1	Childhood physical maltreatment, perceived social isolation, and internalizing symptoms: A longitudinal,
2	three-wave, population-based study
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# 30 Abstract

31 A number of cross-sectional studies have consistently shown a correlation between childhood physical 32 maltreatment, perceived social isolation, and internalizing symptoms. Using a longitudinal, three-wave design, 33 this study sought to assess the mediating role of perceived social isolation in adulthood in the association 34 between childhood physical maltreatment and internalizing symptoms in adulthood. We used data collected 35 from 1994 to 2008 within the framework of the Tromsø Study (N=4,530), a representative prospective cohort 36 study of men and women. Perceived social isolation was measured at a mean age of 54.7 years, and internalizing 37 symptoms were measured at a mean age of 61.7 years. The difference-in-coefficients method was used to assess 38 the indirect effects and the proportion (%) of mediated effects. Childhood physical maltreatment was associated 39 with an up to 68% (relative risk [RR]=1.68, 95% confidence interval [CI]:1.33-2.13) higher risk of perceived 40 social isolation in adulthood. Childhood physical maltreatment and perceived social isolation in adulthood were 41 associated with greater levels of internalizing symptoms in adulthood (p<0.01). A dose-response association 42 was observed between childhood physical maltreatment and internalizing symptoms in adulthood (p < 0.001). Perceived social isolation in adulthood mediated up to 14.89% (p<0.05) of the association between childhood 43 44 physical maltreatment and internalizing symptoms in adulthood. The results of this study indicate the need to 45 take perceived social isolation into account when considering the impact of childhood physical maltreatment on 46 internalizing symptoms. 47 Keywords: psychological well-being; psychological distress; social support; anxiety; depression; childhood 48 abuse; emotional distress; loneliness 49 50 51 52 53 54 55 56 57 58

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# 60 Introduction

61 In recent decades, there has been a great amount of research on how childhood physical maltreatment relates to 62 affective and internalizing disorders in adulthood [1-4], and reviews [5-7] have shown reasonable associations 63 between these variables. Childhood physical maltreatment can have enduring effects on brain development and 64 brain stress regulatory flow systems, which may confer increased vulnerability to internalizing symptoms (i.e., 65 depressive and anxious symptomatology) in later life [8,9,3,1,2]. For instance, childhood physical maltreatment 66 may influence emotional abilities and alter sensory thresholds in ways that undermine effective emotion 67 regulation and create increased susceptibility to internalizing disorders in later life [10,11,4,1,12]. Other reports 68 suggest that childhood physical maltreatment influences internalizing symptoms through the social and 69 emotional impairments it provokes [13,4], such as poor social skills, impulsivity, reactive aggression, 70 behavioural problems, or excessive reassurance-seeking, as individuals with these impairments are likely to be 71 rejected by their normally-functioning peers [14,15,4,1]. 72 An individual's functional social behaviour depends on their ability, capacity, and motivation for social 73 interactions, all of which play an important role in maintaining a social support network and social relationships 74 with peers [4,1]. The biological embedding hypothesis suggests that the childhood physical 75 maltreatment -> internalizing symptoms association is partly driven by embedding or scarring that limits one's 76 ability to develop and maintain social relationships [16,17]. Several reports on the long-term influence of 77 childhood physical maltreatment have suggested it causes high emotional reactivity and decreased social 78 competency, which disturb an individual's ability to make and sustain supportive relationships, thereby 79 contributing to the development of internalizing symptoms [18,15,4,1]. Previous evidence has shown that poor 80 social conformity and negative interaction with friends in adulthood may be a consequence of childhood 81 physical maltreatment [18,10,1,4]. Specifically, childhood physical maltreatment is associated with deficits in 82 attention, mindfulness, and self-referential encoding [19,20,4]. For example, previous studies have indicated that 83 children who were physically maltreated showed errors in judgment in situations that required interpersonal 84 reasoning; in turn, these errors in judgement may lead to inappropriate social behaviour [21,22]. Deficits in 85 emotional self-regulation are also associated with externalizing behaviors, which in turn are associated with 86 peer-rejection [23,24,1,4,25]. Many of these symptoms do not occur in isolation, and most affect the quality and 87 quantity of the social bonds people cultivate in their lifetime [18,26,1,4]. For example, peers may perceive self-88 focused individuals as especially annoying and abrasive [27]. Impaired emotion recognition and a negative 89 emotional bias could contribute further to internalizing symptoms via deficits in receptive communication

