

1 Childhood maltreatment and eating disorder 2 pathology: a systematic review and dose-response 3 meta-analysis

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11 **Background.** Meta-analyses have established a high prevalence of childhood maltreatment (CM) in patients with eating
12 disorders (EDs) relative to the general population. Whether the prevalence of CM in EDs is also high relative to that in
13 other mental disorders has not yet been established through meta-analyses nor to what extent CM affects defining fea-
14 tures of EDs, such as number of binge/purge episodes or age at onset. Our aim is to provide meta-analyses on the asso-
15 ciations between exposure to CM (i.e. emotional, physical and sexual abuse) on the occurrence of all types of EDs and its
16 defining features.

17 **Method.** Systematic review and meta-analyses. Databases were searched until 4 June 2016.

18 **Results.** CM prevalence was high in each type of ED (total $N = 13\,059$, prevalence rates 21–59%) relative to healthy ($N =$
19 $15\,092$, prevalence rates 1–35%) and psychiatric ($N = 7736$, prevalence rates 5–46%) control groups. ED patients reporting
20 CM were more likely to be diagnosed with a co-morbid psychiatric disorder [odds ratios (ORs) range 1.41–2.46, $p < 0.05$]
21 and to be suicidal (OR 2.07, $p < 0.001$) relative to ED subjects who were not exposed to CM. ED subjects exposed to CM
22 also reported an earlier age at ED onset [effect size (Hedges' g) = -0.32 , $p < 0.05$], to suffer a more severe form of the ill-
23 ness ($g = 0.29$, $p < 0.05$), and to binge-purge ($g = 0.31$, $p < 0.001$) more often compared to ED patients who did not report
24 any CM.

25 **Conclusion.** CM, regardless of type, is associated with the presence of all types of ED and with severity parameters that
26 characterize these illnesses in a dose dependent manner.

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28 **Key words:** Anorexia, bulimia, childhood abuse, eating disorders, meta-analysis.

29 Introduction

30 The Diagnostic and Statistical Manual of Mental
31 Disorders 4th edition (DSM-IV-TR; APA, 2000) spe-
32 cified only two types of eating disorders (EDs): anor-
33 exia nervosa (AN) and bulimia nervosa (BN), and a
34 residual category ED not otherwise specified
35 (EDNOS), which included binge-eating disorder
36 (BED) for which in DSM-IV a criteria set has been pro-
37 posed for further study. In the DSM-5 (APA, 2013),
38 BED has been added as an official new diagnostic cat-
39 egory and besides for BED the EDNOS category also

made way for two new residual categories: (1) other 40
specified feeding or ED, including for instance sub- 41
threshold BN and (2) unspecified feeding or ED. The 42
lifetime prevalence of eating disorders according to 43
DSM-5 criteria is estimated to be up to 4% for AN, 44
2% for BN, and 2% for BED (Smink *et al.* 2013). 45

EDs are associated with co-morbidity of other men- 46
tal disorders, high mortality rates and high costs of 47
treatment (Fairburn & Harrison, 2003; Arcelus *et al.* 48
2011; Mischoulon *et al.* 2011; Mitchell *et al.* 2012; 49
Smink *et al.* 2013; Yao *et al.* 2016). Their etiology is com- 50
plex. Heritability estimates for EDs vary between 40% 51
and 60% (Trace *et al.* 2013). Non-shared environmental 52
risk factors such as childhood maltreatment (CM) also 53
contribute to the risk of developing an ED, according 54
to some, an estimated 17–46% (Klump *et al.* 2002). 55

Dozens of studies have been published on the asso- 56
ciation between CM, usually defined as self-reported 57

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58 exposure to emotional abuse or neglect and physical
59 and/or sexual abuse before the age of 18 years, and
60 ED pathology. Among these studies are four
61 meta-analyses (Rind *et al.* 1998; Smolak & Murnen,
62 2002; Chen *et al.* 2010; Caslini *et al.* 2016) and some nar-
63 rative reviews (Kent & Waller, 2000; Brewerton 2007;
64 Röhr *et al.* 2015). The data pooled in the meta-analyses,
65 the highest level of evidence available (Haidich, 2010),
66 clearly shows a relationship between CM and the pres-
67 ence of an ED diagnosis, although heterogeneity in the
68 magnitude of effect-size estimates is apparent (Smolak
69 & Murnen, 2002).

70 However, the meta-analytical results published so
71 far are rather specific, as they pertain to college sam-
72 ples only (Chen *et al.* 2010) or sexual abuse only
73 (Smolak & Murnen, 2002; Chen *et al.* 2010). The most
74 recent meta-analysis summarized the associations
75 between all CM types and all ED types up to
76 January 2014, but did not include a psychiatric control
77 group nor assessed the effect of CM on psychiatric
78 co-morbidity or other severity parameters (Caslini
79 *et al.* 2016). What is lacking is an in-depth overview
80 that summarizes all CM types and all ED types and
81 compares the prevalence not only with a healthy con-
82 trol group, but also with a control group with other
83 mental disorders than EDs. Moreover, from a clinical
84 perspective it is highly relevant to elucidate to what
85 extent CM affects the defining features of EDs, such
86 as its severity or age at onset since these define disease
87 to a large extent (Fairburn & Harrison, 2003; Smink
88 *et al.* 2013), beyond only the presence of a diagnostic
89 entity. To date, this has not been subjected to system-
90 atic review.

91 The aim of this study hence is to provide a system-
92 atic overview on the associations between exposure
93 to CM (i.e. emotional, physical and sexual abuse) on
94 the occurrence of all types of EDs and some of its
95 defining features such as severity and age at onset.

96 Method

97 The methodology that we used adhered to the guide-
98 lines that are recommended by the preferred reporting
99 items for systematic reviews and meta-analyses state-
100 ment (Moher *et al.* 2009).

