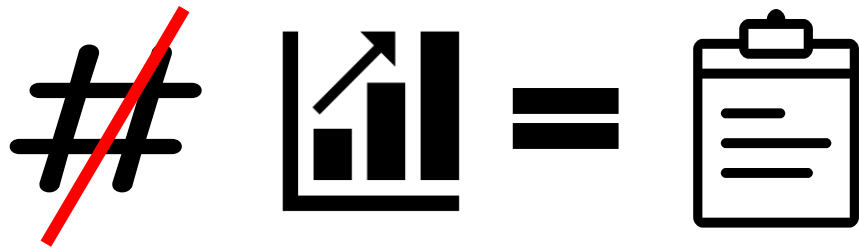
  $=$   CBCL total problems score @ 8years

  $=$   Dose-response between the number of prenatal exposures and the odds of *clinically* elevated CBCL scores

2+



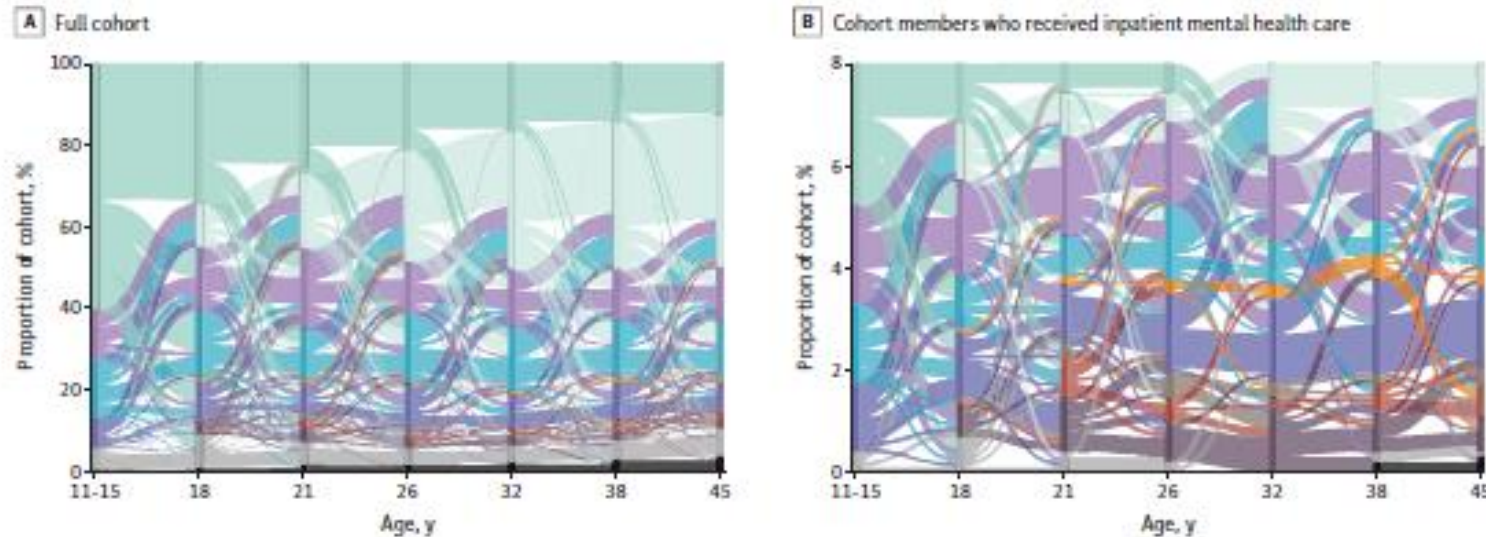
# Subgroups?



# SDQ-total not binary specific diagnosis

Figure 4. Ebb and Flow of Mental Disorders

P



Sankey diagrams show cohort members' shifting diagnoses from 1 assessment phase to the next, from ages 11 to 15 years to age 45 years. The colors of the horizontal bands divide the diagram into different psychiatric statuses, as indicated in the key. The heights of the horizontal bars show the prevalence of different statuses at each assessment

phase. A, Information for the full cohort of 1037 participants. B, Analysis restricted to 83 participants who received inpatient mental-health services (8% of the cohort). Note that it is possible to follow groups across contiguous adjacent assessments, not across the entire panel.

# AIMS OF CURRENT STUDY



## QUESTION 1:

Dose-response  
*cumulative PCs*  
& child mental  
health as  
measured by  
SDQ-total?

## QUESTION 2:

*Cumulative PCs*  
& *clinically*  
*significant*  
SDQ-total  
scores?

## QUESTION 3:

Moderation by  
sex or social  
risk?



11,134  
recruited at 9-  
months

- Exclusions:
- N=1 no PC
- N=38 respondent at 9-months not mother
- N=1794 did not have SDQ @ 5 OR 9 years

9,301 had at  
least one SDQ-  
total

- Exclusions (missing covariates):
- N=663 missing cumulative SR
- N=155 missing smoking

8,483 included  
in fully-adjusted  
models







**Were there any of the following complications with the pregnancy? Tick all that apply:**

Raised blood pressure (In isolation)

Raised blood pressure and protein in the urine (pre-eclampsia)

Urinary or kidney infection

Persistent vomiting or nausea

Gestational diabetes (diet treated)

Gestational diabetes (insulin treated)