The social support deterioration model [29] suggests that stressors such as childhood physical
maltreatment can damage a child's perception of available and helpful social support, which, in turn, can lead to
maladjustment [30,26,4]. Indeed, maltreated children may perceive their social network in a negative manner
[31,14,1,4]. Even when this network is based on mutual giving and receiving, they may not be able to perceive
this because of the difficulties they often face in interpreting the thoughts and feelings of others [31,14,1,4].
They are also less likely to be able to understand negative emotions such as anger and sadness, which can affect
the quantity and quality of the social relationships they maintain throughout their life [31,14,1,4].

99 The vulnerability-stress model suggests that childhood physical maltreatment causes heightened 100 sensitivity to subsequent stressors [32,33,4]. Maltreated children may be more sensitive when it comes to 101 detecting and perceiving threats to their security or cues of rejection [22]. As a result, they may perceive the 102 world as more socially threatening, making them avoid relationships with peers in order to protect themselves 103 from anticipated disappointment [34,20,35]. Sensitivity to peer rejection might result in more acute emotional 104 responses and a more negative interpretation of peer rejection due to an inability to properly regulate the 105 emotions that result from such social encounters [36,28].

106 The concept of 'loneliness loop', proposed by Hawkley and Cacioppo [37], may explain the association 107 between perceived social isolation and internalizing symptoms. Perceived social isolation refers to the 108 distressing feeling that occurs when there are discrepancies between one's desired and actual quantity of social 109 relationships [29,38,39]. Perceived social isolation can lead to dismissive or avoidant behaviour in social 110 relationships, because affected individuals have an expectation of negative social interactions and social 111 rejection, making them withdraw socially [37,40]. When combined with negative social expectations, this 112 inability to develop and maintain social relationships can elicit behaviours from others that confirm these 113 expectations [37]. Thus the self-reinforcing 'loneliness loop' may contribute to internalizing symptoms through 114 feelings of rejection, hostility, and pessimism [34,37,41].

Several studies [1,42-45,34,46,31,47-53,4] have assessed the mediating role of perceived social
isolation in the association between childhood physical maltreatment and internalizing symptoms in adulthood.
However, they had several limitations. First, assessing mediation with cross-sectional studies can lead to
questionable inferences [2,54], as studies have shown reciprocal associations between perceived social isolation

and internalizing symptoms [55,38,56]. Studies with dense temporal sampling are also vulnerable to these biases [54]. A potential solution for establishing temporal order between perceived social isolation and internalizing symptoms is a long follow-up period [54,4]. Other studies have shown that the association between childhood physical maltreatment and internalizing symptoms is over-estimated in cross-sectional studies due to differential misclassification error [2]. Despite this, most previous studies [43,42,1,45,34,47-49,51,31,52] have assessed the association between childhood physical maltreatment and internalizing symptoms, or the association between perceived social isolation and internalizing symptoms, at the same time point.