101 Search strategy

102 The electronic databases PUBMED, PsycINFO, and
103 EMBASE were searched, up to 4 June 2016, to identify
104 relevant articles. The broad set of search terms can be
105 found in section A of the Supplementary material.
106 Additionally, we checked the references that were
107 made to the two seminal papers on the association of
108 interest (Oppenheimer *et al.* 1985; Root & Fallon,

1988) and reviewed the reference lists of the identified 109
articles. All searches, from identification to decision on 110
inclusion, were performed by at least two independent 111
reviewers (M.L.M., M.vS., M.O., V.K.). Inconsistencies 112
between reviewers were resolved through consensus. 113

Inclusion and exclusion criteri

114
115 This review included human studies (all age groups) 115
that assessed the presence of a current or lifetime ED 116
as defined by the DSM (APA, 1980, 2000, 2013) or 117
the International Classification of Diseases (ICD; 118
WHO, 2016a) and CM (before the age of 18 years). 119
We largely followed the definition and categorization 120
of the WHO in what we considered as exposure to 121
CM. The WHO (2016b) distinguishes five subtypes of 122
CM: physical CM, sexual CM, neglect, emotional CM 123
and exploitation. However, given that neglect and 124
emotional abuse are often reported as one entity we 125
decided to pool these categories as 'emotional CM'. 126
Furthermore, since exploitation is hardly a topic in 127
the ED literature, we decided to not include this cat- 128
egory in our meta-analysis. A general exclusion crite- 129
rion was the use of non-diagnostic, quantitative 130
assessment of eating behavior (e.g. disordered eating 131
assessed through a questionnaire). Studies had to be 132
published in English, French, German, Spanish or 133
Dutch in order to be included. 134

Data extraction

135
136 Data that were extracted included: demographic and 136
methodological characteristics (e.g. mean age of the 137
samples that were used and study design), CM preva- 138
lence by diagnostic group, i.e. AN, BN, BED, EDNOS, 139
healthy controls (HCs) and psychiatric controls (PCs) 140
and type of CM (i.e. reported emotional, physical, 141
and sexual abuse), CM characteristics (i.e. type, mean 142
age, assessment method), and ED characteristics (i.e. 143
type, mean age at onset, the number of self-reported 144
binge and purge episodes per week/month, the sever- 145
ity of the ED, assessed for instance by means of the eat- 146
ing attitudes test (Mintz & O'Halloran, 2000), the 147
eating disorder examination (Luce *et al.* 2008), or 148
the eating disorder inventory (Wear & Pratz, 1987), 149
the self-reported mean severity of depressive/anxiety 150
symptoms, psychiatric co-morbidity, suicidal behavior 151
or self-harm, and inpatient *v.* outpatient status) when 152
provided. We contacted by email the corresponding 153
authors of papers who we expected to have data that 154
would fit our purposes, but who did not provide the 155
data in their paper, and requested them to provide 156
us with the necessary data. Data extraction was per- 157
formed independently by at least two of us (M.L.M., 158
M.vS., M.O., V.K.). Inconsistencies were resolved in 159
consensus meetings. 160

161 **Quality assessment**

162 Based on the Newcastle-Ottawa scale (Wells *et al.* 2016)
 163 the methodological quality of the included studies was
 164 assessed. This scale is recommended by the Cochrane
 165 collaboration (Cochrane Community Handbook,
 166 2016). Detailed information on quality assessment is
 167 provided in Supplementary Tables S1 and S2.

168 **Data analyses**

169 Data analyses were performed in Stata version 13
 170 (StataCorp, 2013). All associations were tested for stat-
 171 istical significance at a confidence interval (CI).

172 First, the prevalence rates and 95% CI of CM (by
 173 type) for all EDs and for HCs and PCs were summar-
 174 ized over studies, regardless whether they included a
 175 HC and/or PC reference group. Next, pooled effect-size
 176 estimates were calculated on the association between
 177 CM (coded as yes *v.* no exposure) and the presence
 178 of an ED using data from studies that contained an
 179 ED sample and a HC and/or PC reference group.
 180 Within ED samples pooled effect-size estimates were
 181 calculated on the association between reported CM
 182 exposure and age at onset and severity of the ED, self-
 183 reported number of binge or purge episodes per week/
 184 month, severity of depressive and anxiety symptoms,
 185 psychiatric co-morbidity, suicidal behavior, and
 186 inpatient *v.* outpatient status.

187 The random-effects model, a model that includes
 188 both sampling- and study-level error (Borenstein *et al.*
 189 2009) was adopted in all instances. As effect-size mea-
 190 sures we chose to use odds ratios (ORs) for dichotom-
 191 ous outcome variables and Hedges' *g* for continuous
 192 outcome variables (Hedges & Olkin, 1985).

193 To assess the sensitivity of our results, all analyses
 194 were repeated with the data split-up as a function of
 195 type of childhood maltreatment (i.e. sexual, physical,
 196 and emotional maltreatment) and type of eating disor-
 197 ders [i.e. AN (subdivided by restricted [AN-R] and
 198 binge-and-purge subtype [AN-BP]), BN, EDNOS, and
 199 mixed samples].

200 The I^2 measure (Higgins & Thompson, 2002) was
 201 used to quantify the amount of between-study hetero-
 202 geneity in outcomes. Statistical significance of this mea-
 203 sure was assessed using the χ^2 statistic. Meta-regression
 204 analyses were performed in case of significant between-
 205 study heterogeneity in outcomes. Predictors in these
 206 analyses were: mean age in years and gender distribu-
 207 tion of the sample, inpatient *v.* outpatient status, coun-
 208 try in which the study was performed (coded as
 209 Western *v.* non-Western) and the methodological qual-
 210 ity score of the study.