Bleeding during second half of pregnancy

Vaginal infection during pregnancy

Intrauterine growth restriction

Rhesus incompatibility

Influenza

Placenta Previa

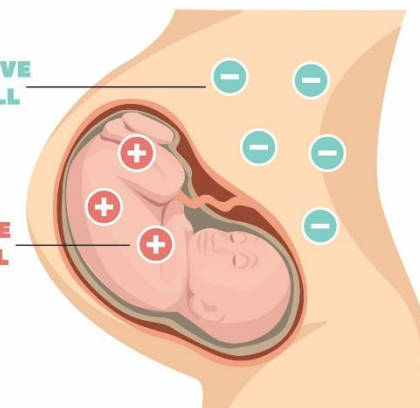
Miscarriage in a multiple pregnancy

Other (please specify)

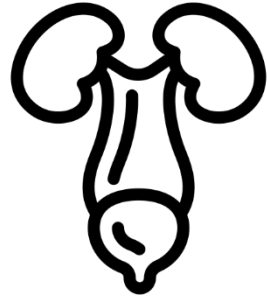


**RH NEGATIVE BLOOD CELL**

**RH POSITIVE BLOOD CELL**

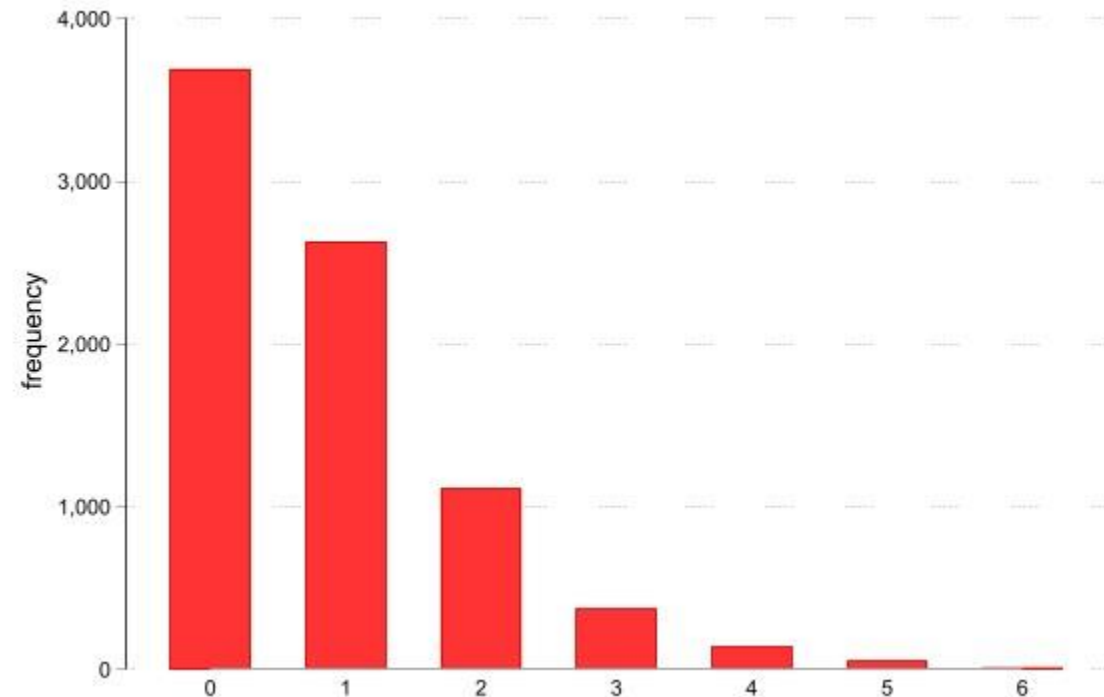






# Cumulative PCs

Specific Pregnancy Complications (n=8483)	(% yes)
Placenta complications	3.24
Pre-eclampsia	7.50
Urinary Tract Infection	15.17
Persistent vomiting	17.64
Late bleeding	6.70
Vaginal infection	3.44
Intra-uterine growth restriction	2.24
Rhesus	3.67
Influenza	3.76
"Other" blood conditions	2.29
"Other" pains	2.44
"Other" high/moderate risk condition	2.39
Maternal condition not thought to impact the fetus	2.83
Gestational Diabetes Mellitus	3.00
Blood pressure problem but not pre-eclampsia	10.32
"Other" problem but not-specified	1.2



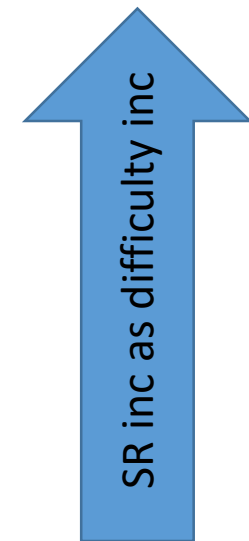
# Cumulative Social Risk

	0 No risk(n%)	1 Low risk(n%)	2 Moderate risk(n%)	3 High risk(n%)
Maternal Age	59.39	40.61	N/A	N/A
Migrancy	81.05	18.95	N/A	N/A
Income	38.60	20.08	41.31	N/A
Maternal Education	10.97	16.35	27.87	44.73
Maternal Relationship	83.90	16.10	N/A	N/A
SR totals	14.55	29.76	31.44	24.25



For validity: "How easy/difficult is it for the household to make ends meet?"