Second, most studies [43-46,48,50-53] based their conclusions on very small and selective samples, 126 127 which makes it impossible to generalise the findings to the general population. Finally, several studies [46,31,47,48,50-52] did not present indirect effect estimates and corresponding confidence intervals. Other 128 studies [57,58] that assessed the associations between childhood physical maltreatment, perceived social 129 130 isolation, and internalizing symptoms in adulthood did not consider the mediating role of perceived social 131 isolation in adulthood in the analysis. 132 A review of the existing literature indicated that the mediating role of perceived social isolation in 133 adulthood in the association between childhood physical maltreatment and internalizing symptoms in adulthood 134 has not been assessed in a thoroughly comprehensive and systematic manner with a large and representative

study sample [4]. Therefore, using longitudinal data from the Tromsø Study, the aim of the present study was to
assess the mediating role of perceived social isolation in adulthood in the association between childhood
physical maltreatment and internalizing symptoms in adulthood.

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#### 149 Material and Methods

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### 151 Study sample

152 The Tromsø Study is a longitudinal prospective cohort study and its participants are considered representative of

the adult population residing in the municipality of Tromsø [59]. With more than 60,000 inhabitants, Tromsø is

the largest city in Northern Norway. The present study has a three-wave design and includes data collected from

155 1994 to 2008 [59,4,3]. To be eligible for the present analyses, participants had to have participated in all of the

156 following surveys: Tromsø IV (1994-95), Tromsø V (2001-02), and Tromsø VI (2007-08) (N=4,530) [3,4]. The

157 exposure (childhood physical maltreatment) was measured retrospectively in 2007-08, the mediator (perceived

social isolation in adulthood) was measured in 1994-95, and the outcome (internalizing symptoms in adulthood)

was measured in 2001-02 [4]. The limitations of the study design are discussed at the end of this article.

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#### 161 Ethical approval

162 This investigation was carried out in accordance with the latest version of the Declaration of Helsinki. The

163 Tromsø Study has been approved by the Regional Committee for Medical and Health Research Ethics, the Data

164 Inspectorate, and the Norwegian Directorate of Health. Written informed consent was obtained from all

- 165 individual participants included in the study.
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## 167 Study variables

### 168 Exposure (childhood physical maltreatment)

169 Self-reported information on childhood physical maltreatment was measured retrospectively in the Tromsø VI

170 questionnaire by two questions [1,2,9,3,4]: "Have you over a long period experienced any of the following as a

171 child?". The possible responses were: i) 'Being tormented, or threatened with violence'; and ii) 'Being beaten,

172 kicked, or the victim of other types of violence'. Using responses to both questions, a separate cumulative

- variable of childhood physical maltreatment was constructed as: 0=not exposed to childhood physical
- 174 maltreatment, 1= exposed to one childhood physical maltreatment (some), and 2= exposed to both childhood
- 175 physical maltreatments (severe) [2,9]. The internal reliability of childhood physical maltreatment was good in

the Tromsø Study [9].

# 177 Mediator (perceived social isolation in adulthood)

Perceived social isolation (i.e., a subjective judgement of the adequacy of social relationships) was measured in
Tromsø IV (mean age: 54.7 years) [4,3]. The operational definition of perceived social isolation in this study
was the perception that one's social needs were not met by the quantity of good friends [4]. Perceived social
isolation was measured in the questionnaire by the question "Do you feel that you have enough good friends?"
(yes=0, no=1) [4,3]. Those responding 'no' were categorised as socially isolated, while those responding 'yes'
were categorised as *not* socially isolated [4,3]. We recognize that this is a weak measure; the implications of this

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# 186 *Outcome (internalizing symptoms in adulthood)*

187 Depressive and anxious symptomatology over the last two weeks were considered as indicators of internalizing 188 symptoms [1-4]. The Hopkins Symptom Checklist (HSCL-10) scale in the Tromsø V questionnaire (mean age: 189 61.7 years) was used to measure internalizing symptoms, which is widely used in epidemiological studies [1-4]. 190 Respondents rated each of the 10 items in the HSCL-10 on a four-point scale, ranging from not at all (1) to 191 extremely (4) [1-4]. The HSCL-10 had an acceptable degree of internal consistency in this sample (Cronbach's 192 alpha: 0.86, mean inter-item correlation: 0.42, McDonald's omega coefficient for composite reliability: 0.87) [1-193 4]. An HSCL-10 score between 1.0 and 4.0 was assigned by dividing the total score (sum of the 10 indicators) 194 by 10, where 4 represented the highest and 1 represented the lowest score on internalizing symptoms [3] (mean: 195 1.22, standard deviation [SD]: 0.33).