211 Publication-bias was assessed by means of Egger's
 212 test (Egger *et al.* 1997). Where publication bias was evi-
 213 dent, trim-and-fill procedures were applied to estimate

pooled effect sizes while taking bias into account 214
 (Duval & Tweedie, 2000). 215

Results 216

The electronic searches yielded 3938 publications after 217
 duplicates were removed and 22 additional records 218
 through backward searches and reference lists. Of 219
 these, 3878 were excluded leaving a number of 82 stud- 220
 ies that reported on at least one effect size that met our 221
 inclusion criteria. Study selection, from search to inclu- 222
 sion, is presented in the flowchart in Supplementary 223
 Fig. S1. Table 1 and Supplementary Table S3 list the 224
 papers that were included for analyses and provide 225
 information on the general characteristics of them. 226
 Full references to the included papers also are pre- 227
 sented in the Supplementary reference list (pp. 14–18). 228

Prevalence rates of CM in EDs and healthy and psychiatric reference samples 229 230

Table 2 provides the prevalence rates (in percentage 231
 and 95% CIs) of self-reported CM exposure (by type) 232
 for all the EDs [total number of estimates (k)=214, 233
 N =13 059 unique individuals], for healthy and psychi- 234
 atric reference samples (k =50, N =15 092 and k =26, 235
 N =7736, respectively). In general, prevalence rates of 236
 all types of CM appeared to be two- to fourfold higher 237
 in ED samples as HC samples. However, in patients 238
 with AN of the restrictive subtype the prevalence 239
 rates of reported CM exposure were less pronounced 240
 (see Table 2) and for some ED subtypes there was over- 241
 lap in 95% CIs when compared to HCs (e.g. in the case 242
 of sexual, physical, and emotional maltreatment in 243
 samples of patients with AN of the restrictive subtype). 244
 Prevalence rates of emotional CM and exposure to >1 245
 type of CM was at least twofold higher in ED samples 246
 as compared to those reported in PC samples. ED sam- 247
 ples did not differ significantly from PC samples 248
 regarding the prevalence of physical and sexual CM. 249

CM in EDs *v.* healthy reference samples: direct comparisons 250 251

The data presented in Table 2 are derived from studies 252
 that reported CM prevalence rates in ED samples only, 253
 excluding the possibility for between-group compari- 254
 sons. In a next step, we excluded studies that lacked 255
 a reference group and ran meta-analyses on the differ- 256
 ence in prevalence rates of CM in ED samples *v.* 257
 healthy reference groups. Analyses were run stratified 258
 by type of CM and type of ED. From the data it 259
 appeared that in ED samples (any type), a history of 260
 CM (any type) is much more often reported compared 261
 to HC reference groups (ORs in most cases >2). In some 262
 specific instances there were no significant differences 263

Table 1. Basic characteristics of the included studies

Author, year	Analysis ^a	Diagnosis (n)	In/ outpatient	Maltreatment	% Female	Mean age	Country
Oppenheimer <i>et al.</i> (1985)	[1, 2, 3]	AN (36), BN (33), Mix (9)	N.K.	SA	100	24	UK
Root & Fallon (1988)	[1, 3]	BN (172)	Outpatients	PA, SA	100	25	USA
Palmer <i>et al.</i> (1990)	[1, 2, 3]	AN (80), BN (78)	Outpatients	SA	100	24	UK
Steiger & Zanko (1990)	[1, 2, 3]	AN-BP (12), AN-R (16), BN (45), HC (24), PC (21)	N.K.	SA	100	28	Canada
Stuart <i>et al.</i> (1990)	[1, 3]	BN (30), PC (15), HC (100)	N.K.	EA, SA	100	35	USA
Waller (1992)	[1, 3, 5]	BN (40)	N.K.	SA	100	25	UK
Folsom <i>et al.</i> (1993)	[1, 5]	AN (15), BN (57), Mix (21), ED-NOS (9), PC (49)	Inpatients	PA, SA	100	26	USA
Herzog <i>et al.</i> (1993)	[1, 5]	Mix (22)	Outpatients	SA	100	24	USA
Pitts & Waller (1993)	[1, 5]	Mix AN-BP/BN (41)	N.K.	SA	100	25	UK
Schmidt <i>et al.</i> (1993)	[1, 2, 3]	AN-BP (23), AN-R (63), BN (116)	Outpatients	SA	100	24	UK
Pope <i>et al.</i> (1994)	[1, 3]	BN (91)	Outpatients	SA	100	23	USA
Rorty <i>et al.</i> (1994)	[1, 5]	BN (80)	Outpatients	EA, PA, SA	100	24	USA
Waller (1994)	[1, 5]	AN (47), BN (68)	N.K.	SA	100	24	UK
Fullerton <i>et al.</i> (1995)	[1, 2, 3, 4, 5]	AN (98), BN (243), EDNOS (353), Mix AN/BN (18)	N.K.	PA, SA	100	24	USA
Garfinkel <i>et al.</i> (1995)	[1, 3]	BN (62), HC (585)	N.K.	SA	100	37	Canada
Kern & Hastings (1995)	[1, 3]	BN (30), HC (50)	Outpatients	SA	100	22	USA
Olivardia <i>et al.</i> (1995)	[1, 2, 3]	AN (6), BED (6), BN (13), HC (26)	Outpatients	PA, SA	0	18–25	USA
Schmidt <i>et al.</i> (1995)	[1]	AN-BP (23), AN-R (40), BN (95)	Outpatients	SA	100	23	UK
Sullivan <i>et al.</i> (1995)	[1, 3, 5]	BN (87)	N.K.	SA	100	27	New Zealand
Steiger <i>et al.</i> (1996)	[1, 3, 5]	BN (61)	Outpatients	PA, SA	100	26	Canada
Anderson <i>et al.</i> (1997)	[1, 2, 3]	BN (74)	Inpatients	SA	100	27	USA
Brown <i>et al.</i> (1997)	[1, 5]	AN/BN (117), HC (21)	Mix	PA, PA + SA, SA	97	28	Australia
Casper & Lyubomirsky (1997)	[1, 3]	BN (69), HC (92)	Outpatients	SA	100	N.K.	USA
Fairburn <i>et al.</i> (1997)	[1, 3]	BN (102), HC (104), PC (102)	Outpatients	PA, SA	100	24	UK
Favaro & Santonastaso (1997)	[1, 2, 3, 5]	AN (98), BN (111), ED-NOS (74)	Outpatients	SA	100	24	Italy
Friedman <i>et al.</i> (1997)	[1, 3]	BN (37), HC (48)	Inpatients	PA/SA Mix	100	29	USA
Welch <i>et al.</i> (1997)	[1, 3]	BN (102), HC (204)	Outpatients	PA, SA	100	24	UK
Fairburn <i>et al.</i> (1998)	[1, 4]	BED (52), HC (104), PC (102)	Outpatients	PA, SA	100	25	UK
Favaro <i>et al.</i> (1998)	[1, 2, 3, 5]	AN-BP (48), AN-R (38), BN (69), HC (81)	Mix	PA, SA	100	23	Italy
Waller (1998)	[1, 5]	AN (15), BN (40)	Outpatients	SA	100	23	UK
Deep <i>et al.</i> (1999)	[1, 2, 3, 5]	AN (26), BN (47), HC (44)	Outpatients	SA	100	25	USA
Matsunaga <i>et al.</i> (1999)	[1, 5]	BN (44)	Outpatients	PA, SA	100	25	Japan
Steiger <i>et al.</i> (2000)	[1, 3, 5]	BN (40), HC (25)	Outpatients	PA, SA	100	24	Canada
Webster & Palmer (2000)	[1, 2, 3]	AN (28), BN (32), HC (40), AN/BN (20), PC (40)	Mix	PA, SA	100	N.K.	UK
Grilo & Masheb (2001)	[1, 4, 5]	BED (145), HC (1125)	Outpatients	EA, PA, SA	77	38	USA