	Wave 1 – 9 months SR mean(SD)	Wave 3 – 5 years SR mean(SD)	Wave 5 – 9 years SR mean(SD)
With great difficulty	2.48 (.62)	2.22 (.78)	2.23 (.79)
With difficulty	2.32 (.71)	1.91 (.91)	2.18 (.84)
With some difficulty	1.96 (.88)	1.69 (.95)	1.85 (.93)
Fairly easily	1.39 (.98)	1.33 (1.04)	1.50 (.98)
Easily	1.20 (1.02)	1.14 (1.04)	1.26 (1.02)
Very easily	.85 (.93)	1.02 (1.09)	1.01 (1.05)


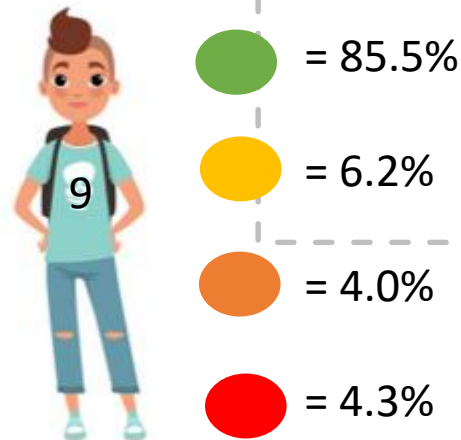
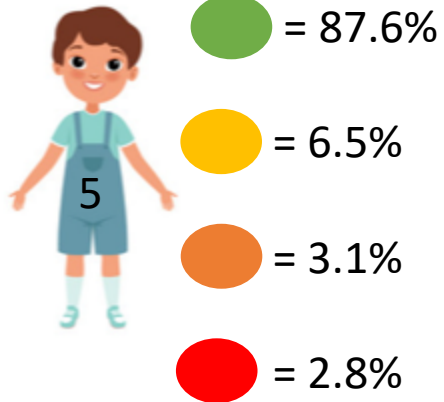


"Answer on your child's behaviour over the past 6 months/school year"

**Response options:**


Not true, somewhat true, certainly true

# OUTCOME: SDQ-total x/40




### EMOTIONAL PROBLEMS X10

Often unhappy, nervous in new situations, many fears, many worries, gets headaches often.



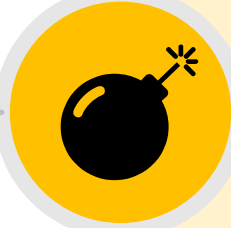
### HYPERACTIVITY/INATTENTION X10

Doesn't finish what they start, doesn't think before acting, restless, easily distracted, constantly fidgeting.



### PEER PROBLEMS X10

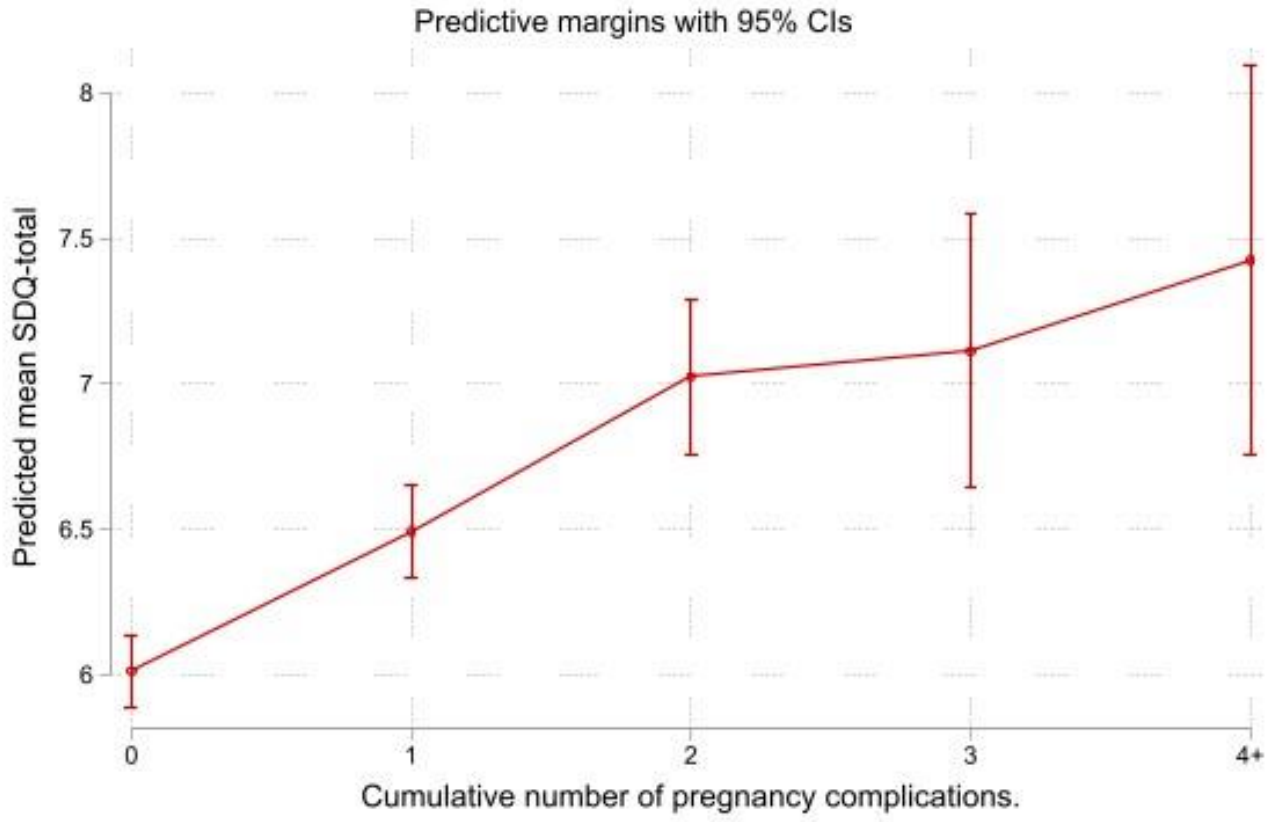
Has no good friends, usually on their own, not generally liked by peers, picked on, gets on better with adults.