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## 197 Confounding variables (Tromsø IV)

The associations between childhood physical maltreatment, perceived social isolation, and internalizing symptoms in adulthood are likely confounded by age, gender, living in Norway at age 1 year (yes, no), childhood financial conditions, and mother's/father's history of psychiatric disorders [60,61,1,2,9,4,3]. Previous studies have indicated that parental psychopathology may partially explain the association between childhood maltreatment and internalizing symptoms [62,9]. Being raised in an environment with ample monetary and parental resources may expose children to experiences that may aid in development of capacities, skills, and connections that are conducive to social engagement and forming relationships [2,63,64,4].

Valid information on age and gender was obtained from Statistics Norway by using the unique personal
identification number of each participant. Mother's/father's history of psychiatric disorders was measured as:
'Does your mother/father have/has or your mother/father ever had psychiatric disorders? (yes, no) [3,4]. The

234	physical maltreatment on internalizing symptoms in adulthood
233	Assessing direct and indirect effects (through perceived social isolation in adulthood) of childhood
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231	robust error variance. OLS estimates ( $\beta$ ) and 95% CIs are presented.
230	internalizing symptoms in adulthood was assessed by ordinary least square (OLS) regression analysis with a
229	confidence intervals (CIs) are presented. The association between perceived social isolation in adulthood and
228	assessed by Poisson regression analysis with a robust error variance [54]. Relative risks (RRs) and 95%
227	The association between childhood physical maltreatment and perceived social isolation in adulthood was
226	perceived social isolation, age and gender were observed (regressed on internalizing symptoms) in this sample.
225	No statistically significant multiplicative interactions between childhood physical maltreatment,
224	presented.
223	statistical analyses were performed on the MI datasets and both unadjusted (crude) and adjusted estimates are
222	missing) and the MI datasets is presented with proportions (%), and mean (standard error) (Table 1). All
221	HSCL-10 were included in the imputation models. A comparison between the complete-case (excluding
220	imputation procedure. In order to increase the predictive power of the imputation procedure, all indicators of
219	with missing data [54]. One hundred MI datasets were generated to help account for the uncertainty in the
218	(MI) with chained equations to avoid any bias in the associations of interest introduced by excluding individuals
217	All analyses were conducted using Stata version 15. Imputed values were generated with multiple imputation
216	Statistical analysis
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214	[3,4].
213	reliability of childhood financial conditions was good (Kappaweighted: 0.61, 95% CI: 0.59-0.63) in this sample
212	on a 4-point scale (1=very good, 2=good, 3=difficult, 4=very difficult) [64,1,2,9,3,54,4]. The test-retest
211	measured retrospectively by the question: "How was your family's financial situation when you were a child?"
210	The variable childhood financial conditions was used as a proxy for childhood socioeconomic status, and was
209	this sample were Kappa: 0.57 (95% CI: 0.52–0.62) and Kappa: 0.61 (95% CI: 0.53–0.69), respectively [3,4].
208	test-retest reliability of mother's history of psychiatric disorders and father's history of psychiatric disorders in

235 The association between childhood physical maltreatment and internalizing symptoms in adulthood was

assessed by OLS regression analysis. Mediation analysis with the difference-in-coefficients method [54] was

used. Perceived social isolation in adulthood was included in the models to assess the indirect effects and the