Table 1 (cont.)

Author, year	Analysis ^a	Diagnosis (n)	In/ outpatient	Maltreatment	% Female	Mean age	Country
Karwautz <i>et al.</i> (2001)	[1, 2]	AN (45), HC (45)	Outpatients	PA, SA	100	28	Austria
Mahon <i>et al.</i> (2001)	[1, 3]	BN (114)	Outpatients	PA, SA	100	27	UK
Nagata <i>et al.</i> (2001)	[1, 2, 3]	AN-BP (80), AN-R (67), BN (92), HC (99)	Mix	SA	100	23	Japan, USA
Romans <i>et al.</i> (2001)	[1]	AN (19), BN (26), AN/ BN (6), HC/PC (2522)	Outpatients	SA	100	22	New Zealand
Waller <i>et al.</i> (2001)	[1, 3, 5]	BN (61)	Outpatients	SA	100	23	UK
Grilo & Masheb (2002)	[1, 4, 5]	BED (116), HC (1125), PC (309)	Outpatients	EA, PA, SA	78	44	USA
Hartt & Waller (2002)	[1, 2, 3]	BN (15), BED (3), AN-BP (5)	N.K.	EA, PA, SA	100	29	UK
Schoemaker <i>et al.</i> (2002)	[1, 3, 5]	BN (38), HC (1350), PC (589)	Outpatients	EA, PA, PA + SA, SA	100	33	The Netherlands
Striegel-Moore <i>et al.</i> (2002)	[1, 3]	BED (102), HC (164), PC (86)	Outpatients	PA, SA	100	30	USA
Léonard <i>et al.</i> (2003)	[1, 3, 5]	BN (51), HC (25)	Outpatients	PA, SA	100	24	Canada
Basurte <i>et al.</i> (2004)	[1, 5]	Mix (25)	N.K.	PA, SA	100	N.K.	Spain
Lockwood <i>et al.</i> (2004)	[1]	AN-BP (8), AN-R (14), BN (21), EDNOS (19)	N.K.	PA, EA, SA	100	29	UK
Rayworth <i>et al.</i> (2004)	[1]	Mix (49), PC (515)	N.K.	PA, SA	100	40	USA
Steiger <i>et al.</i> (2004)	[1, 3]	BN (73), HC (50)	Outpatients	PA, SA	100	25	Canada
Van Gerko <i>et al.</i> (2005)	[1, 2, 3, 4, 5]	AN-BP (61), AN-R (62), BN (92), EDNOS (84)	N.K.	SA	100	29	UK
Vaz Leal <i>et al.</i> (2005)	[1, 3, 5]	BN (70)	N.K.	SA	100	N.K.	Spain
Wentz <i>et al.</i> (2005)	[1]	AN (37), HC (44)	N.K.	SA	95	16	Sweden
Carter <i>et al.</i> (2006)	[1, 2, 5]	AN-BP (41), AN-R (54)	Inpatients	SA	100	N.K.	Canada
Kugu <i>et al.</i> (2006)	[1]	BED/BN (21), HC (21)	Outpatients	EA, PA, SA	85	21	Turkey
Nickel <i>et al.</i> (2006)	[1, 3, 5]	BN (211)	N.K.	SA	100	18	Germany
Allison <i>et al.</i> (2007)	[1, 4]	BED (176), HC (38)	Outpatients	EA, PA, SA	79	45	USA
Claes & Vandereycken (2007)	[1, 2, 3, 5]	AN-BP (15), AN-R (22), BN (28)	Inpatients	PA, SA	100	22	Belgium
Corstorphine <i>et al.</i> (2007)	[1, 2, 3, 4, 5]	AN-BP (19), AN-R (23), BN (40), EDNOS (20)	N.K.	EA, PA, SA	99	29	UK
Feldman & Meyer (2007)	[1]	AN/BN (30), HC (163)	Outpatients	SA	0	N.K.	USA
Hepp <i>et al.</i> (2007)	[1]	AN (84), BN (152), EDNOS (41)	Mix	SA	100	28	Switzerland
Wonderlich <i>et al.</i> (2007)	[5]	BN (123)	Outpatients	EA, PA, SA	100	25	USA
Bardone-Cone <i>et al.</i> (2008)	[5]	BN (138)	N.K.	EA, PA, SA	100	26	USA
Cumella & Kally (2008, older)	[1]	Mix (302)	N.K.	SA	100	46	USA
Cumella & Kally (2008, younger)	[1]	Mix (302)	N.K.	SA	100	21	USA
Richardson <i>et al.</i> (2008)	[1, 5]	BN (89)	N.K.	PA, SA	100	17–49	Canada
Sanci <i>et al.</i> (2008)	[1, 2, 3]	AN (32), BN (35), HC (999)	Outpatients	SA	100	17	Australia
Steiger <i>et al.</i> (2008)	[1]	BN (90)	N.K.	PA, SA	100	25	Canada
Kong & Bernstein (2009)	[1, 5]	AN (29), BN (39), EDNOS (5)	N.K.	EA, PA, SA	97	24	South Korea
Favaro <i>et al.</i> (2010)	[1, 2]	AN (109), HC (554)	N.K.	PA/SA	100	~35	Italy
Mangweth-Matzek <i>et al.</i> (2010)	[1, 2, 3]	AN (9), BN (15), EDNOS (8), HC (43)	Mix	PA, SA	0	25	Austria