### CONDUCT PROBLEMS X10

Disobedient, often gets angry, steals, lies or cheats, fights with peers.

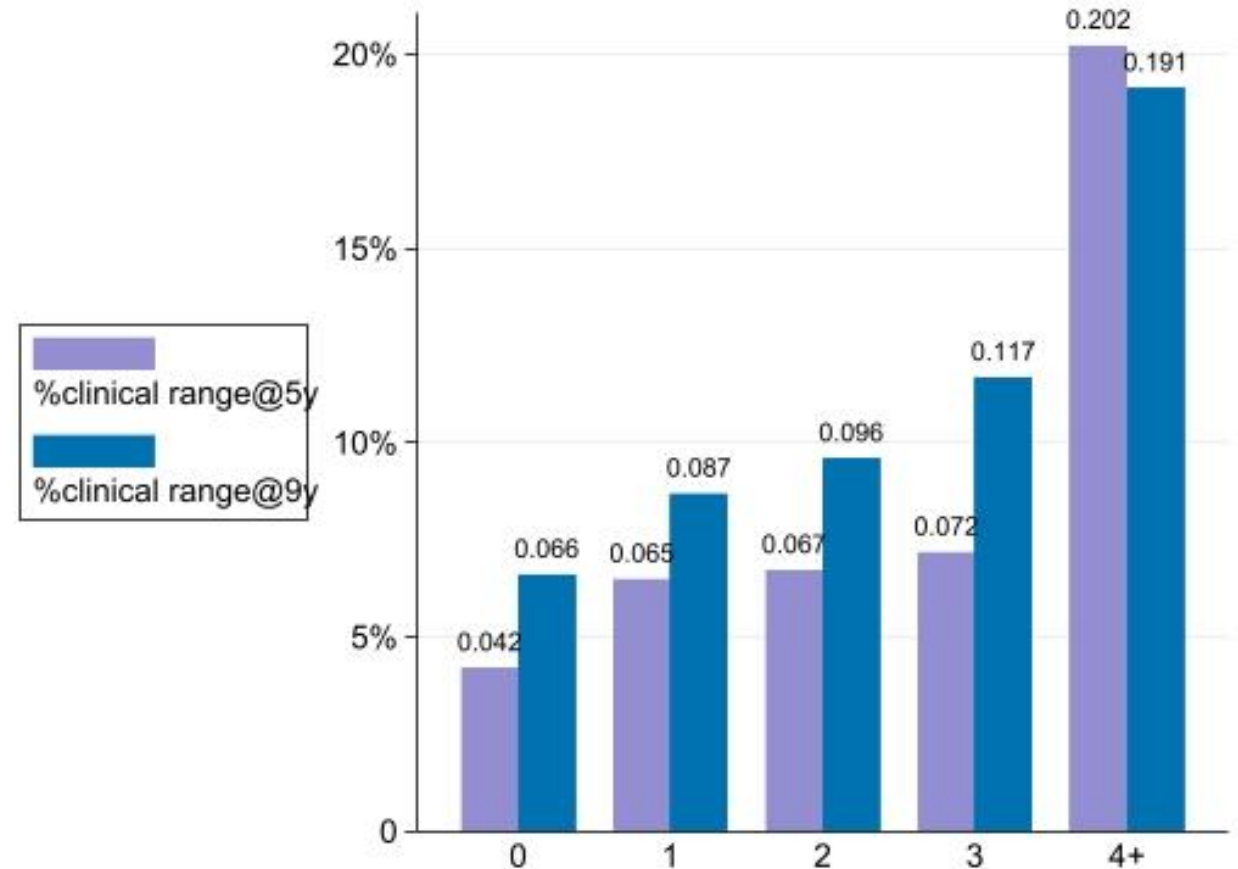
# Aim 1: dose-response relationship between cumulative pregnancy complications and offsprings mental health in middle childhood