238	proportion of mediated effects (%) [4,54]. If perceived social isolation in adulthood is an important mediator of
239	the childhood physical maltreatment $\rightarrow$ internalizing symptoms in adulthood associations, the effects of
240	childhood physical maltreatment ( $\beta_{Total Effect}$ ) should decline when it is added to the regression model [4,54].
241	Four estimates are presented: total effects (adjusted for confounding variables), direct effects (adjusted for
242	confounding variables and perceived social isolation in adulthood), indirect effects (difference between total
243	effect and direct effect) [54], and proportion mediated (%) [2]. Standard errors were derived with bias-corrected
244	bootstrapping [54] for hypothesis testing, and 95% CIs are presented.
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### 268 Results

269 Compared to the respondents that did not participate in all three considered waves of the Tromsø Study (Tromsø 270 IV, Tromsø V, and Tromsø VI), the respondents in this study sample were likely to be female and older 271 (p<0.05). Among the 4,530 individuals in this study sample, the majority were aged 55 years or older (56.2%) at 272 baseline (Tromsø IV) and 59.2% were women. Mother's history of psychiatric disorders was reported more 273 frequently than father's history of psychiatric disorders (5.8% and 2.0%, respectively). The majority (60.3%) of 274 the study sample reported having good or very good financial conditions in childhood; 4.9% reported some (any 275 type) childhood physical maltreatment, and 2.6% reported severe (both) childhood physical maltreatment (Table 276 1). A substantial proportion (16.1%) of respondents reported perceived social isolation in adulthood (Table 1).

277 As a first analytic step in testing direct and indirect effects, the associations between childhood physical 278 maltreatment, perceived social isolation in adulthood, and internalizing symptoms in adulthood were assessed. 279 In the fully-adjusted model, exposure to some (any type) childhood physical maltreatment was associated with a 280 68% (RR<sub>adjusted</sub>=1.68, 95% CI: 1.33-2.13) higher risk of perceived social isolation in adulthood, while exposure to severe (both) childhood physical maltreatment was associated with a 57% (RR<sub>adjusted</sub>=1.57, 95% CI: 1.12-281 282 2.21) higher risk (Table 2). In turn, perceived social isolation in adulthood was associated with greater levels  $(\beta_{adjusted}=0.14, p<0.001)$  of internalizing symptoms in adulthood (Table 3). 283 284 285 Direct and indirect effect of childhood physical maltreatment on internalizing symptoms in adulthood

A dose-response association was observed between childhood physical maltreatment and internalizing symptoms in adulthood (p<0.001). After controlling for confounding variables, exposure to some or severe childhood physical maltreatment was associated with greater levels of internalizing symptoms in adulthood (p<0.001) (Table 4). Decomposition of total effects showed that there was a direct and an indirect effect of childhood physical maltreatment on internalizing symptoms in adulthood (p<0.05). Perceived social isolation in adulthood mediated 7.25-14.89% (p<0.05) of the association between childhood physical maltreatment and internalizing symptoms in adulthood (Table 4).

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# 298 Discussion

In this study, we examined the relationship between childhood physical maltreatment, perceived social isolation, and internalizing symptoms in a large, representative, and population-based sample of Norwegian adults. Childhood physical maltreatment was associated with an increased risk of perceived social isolation, and both childhood physical maltreatment and perceived social isolation were associated with greater levels of internalizing symptoms. The results of this study showed that perceived social isolation mediates the association between childhood physical maltreatment and internalizing symptoms.

305 It must be noted that perceived social isolation as conceptualized in this study does not refer to actual 306 social isolation [4]. Indeed, maltreated individuals may perceive social isolation even when social opportunities 307 and relationships exist, because they lack the capacity to utilize these resources [40,4,1]. Accordingly, the results of this study may not support the conclusion that "social support" mediates the childhood physical 308 309 maltreatment-internalizing symptoms association. Indeed, in a recent longitudinal population-based study [4], 310 "number of friends in adulthood" mediated only 3% of the association between childhood adversity and 311 internalizing symptoms in adulthood. The results of this study suggest that respondents with a history of 312 childhood physical maltreatment may develop internalizing symptoms in adulthood partly because they feel 313 dissatisfied or unfulfilled by the social connections they have, and not necessarily because they have fewer 314 social contacts or sources of support per se [65,4].