Table 1 (cont.)

Author, year	Analysis ^a	Diagnosis (n)	In/ outpatient	Maltreatment	% Female	Mean age	Country
Steiger et al. (2010)	[1, 2, 3]	AN-BP (8), AN-R (9), BN (108), EDNOS (13), HC (93)	Outpatients	EA, PA, SA	100	26	Canada
Becker & Grilo (2011)	[1, 4, 5]	BED (137)	Outpatients	EA, PA, SA	100	43	USA
Jaite et al. (2012)	[1, 2, 3]	AN-BP (27), AN-R (50), HC (44)	N.K.	EA, PA, SA	100	16	Germany
Sachs-Ericsson et al. (2012)	[1, 4]	BED (137), HC (2823)	Outpatients	PA, SA	58	42	USA
Backholm et al. (2013)	[1]	AN (891), BED (366), BN (1511), EDNOS (1760)	N.K.	PA, SA	97	26	Sweden
Castellini et al. (2013)	[1, 2, 3, 5]	AN (27), BN (31)	Outpatients	PA, SA	100	27	Italy
Tasca et al. (2013)	[1]	AN/BN/EDNOS (267)	N.K.	PA/SA	N.K.	27	Canada
Brewerton et al. (2014)	[1, 5]	BED/BN (139)	Outpatients	PA, SA	100	46	USA
Groleau et al. (2014)	[1]	BN (52)	Outpatients	PA, SA	100	22	Canada
Machado et al. (2014)	[1]	AN-R (58), AN-BP (28), HC (86), PC (86)	N.K.	EA, PA, SA	100	20	Portugal
Monteleone et al. (2015)	[1, 2, 3, 5]	AN (23), BN (21), HC (29)	Outpatients	EA, PA/SA	100	28	Italy
Racine & Wildes (2015)	[5]	AN (188)	Outpatients	EA, PA, SA	96	26	USA

AN, Anorexia nervosa; AN-BP, anorexia nervosa binge-purge subtype, AN-R, anorexia nervosa restrictive subtype; BED; binge-eating disorder, BN, bulimia nervosa; EA, emotional abuse (including emotional abuse and neglect and psychological abuse); HC, healthy controls; N.K. not known, PA, physical abuse; PC, psychiatric controls; SA, sexual abuse.

^a This column indicates in which meta-analysis the study in the corresponding row is included:

[1] Childhood maltreatment (CM) exposure in patients with an ED.

[2] CM exposure in patients with AN *v.* that in HCs.

[3] CM exposure in patients with BN *v.* that in HCs.

[4] CM exposure in patients with BED/EDNOS *v.* that in HCs.

[5] CM exposure and characteristics of EDs such as age at onset or psychiatric co-morbidity (for more information we refer to the Method section of the main text).

264 in CM exposure between ED and HC samples (e.g. in
265 sexual, and physical maltreatment in samples of
266 patients with AN of the restrictive subtype and
267 ED-NOS, respectively). However, in all these instances
268 that were based on a limited number of studies, the
269 effect-size pointed in the direction of higher reported
270 CM in ED samples. Table 3 provides an overview of
271 the results of these analyses together with data on
272 between-study heterogeneity and publication bias.
273 Forest plots, by type of ED, are provided in
274 Supplementary Figs S2–S5. For most pooled effect-size
275 estimates there was little evidence for between-study
276 heterogeneity or publication bias (see Table 3).