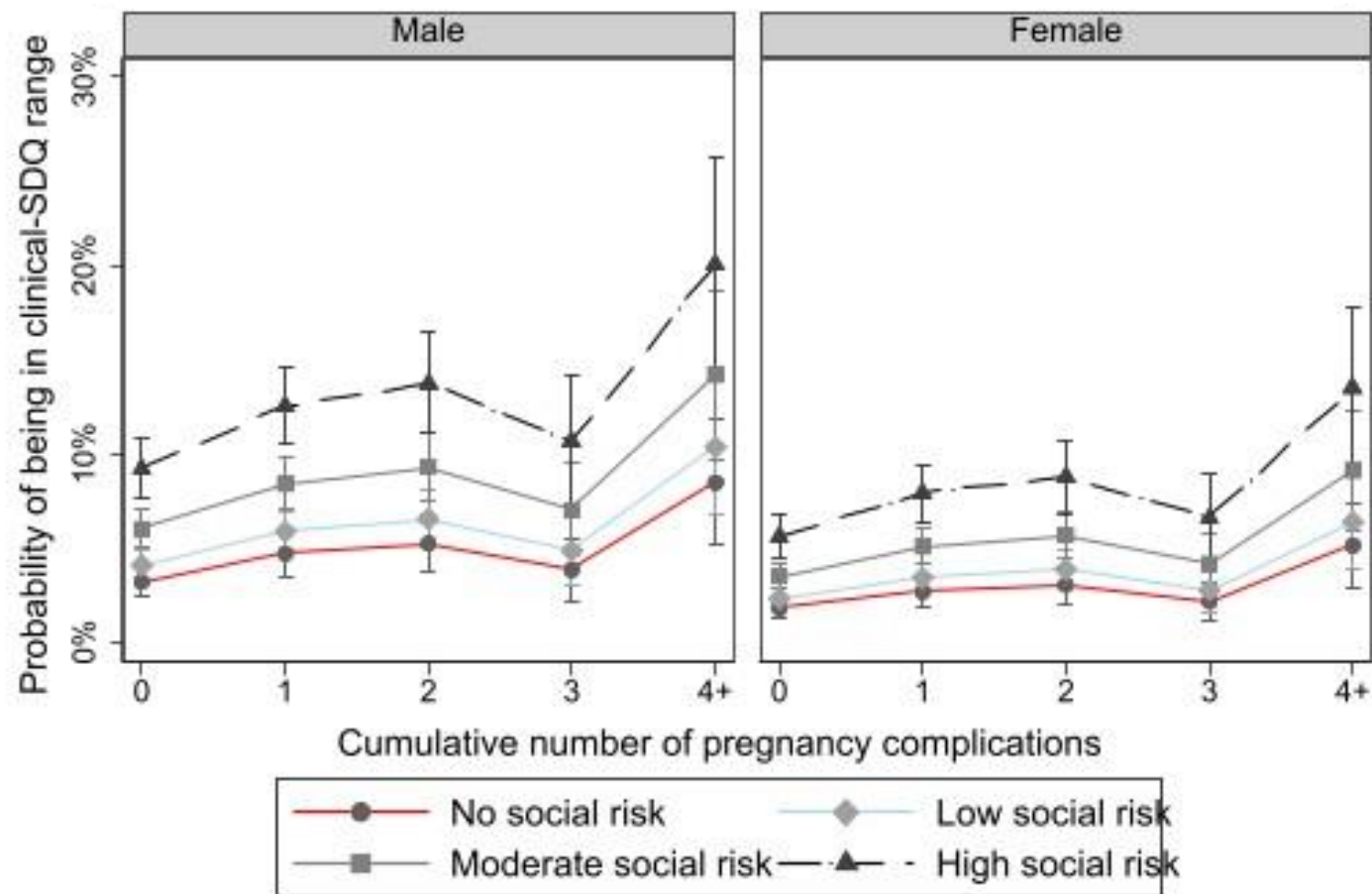
	Unadjusted (fixed PCs + random effect)	Fully Adjusted model
N=	8483	8483
1 complication	1.09(1.05-1.13) Z=4.82, p<.001	1.10(1.06-1.14) Z=5.12, p<.001
2 pregnancy complications	1.21(1.15-1.27) Z=7.65, p<.001	1.20(1.15-1.26) Z=7.63, p<.001
3 pregnancy complications	1.22(1.14-1.31) Z=5.37, p<.001	1.20 (1.12-1.29) Z=5.16, p<.001
4+ pregnancy complications	1.39(1.25-1.54) Z=6.13, p<.001	1.34 (1.21-1.48) Z=5.64, p<.001
Sex		.84 (.81-.87) Z=-11.15, p<.001
Low Social Risk		1.05(1.01-1.10) Z=2.39, p=.02
Moderate Social Risk		1.15 (1.09-1.20) Z=5.86, p<.001
High Social Risk		1.29 (1.22-1.36) Z=9.15, p<.001
Number of people smoking in pregnancy in household		1.12 (1.08-1.16) 1.23 (1.16-1.31) 1.36 (1.20-1.55)

## Aim 2: whether cumulative pregnancy complications were associated with *clinically significant* mental health scores

- Generalised linear mixed models (GLMMs) with a random effect of participant ID.
- Binomial distribution with its default link function.
- Predicted probabilities calculated for being in clinical range for different “types”