315 The stress buffering model and the stress process model suggest that perceived social isolation may 316 mitigate the psychological impact of childhood physical maltreatment on internalizing symptoms by attenuating 317 the stress appraisal response [66,67]. However, no statistically significant interaction was observed between 318 childhood physical maltreatment and perceived social isolation (data not shown), which is in agreement with 319 some [1,48,68], though not all [44,46,58] previous studies. Age and childhood financial conditions did not play 320 a moderating role in the association between childhood physical maltreatment and internalizing symptoms, 321 which is consistent with most [42,1,2,9], though, not all [69] previous studies. Some studies [34] have used 322 confounding variables to control for the association between this exposure (childhood maltreatment) and 323 outcome (internalizing symptoms in adulthood), but without adjusting for the association between the exposure 324 and the mediator (perceived social isolation), or between the mediator and the outcome. It is well established 325 that the decomposition of total effects into direct and indirect effects assumes no unmeasured or unaccounted-326 for exposure-mediator, exposure-outcome, or mediator-outcome confounding [2,3,54]. Consistent with previous studies, childhood physical maltreatment was associated with perceived social isolation [1,4]. Finally, consistent
 with most [1,34,47,44], though not all [31,49,53,45] previous studies, perceived social isolation mediated the
 association between childhood physical maltreatment and internalizing symptoms.

330 These results should be interpreted in the context of some limitations. Measurement of childhood 331 physical maltreatment, perceived social isolation, and internalizing symptoms relied on self-reports. Childhood 332 physical maltreatment was measured retrospectively with two items, and this occurred 13 years after the 333 measurement of perceived social isolation and 6 years after the measurement of internalizing symptoms. 334 Therefore, the possibility that perceived social isolation and chronic internalizing symptoms affected the recall 335 of childhood physical maltreatment is a potential concern for the interpretation of these findings. However, the 336 prevalence of childhood physical maltreatment in this study (4.9%) corresponds with that reported in other studies from Norway [1,70-72]. Misclassification of childhood physical maltreatment maybe influenced by age, 337 338 state of mind, and current psychopathology [73,2,9,74]; however, other evidence suggests that these biases 339 should be fairly low [75-77,9,78-81]. Moreover, some other reports have suggested that subjects under-report 340 childhood physical maltreatment when it is measured retrospectively [82,83]. Recent studies that compared 341 associations of retrospective and prospective assessments of childhood physical maltreatment and health have shown that the associations remained in the same direction; however, the associations between retrospective 342 343 childhood physical maltreatment and self-reported outcomes were over-estimated [74,84]. Accordingly, since 344 the measurement of internalizing symptoms in this study was self-reported, it is plausible that the total effects 345 and direct effects presented here are over-estimated, while the indirect effects may be under-estimated. Another 346 limitation of the study is that the precise timing or persistence of childhood physical maltreatment was not 347 measured [9,2]. The use of a single binary variable for perceived social isolation has methodological and 348 conceptual limitations [4,44,3]. Although the results of this study are consistent with most previous reports, 349 potential misclassification of perceived social isolation should be considered [2,9,54,44]. For instance, non-350 differential misclassification of perceived social isolation would lead to an under-estimation of indirect effect 351 estimates (biased downwards); however, differential misclassification of perceived social isolation would lead to 352 an over-estimation of indirect effect estimates (biased upwards) [2,54,4]. Although perceived social isolation 353 was measured approximately 7 years earlier than internalizing symptoms, it is still possible that some 354 individuals might have been suffering from chronic internalizing disorders when they participated in Tromsø IV. 355 Indeed, history of internalizing symptoms is a strong predictor of internalizing symptoms in later life [85-87]. 356 However, internalizing symptoms were not measured in Tromsø IV, and perceived social isolation was not