277 CM in EDs *v.* psychiatric reference samples: direct 278 comparisons

279 Direct comparisons of CM prevalence rates in ED sam-
280 ples ($N=1809$) *v.* psychiatric reference samples ($k=36$,
281 $N=13\,186$; composed mostly of depressed and
282 substance-dependent persons) showed that a history

of CM was more often reported in the EDs as relative 283
to other psychiatric illnesses (OR 1.31, 95% CI 1.08– 284
1.58, $p<0.001$). There were no marked differences for 285
type of CM or for type of ED. A forest plot is provided 286
(Supplementary Fig. S6) by type of ED. There was 287
evidence for between-study heterogeneity in outcome 288
($I^2=55.7$, $\chi^2=76.8$, $p<0.001$) and publication bias 289
(Egger's $t=3.22$, $p<0.01$). When we accounted for pub- 290
lication bias by means of trim-and-fill estimates, the 291
difference in CM exposure was no longer significant. 292

293 Association between psychiatric co-morbidity and 294 CM severity within ED patients

295 Within ED patients we assessed whether CM exposure 295
was associated with the presence of psychiatric 296
co-morbidity and suicidal/self-harm behavior. We 297
found CM to be associated with a statistically significant 298
increase in the odds on psychiatric co-morbidity and sui- 299
cidal/self-harm behavior. Results by type of co-morbidity 300
are provided in Table 4a and in the Supplementary 301

Table 2. Prevalence (in bold) of Child maltreatment (CM) exposure (by type) and estimated 95% CI and diagnostic status (by type)

	Sexual CM	Physical CM	Emotional CM	Double CM counts ^a
All EDs	31% (27–35%) (k = 121, N = 12 294)	26% (21–32%) (k = 63, N = 8620)	45% (38 — 54%) (k = 17, N = 1170)	46% (31 — 59%) (k = 13, N = 1173)
AN	26% (21–33%) (k = 42, N = 2689)	17% (11–25%) (k = 19, N = 1666)	34% (23–46%) (k = 8, N = 437)	No data available
AN binge-purge subtype	37% (30–45%) (k = 13, N = 424)	25% (16–35%) (k = 5, N = 117)	48% (34 — 62%) (k = 2, N = 46)	No data available
AN restrictive subtype	19% (12–27%) (k = 14, N = 553)	17% (9–30%) (k = 7, N = 251)	24% (18 — 32%) (k = 4, N = 182)	No data available
BN	35% (29–41%) (k = 45, N = 4395)	33% (23–45%) (k = 20, N = 2998)	81% (18–98%) (k = 2, N = 63)	49% (38–60%) (k = 3, N = 326)
BED	24% (16–33%) (k = 8, N = 1233)	23% (14–35%) (k = 8, N = 1233)	59% (48–70%) (k = 4, N = 574)	No data available
EDNOS	21% (9–45%) (k = 6, N = 2351)	19% (5–54%) (k = 4, N = 2193)	45% (–) ^b (k = 1, N = 20)	No data available
Mix EDs	42% (32–52%) (k = 20, N = 1626)	41% (33–50%) (k = 12, N = 530)	46% (35–57%) (k = 2, N = 83)	40% (16–56%) (k = 2, N = 122)
Healthy controls	13% (9–17%) (k = 32, N = 9245)	10% (7–15%) (k = 18, N = 6515)	13% (8–20%) (k = 9, N = 2737)	7% (1–35%) (k = 4, N = 1375)
Psychiatric controls	27% (19–38%) (k = 11, N = 4197)	30% (18–44%) (k = 8, N = 1639)	21% (8–46%) (k = 3, N = 847)	12% (5–30%) (k = 3, N = 1053)

AN, Anorexia nervosa; BN, bulimia nervosa; BED, binge-eating disorder; ED, Eating disorder; EDNOS, eating disorder not otherwise specified.

^a Double maltreatment counts entails exposure to >1 type of CM (regardless of the exact CM type).

^b No confidence interval could be calculated because only one estimate was available.

The number of studies and individuals given here in this table are not ‘unique’. Individuals may appear in various cells (e.g. subjects on which data is presented in an ‘emotional CM cell’ may also appear in a ‘sexual CM cell’. N’s thus do not add up to the unique N, just as the number of effect sizes does not add up to the number of studies.

material as a forest plot by type of ED (Supplementary Fig. S7). These associations were not due to a particular CM or ED type although statistical power to show differences may have been too low due to a relatively limited number of studies (range k = 7–16, range N = 497–2109). We did not detect evidence for between-study heterogeneity (p = 0.38), but publication bias was observed (Egger’s t = 2.4, p = 0.02). Effect-size estimates were not markedly different after we applied trim-and-fill estimates (data not shown).

Association between ED psychopathology and CM exposure within ED patients

We aggregated the reported associations between CM exposure and (i) severity of ED pathology measured on continuous scales such as the EAT, (ii) frequency of binge-purge episodes per week/month, (iii) age at ED onset, (iv) continuous scores on depression/anxiety severity within patients with an ED, and (v) the use of diuretics and laxatives. In general CM exposure was associated with higher severity of ED pathology, a larger number of binge/purge episodes, an earlier age at ED onset, higher symptom level of depression/anxiety, and a higher frequency of use of diuretics and laxatives (see Table 4b and forest plots; Supplementary Figs S8 and S9). In line with the idea that a history of CM is related with a more severe ED pathology was that prevalence rates of CM were in general higher in inpatients as compared to outpatients (prevalence rates of 0.45 v. 0.29 respectively, p for the difference <0.01).

Association between psychiatric co-morbidity and severity measures of CM exposure within ED patients

Forty-one effect-size estimates (N = 4683) used a continuous CM exposure measure (e.g. the sum score on the childhood trauma questionnaire) to predict general psychopathology (e.g. depression severity) and ED severity features (e.g. EAT score) within ED patients. Pooling these estimates showed that the severity of CM was positively correlated with continuous indices for depression and anxiety (weighted Pearson’s r = 0.27, 95% CI 0.10–0.53, k = 19, N = 2197) but not with ED severity features (r = 0.26, 95% CI –0.11 to 0.67, k = 13, N = 1067).