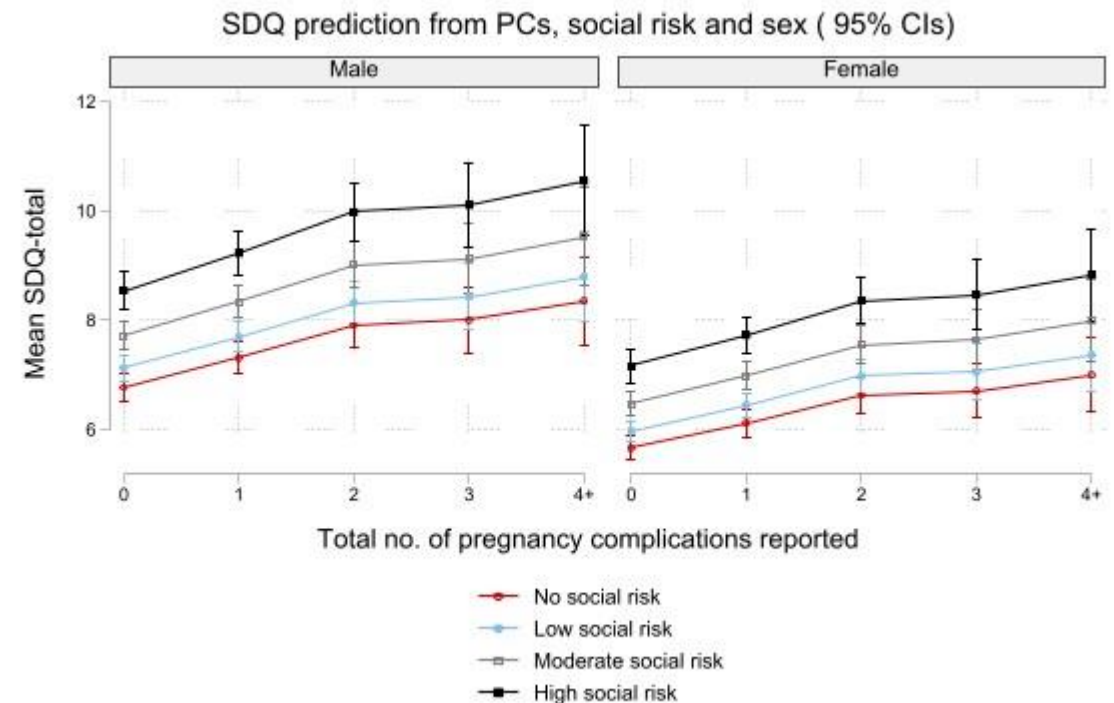
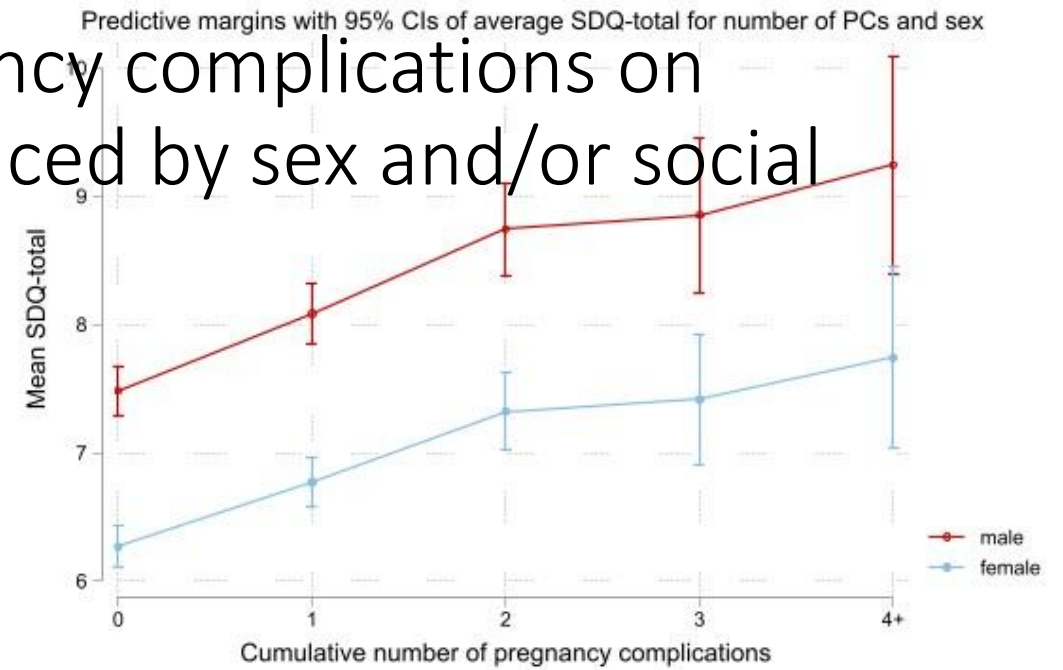


	Unadjusted Model (fixed PCs+ random effect)	Fully Adjusted Model
N=	8483	8483
1 complication	1.88 (1.37-2.59) Z=3.88, p<.001	1.88 (1.37-2.59) Z=3.90, p<.001
2 Pregnancy complications	2.35 (1.56-3.54) Z=4.08, p<.001	2.31 (1.53-3.50) Z=3.96, p<.001
3 Pregnancy complications	2.03 (1.01-4.05) Z=2.00, p=.05	1.77 (.89-3.52) Z=1.62, p=.11
4+ Pregnancy complications	8.50 (4.01-18.03) Z=5.58, p<.001	6.88 (3.29-14.40) Z=5.12, p<.001
Sex (female)		.43 (-.32-.57) Z=-5.86, p<.001
Low Social Risk		1.56 (1.00-2.43) Z=1.95, p=.05
Moderate social Risk		2.84 (1.81-4.46) Z=4.55, p<.001
High Social Risk		6.83 (4.16-11.22) Z=7.58, p<.001
Number of people smoking in pregnancy in household		1.49 (1.24-1.78) Z=4.25, p<.001



# Aim 3: whether the effect of pregnancy complications on childhood mental health was influenced by sex and/or social risk