357 measured in Tromsø V; therefore, the change in the mediator (perceived social isolation) or outcome 358 (internalizing symptoms) was not assessed [3,4]. It is plausible that childhood physical maltreatment may have 359 led to the development of internalizing symptoms in young adulthood, which in turn could have compromised 360 the self-reports of perceived social isolation measured later (i.e., those suffering from chronic internalizing 361 disorders could also be more likely to perceive social isolation later in life). Other evidence indicates that 362 respondents with a history of maltreatment attribute sadness to both positive and negative social situations, which may affect their social skills over the life course [88]. Although childhood physical maltreatment was the 363 364 focus of the study, we cannot rule out that exposure to adverse events later in life (concern for intermediate 365 confounding [2,9,54]) might also have affected perceived social isolation and internalizing symptoms in 366 adulthood [1,4]. Sociable disposition is, at least in part, based on genetic factors [89,90]. Indeed, a substantial 367 proportion of one's risk of internalizing symptoms is also genetically determined [91,92,3]. The test-retest 368 reliability of self-reported parental history of psychiatric disorders was not high, which raises the concern that 369 we were not able to fully control for genetic dispositions [4]. However, other evidence suggests that childhood 370 physical maltreatment is associated with internalizing symptoms in childhood [93] and adulthood [2,3], 371 independent of genetic dispositions.

The strengths of this study are its three-wave design, and a representative sample of the adult population of Tromsø. By determining childhood physical maltreatment, perceived social isolation in adulthood, and internalizing symptoms in adulthood at different time points, spurious associations due to state of mind and mood congruency bias were avoided [2,9,54]. Finally, missing values were imputed, which avoids the possibility that attenuations in the coefficient for childhood physical maltreatment were due to selection bias rather than mediation [9,54,4].

The results of this study indicate the need to take perceived social isolation over the life course into account when considering the long-term impact of childhood physical maltreatment on internalizing symptoms in adulthood [1,4]. As the mean age of the respondents was 61.7 years (median: 63) at the time internalizing symptoms were assessed (in Tromsø V), the associations between childhood physical maltreatment and internalizing symptoms in adulthood suggests that the consequences of childhood physical maltreatment persists for several decades after its occurrence [94,9,1,2]. In summary, a dose-response relationship [9,1] was observed between childhood physical maltreatment and

internalizing symptoms, and this association appeared to be mediated substantially by perceived social isolation.

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Table 1. General characteristics of the study sample (n=4,530).

υ	5	2
6	5	3

	Characteristics	Complete- case dataset	Imputed dataset
		n (%)	(%)
Age (in 1994)	Mean (standard error,	54.69 (0.15)	_b
	SE)		
	25-34	302 (6.7)	_b
	35-44	352 (7.8)	_b
	45-54	1327 (29.3)	_b
	55-64	1852 (40.8)	_b
	65-74	697 (15.4)	_b
Gender	Male	1849 (40.8)	_b
	Female	2681 (59.2)	_b
History of psychiatric disorders, mother	Yes	262 (5.8)	_b
	No	4268 (94.2)	_b
History of psychiatric disorders, father	Yes	90 (2.0)	_b
	No	4440 (98.0)	_b
Living in Norway at age 1year <sup>a</sup>	Yes	4081 (98.4)	98.4
	No	66 (1.6)	1.6
Childhood financial conditions <sup>a</sup>	Mean (SE)	2.40 (0.01)	2.40 (0.01)
	Very good	120 (2.8)	2.8
	Good	2437 (57.5)	57.5
	Difficult	1547 (36.5)	36.5
	Very difficult	136 (3.2)	3.2
Childhood physical maltreatment	None	4198 (95.1)	_b
	Some	218 (4.9)	_b
	Severe	114 (2.6)	_b
Perceived social isolation	Not socially isolated	3485 (83.9)	83.9
	Socially isolated	669 (16.1)	16.1
Internalizing symptoms (HSCL-10) <sup>a</sup>	Mean (SE)	1.22 (0.01)	1.27 (0.01)

<sup>a</sup>The numbers for some variables do not add up to 4,530 due to missing values.