Given the limited amount of studies, we were not able to formally test whether the above-reported dose-response associations were particularly due to one type of CM or ED.

Predictors of CM prevalence rates and between-study heterogeneity in outcomes

By means of meta-regression analyses we identified sources of between-study heterogeneity in prevalence

Table 3. Child maltreatment (CM) exposure (overall and by type) and the odds (in bold) on ED (by type) v. healthy controls by type

	k	N	OR (95% CI)	Heterogeneity		Publication bias Egger's <i>t</i>
				<i>I</i> ²	χ^2	
Overall CM						
Any ED	96	34 521	2.47 (2.15–2.84)***	57.4	222.6***	1.84
AN	33	5665	2.25 (1.66–3.06)***	51.0	65.3**	0.34
Binge-purge subtype	11	852	3.49 (1.66–7.33)***	63.7	22.0*	2.05*
Restrictive subtype	17	1571	2.38 (1.42–3.99)*	45.1	25.5	1.96*
BED ^a	14	11 979	2.23 (1.88–2.64)***	23.2	23.2*	0.9
BN	30	10 499	3.05 (2.20–4.23)***	73.1	107.8***	2.08***
EDNOS ^a	2	304	2.51 (0.70–9.05)	0.81	5.0*	n.a. ^b
Mix EDs	16	7704	2.61 (2.01–3.38)***	6.3	17.1	0.62
Sexual CM						
Any ED	49	15 006	2.23 (1.79–2.78)***	63.8	132.7***	1.92
AN	19	3251	1.98 (1.31–2.98)***	61.0	46.1*	0.49
Binge-purge subtype	5	481	3.09 (1.57–6.08)*	77.5	17.8**	2.1*
Restrictive subtype	7	715	1.50 (0.89–2.54)	61	15.1	1.8
BED	5	3147	1.88 (1.38–2.55)***	34.2	6.1	0.98
BN	16	4911	2.57 (1.62–4.01)***	76.2	63.0**	1.13
EDNOS	1	152	5.11 (1.94–13.64)**	n.a. ^b	n.a. ^b	n.a. ^b
Mix EDs	8	3516	2.37 (1.53–2.68)***	29.7	9.9	0.55
Physical CM						
Any ED	32	10 347	2.66 (2.18–3.25)***	29.3	43.8	1.40
AN	10	996	2.42 (1.34–4.35)***	32.4	13.3	0.64
Binge-purge subtype	3	300	3.06 (1.22–7.67)**	0.12	2.2	1.43
Restrictive subtype	5	538	2.78 (1.13–6.78)*	33.5	6.0	0.98
BED	5	3147	2.57 (1.99–3.31)***	16.0	4.8	0.40
BN	8	2771	3.78 (2.25–6.32)***	52.6	14.8*	1.12
EDNOS	1	152	1.38 (0.74–2.54)	n.a. ^b	n.a. ^b	n.a. ^b
Mix EDs	8	3516	2.64 (1.82–3.82)**	1.0	6.2	0.17
Emotional CM						
Any ED	10	4594	2.98 (2.30–3.87)***	25.0	11.9	0.81
AN	4	254	3.81 (2.05–7.08)***	0.1	1.1	0.56
BED	3	2725	2.44 (1.73–3.43)***	55.7	4.5	1.47
BN	2	1438	5.13 (2.80–9.40)***	n.a. ^b	n.a. ^b	n.a. ^b
Mix EDs	1	42	8.00 (0.92–69.72)	n.a. ^b	n.a. ^b	n.a. ^b
<i>No analyses were run for the AN subtypes, and ED-NOS given a lack of data</i>						
Double CM counts ^c						
Any ED	5	4574	2.96 (1.512.55)*	0.2	0.9	0.22
<i>No analyses were run for Any ED, AN, the AN subtypes, ED-NOS, and Mix ED given a lack of data</i>						

AN, Anorexia nervosa; BN, bulimia nervosa; BED, binge-eating disorder; ED, eating disorder; EDNOS, eating disorder not otherwise specified; n.a., not available; OR, odds ratio; CI, confidence interval.

^a Although BED was subsumed in the EDNOS category in DSM-IV we decided to report data separately for these illnesses where possible because EDNOS is not necessarily BED.

^b This could not be calculated because only 1 or 2 estimates were available.

^c Double maltreatment counts entails exposure to >1 type of CM (regardless of the exact CM type).

*Statistically significant at $p < 0.05$; **statistically significant at $p < 0.01$; ***statistically significant at $p < 0.001$.

352 rates of CM. We found that a small part of the hetero-
353 geneity could be explained by (1) the methodological
354 quality of a study; studies of higher quality reported
355 somewhat lower prevalence rates ($R^2 = 0.05$, $p < 0.01$);

(2) sample size; studies that used a larger sample size
reported somewhat lower prevalence rates ($R^2 = 0.07$,
 $p < 0.01$), (3) a lower age threshold for maltreatment
to count as CM; studies that applied a lower age

356
357
358
359

Table 4. Child maltreatment exposure, co-morbidity (part a), and eating disorder (ED) characteristics (part b) in ED patients

	k	N	OR (95% CI)	Heterogeneity		
				I ²	χ ²	Publication bias Egger's t
<i>(a) Psychiatric co-morbidity</i>						
Alcohol/substance dependence	8	990	1.81 (1.39 to 2.36)***	1.0	2.5	2.1
Axis I disorder	8	497	1.54 (1.08 to 2.20)*	2	6.0	1.3
Axis II disorder	7	550	2.33 (1.32 to 4.13)***	61.8	15.7* ^b	1.9
Suicidality/self-harm	16	2109	2.59 (2.02 to 3.31)***	32	21.5	0.8
Hedges' g (95% CI)						
<i>(b) ED features</i>						
ED severity ^a	9	887	0.27 (0.14 to 0.41)***	85.8	63.3* ^b	1.3
No. of binge/purge episodes	22	1580	0.32 (0.22 to 0.42)***	40.8	35.5* ^b	1.5
Age at onset	4	175	-0.32 (-0.62 to 0.02)*	1.0	1.2	0.3
Depression/anxiety severity	7	546	0.67 (0.48 to 0.86)***	72.5	21.8* ^b	1.4
Diuretic/laxative use	4	583	0.35 (0.20 to 0.51)**	0.7	4.1	0.9

OR, Odds ratio; CI, confidence interval.