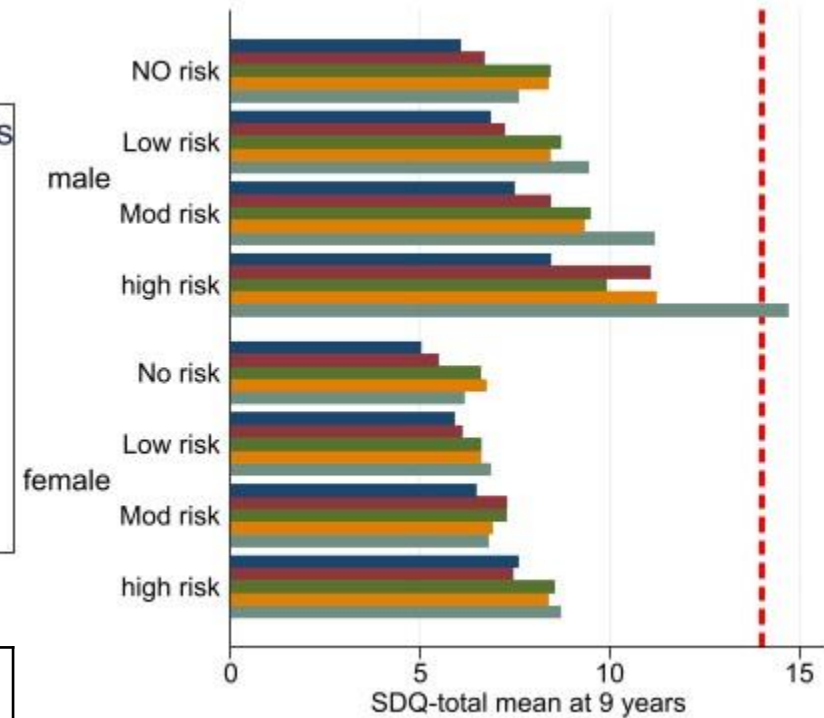
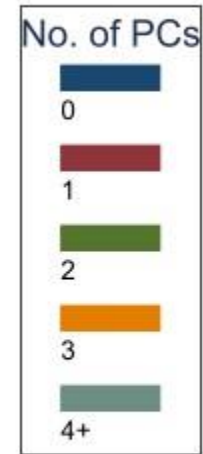
	Effect estimates and 95% CI's after adjustment		
Total no. of PCs	Total (n=8483)	Male (n=4277)	Female (n=4206)
None	REFERENCE		
1	1.89(1.37-2.59)	2.65(1.72-4.10)	1.18(.74-1.89)
2	2.31(1.53-3.50)	3.23(1.83-5.69)	1.50(.82-2.75)
3	1.77(.89-3.52)	2.27(.90-5.69)	1.25(.44-3.58)
4+	6.88(3.29-14.40)	10.73(3.73-30.86)	4.23(1.51-11.83)





## Number needed to SCREEN

Using total complications of none 'v' any with clinical SDQ-total @ 9-years – we would need to screen 17 males to find 1 male with clinical level of problems but 100 females to find 1.



	Sex	Pregnancy complications	Social Risk	Predicted probability of being in the clinical range on SDQ (95%CI)
1	Males	4+	High	22.6% (15.4-29.7%)
2	Females	4+	High	15.6% (10.0-21.2%)
3	Males	0	Low	3.7% (2.9-4.5%)
4	Females	0	Low	2.1% (1.6-2.6%)
5	Males	0	High	9.0% (7.1-10.9%)
6	Females	0	High	5.5% (4.2-6.9%)
7	Males	1	Low	5.5% (4.4-6.7%)
8	Females	1	Low	3.2% (2.5-4.0%)

Note: “ideal types are particularly illustrative for interpretation when independent variables are substantially correlated” (Long & Freese, 2014).

# Implications



## Fás Aníos in Éirinn Growing Up in Ireland

- The majority of women (78.6%) experienced none or only one pregnancy complication. However, 21.4% of women experienced 2 or more PCs, and their children had **double to six fold** increased odds of experiencing clinically significant levels of mental health symptoms in middle childhood compared to children whose mothers experienced no complications
- We cannot infer a causal relationship underlying the association between PCs and child mental health however cumulative PCs could be considered as a possible **prognostic** factor for child mental health. Our findings highlight the potential of the cumulative number of PCs experienced to be used to screen/identify children at-risk of poorer mental health outcomes who may benefit from early intervention.
- Although we know there is a relationship between **social-risk** and outcomes such as cognition in high-risk groups e.g. preterm birth (Beauregard et al., 2017), our findings highlight a relationship between social-risk and *mental health* outcomes at population-level, that is, for *all* children.
- **Economically**, the costs of intervening for children at-risk preventatively has been shown to be lower and require less intense treatment than the cost of providing interventions targeting children who already present in the clinical range (Campbell...& Pan, 2014). Screening for PCs and SR at baby developmental appointments may help identify vulnerable babies during a unique window of opportunity for early interventions promoting both maternal and child well-being **prior** to mental health difficulties being evident.
- It is important to note that although there was increased risk among children exposed to PCs, the majority of children (ranging from 80-95%) of mothers with PCs did not have mental health problems. Further research into determinants of such resilience might be important for designing interventions for children at-risk. Additionally, future research should consider cumulative exposures in pregnancy across **biopsychosocial** domains not just biological, psychological or social exposures independently.