<sup>b</sup>There were no missing values, so no imputations were made for these variables.

656 SE: standard error; HSCL-10: Hopkins Symptom Check List-10; scale (1.0–4.0), where 1.0 represents lowest

657 score on internalizing symptoms, and 4.0 represents highest score on internalizing symptoms.

Table 2. Association between childhood physical maltreatment and perceived social isolation in adulthood (n=4,530).

		Perceived social isolation			
		(	Crude	Ad	ljusted
		RR	95% CI	RR <sup>a</sup>	95% CI
Childhood physical maltreatment	None (ref)	1.00 (ref)		1.00 (ref)	
	Some	1.85 <sup>b</sup>	1.47-2.32	1.68 <sup>b</sup>	1.33-2.13
	Severe	1.78 <sup>b</sup>	1.29-2.47	1.57°	1.12-2.21

<sup>a</sup>Adjusted for age, gender, childhood financial conditions, living in Norway at age 1 year, and mother's/father's history of psychiatric disorders.

<sup>b</sup> p<0.001 <sup>c</sup> p<0.01

RR: relative risk; CI: confidence interval.

		Internalizing symptoms			
		Crude Adjusted			
		β <sup>b</sup>	95% CI	β <sup>a,b</sup>	95% CI
Perceived social isolation	Not socially isolated	reference		reference	
	Socially isolated	0.14	0.10-0.18	0.14	0.11-0.18

Table 3. Association between perceived social isolation and internalizing symptoms in adulthood (n=4,530).

<sup>a</sup>Adjusted for age, gender, childhood financial conditions, childhood physical maltreatment, living in Norway at age 1 year, and mother's/father's history of psychiatric disorders. <sup>b</sup>p<0.001

CI: confidence interval; HSCL-10: Hopkins Symptom Check List-10; scale (1.0–4.0), where 1.0 represents lowest score on internalizing symptoms, and 4.0 represents highest score on internalizing symptoms. Perceived social isolation was measured in 1994-95, while internalizing symptoms was measured in 2001-2002.

Table 4. Direct and indirect effect (mediated through perceived social isolation) of childhood physical maltreatment on internalizing symptoms in adulthood (n=4,530).

		Internalizing symptoms					
		Total effect <sup>a</sup> Direct effect <sup>b</sup> Indirect effect <sup>a</sup> Proportion mediated <sup>a</sup>					
		β (95% CI)	β (95% CI)	β (95% CI)	% (95% CI)		
Childhood physical maltreatment	Some <sup>c</sup>	0.11 (0.09-0.12) <sup>d</sup>	0.09 (0.09-0.11) <sup>d</sup>	$0.02 (0.01 - 0.02)^d$	14.89 (5.81-16.86) <sup>d</sup>		
	Severe <sup>c</sup>	0.20 (0.15-0.20) <sup>d</sup>	0.18 (0.13-0.18) <sup>d</sup>	$0.01 (0.01 - 0.02)^d$	7.25 (6.59-15.93) <sup>e</sup>		

<sup>a</sup>Adjusted for age, gender, childhood financial conditions, living in Norway at age 1 year, and mother's/father's history of psychiatric disorders.

<sup>b</sup>Adjusted for age, gender, childhood financial conditions, living in Norway at age 1 year, and mother's/father's history of psychiatric disorders + perceived social isolation in adulthood.

<sup>c</sup>Reference: None

<sup>d</sup> p<0.001

<sup>e</sup>p<0.05

CI: confidence interval; HSCL-10: Hopkins Symptom Check List-10; scale (1.0–4.0), where 1.0 represents lowest score on internalizing symptoms, and 4.0 represents highest score on internalizing symptoms.