# GOAL B

Optimum physical and mental health

# OBJECTIVE 6

Babies, young children and their parents enjoy positive mental health

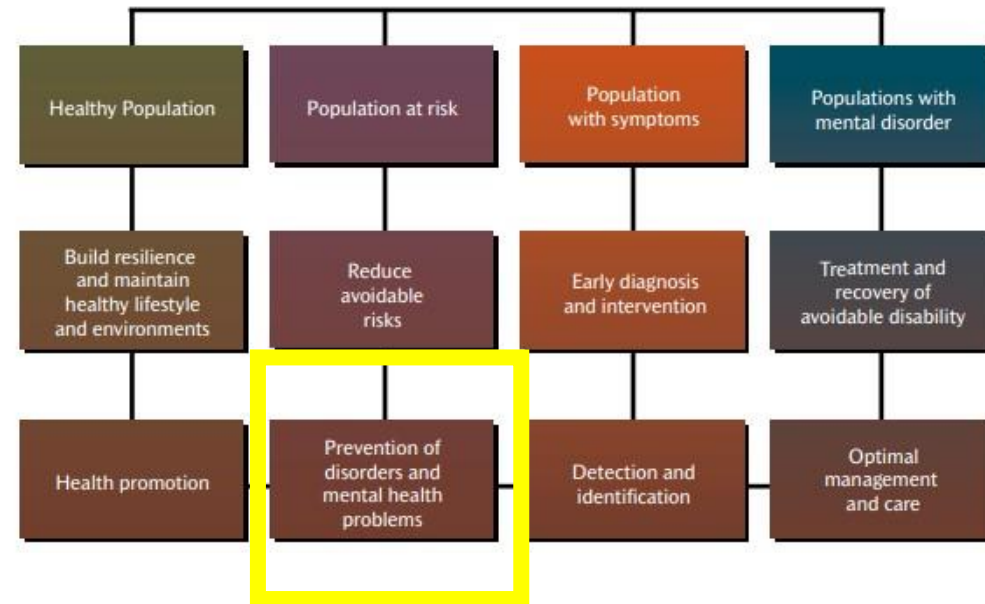


Strategic Action	Action	Lead and Partner(s)	Key Output by 2021
<b>6.1</b> Improve the early identification of mental health problems among babies, young children and families.	B.6.1.1. Provide information and guidance on promoting and supporting positive mental health among pregnant women, babies, young children and families and carry out a public information campaign to raise awareness of the importance of promoting social-emotional development, positive mental health and reducing the impact of early childhood trauma in babies and young children.	Lead: HSE Partner: DoH	Infant mental health promoted and supported through better information sources.  See also actions B.6.1.6 and B.6.2.6.
	B.6.1.2. Enhance efforts to promote positive mental health among pregnant women, babies, young children and their families in all resources and service contacts by extending the Making Every Contact Count: A Health Behaviour Change Framework and Implementation Plan for Health Professionals in the Irish Health Service to include mental health.	Lead: HSE	Scoping document to outline the expansion of Making Every Contact Count (MECC) to include the promotion of mental health and wellbeing as part of the revised implementation plan for the programme from 2021.



**Figure 5.1 Opportunities for mental health promotion: A population perspective**

*Adapted from Building capacity to promote mental health of Australians.<sup>66</sup>*



# References

- Beauregard et al (2017). Preterm birth, poverty and cognitive development. *Pediatrics*, 141(1).
- Buss & Genuneit (2023). How many are too much? Clinical and epidemiological considerations for the Era of Poly-, Pre- & postnatal exposures relevant for the developing brain. *Biological Psychiatry*, 93(10), 861-863.
- Campbell...& Pan (2014). Early childhood investments substantially boost adult health. *Science*, 343, 1478-85.
- Caspi et al (2020). Longitudinal assessment of mental health disorders and comorbidities in the Dunedin birth cohort study. *JAMA Network Open*, 3(4), e203221.
- Fuchs et al (2022). Early parenting stress links obstetric complications and child psychopathology in middle childhood in at-risk sample. *J of Dev and Beh Pediatrics*, 43(1), 48-55.
- Healy et al (2022). Person-Centered Trajectories of Psychopathology From Early Childhood to Late Adolescence. *JAMA Network Open*, 5(5). E229601
- HSE <https://www.hse.ie/eng/about/who/healthwellbeing/our-priority-programmes/child-health-and-wellbeing/nationalhealthychildhoodprogramme/>
- Laucht et al (2000). Behavioral sequelae of perinatal insults and early family adversity at 8 years of age. *J Am Acad Child Adolesc Psychiatry*, 39, 1229-1237.
- Lee et al(2020). Maternal Bacterial Infection During Pregnancy and Offspring Risk of Psychotic Disorders: Variation by Severity of Infection and Offspring Sex. *Am J Psychiatry*, 177(1), 66-75.
- Li et al (2020). Exploring the validity of allostatic load in pregnant women. *Midwifery*, 82, 102621.
- Long & Freese (2014). *Regression models for categorical dependent variables using STATA*. STATA press, 3<sup>rd</sup> ed.
- Mac Giollabhui et al. (2019). Maternal inflammation during pregnancy and offspring psychiatric symptoms in childhood: Timing and sex matter. *J Psychiatr Res*, 111, 96-103.
- Milberger et al (1997). Pregnancy, delivery and infancy complications and attention deficit hyperactivity disorder: issues of gene-environment interaction. *Biological Psychiatry*, 41(1), 65-75.
- Morales et al. (2023). Developmental trajectories of aggression, hyperactivity/inattention, and anxious depressed mood: co-occurring problems within a Chilean context. *Current Psychology*, doi: 10.1007/s12144-023-04530-3
- Nees et al. (2021). Prediction Along a Developmental Perspective in Psychiatry: How Far Might We Go? *Front Syst Neurosci*, 15, 670404.
- Nielsen et al. (2016). Individual and combined effects of maternal anemia and prenatal infection on risk for schizophrenia in offspring. *Schizophr Res*, 172(1-3), 35-40.
- Nomura et al. (2012). Exposure to gestational diabetes mellitus and low socioeconomic status: effects on neurocognitive development and risk of attention-deficit/hyperactivity disorder in offspring. *Arch Pediatr Adolesc Med*, 166(4), 337-43.
- Parboosing et al. (2013). Gestational influenza and bipolar disorder in adult offspring. *JAMA Psychiatry*, 70(7), 677-85.
- Robinson et al. (2021). Maternal Hypertensive Pregnancy Disorders and Mental and Behavioral Disorders in the Offspring: a Review. *Curr Hypertens Rep*, 23(5).
- Roffman et al (2021). Association of adverse prenatal exposure burden with child psychopathology in the Adolescent Brain Cognitive Development (ABCD) Study. *PLoS One*, 16(4), e0250235.
- Stroud, Mc Callum & Saisbury (2018). Impact of maternal prenatal smoking on fetal to infant neurobehavioral development. *Dev Psychopathol*, 30(3), 1087-1105.
- Wagner et al (2009). The limited effects of obstetrical and neonatal complications on conduct and attention-deficit hyperactivity disorder symptoms in middle childhood. *J Dev Behav Pediatr*, 30(3), 217-25.