^a ED severity was measured eight times with the EAT (Mintz & O'Halloran, 2000), four times with the EDE (Luce & Crowther, 1999); three times with the EDI (Wear & Pratz, 1987) and one time with the BITE (Henderson & Freeman, 1987).

^b In case there was between-study heterogeneity in outcomes, we tested whether this could be explained by the pooling together of several related but distinct constructs (e.g. different Axis II disorders or different questionnaires to measure ED severity with). Heterogeneity in outcomes, however, were in all cases independent of pooling constructs.

360 threshold (e.g. 16 *v.* 18 years age) reported somewhat
 361 lower prevalence rates ($R^2=0.03$, $p<0.01$), and (4)
 362 year of publication; studies that were published more
 363 recently also yielded somewhat lower prevalence
 364 rates ($R^2=0.04$, $p<0.01$). Besides, prevalence rates of
 365 CM were in general somewhat lower when they were
 366 acquired by means of an interview *v.* self-report ques-
 367 tionnaire (prevalence rates of CM of 0.28 *v.* 0.34
 368 respectively, $p<0.05$). Mean age of the sample at
 369 assessment, gender distribution of a particular study,
 370 and whether the data was gathered in a Western *v.* a
 371 non-Western culture were not related to between-study
 372 heterogeneity in prevalence rates.

373 There were no statistically significant associations
 374 between the pre-specified moderators and between-
 375 study variation in OR estimates on the association
 376 between CM in ED samples *v.* HC and PC reference
 377 samples.

378 Discussion

379 The purpose of this study was to provide a quantita-
 380 tive overview of studies that report on the association
 381 between CM exposure (including type and severity)
 382 and a lifetime diagnosis of an ED (including type
 383 and severity parameters). Eighty-two studies, report-
 384 ing on 13 059 individuals with an ED, 15 092 HC and
 385 7736 PC subjects were pooled. This pool of data, the
 386 largest of its kind to date, indicated that reported
 387 CM, regardless of type (emotional, physical, or sexual),

was strongly associated with the presence of all types 388
 of EDs. Overall, the lifetime prevalence rates of CM 389
 ranged from 1–35% in HCs, 5–46% in PCs, and 21– 390
 59% in individuals with any ED. The difference in 391
 prevalence rates of CM between patients with an ED 392
 and HCs is largely in line with the findings of previous 393
 meta-analyses (Rind *et al.* 1998; Smolak & Murnen, 394
 2002; Chen *et al.* 2010; Caslini *et al.* 2016). Some discrep- 395
 ancies were found when considering the specific types 396
 of CM. Caslini *et al.* (2016) for instance did not find a 397
 significant difference with respect to the prevalence 398
 of emotional CM among ED patients *v.* HCs whereas 399
 we do. The reason for this discrepancy probably is statist- 400
 ical power; we report on 10 studies that assessed this 401
 effect and Caslini *et al.* (2016) reported on six. Besides, 402
 their estimated point estimate on the association 403
 between emotional CM and EDs (OR range 2.13– 404
 4.15) lies within the CI that we estimate on this associ- 405
 ation (i.e. 2.05–7.08). In addition, we found that the 406
 associations between CM and ED were particularly 407
 strong for BN, BED and for AN of the *binge-purge* sub- 408
 type, whereas this seemed to be less the case for AN of 409
 the restrictive subtype. This is in line with the finding 410
 that ED patients with CM reported more bingeing and 411
 purging. Our findings that ED patients reporting CM 412
 had an earlier age at ED onset, suffered a more severe 413
 form of the illness, and binge-purged more often as 414
 compared to their non-maltreated counterparts is 415
 new. Also new is the finding that individuals with an 416
 ED reporting CM were more likely to be diagnosed 417

with a co-morbid psychiatric disorder and to be suicidal relative to ED patients who did not report exposure to CM is new. The finding of higher psychiatric co-morbidity in CM exposed patients with EDs is remarkable because this kind of co-morbidity tends to be the rule rather than the exception in individuals with EDs (Ulfvebrand *et al.* 2015; Keski-Rahkonen & Mustelin, 2016). The fact that both CM exposed and non-exposed groups allegedly have similar co-morbidities makes a potential difference between the two less likely to be detected.

We found some evidence suggesting higher prevalence rates of CM in patients with an ED compared to PCs. This finding, however, did not remain statistically significant when we corrected for the likely presence of publication bias. So, the difference in CM exposure between patients with an ED as compared to PCs may reflect the effect of some small-scale studies reporting overly large effect-size estimates (and hence are more likely to get published). In general, these *new* findings should be considered with some caution because with each further breakdown of variables (i.e. subtypes of eating disorder, trauma type, or severity measure) analyses were run on smaller number of studies, and results become less reliable (Cochrane Community Handbook, 2016).

The consistency of our findings across most types of EDs and forms of maltreatment (see the forest plots), the strength and dose-response nature of the association, and the likely temporal precedence suggests a causal link of CM exposure on ED pathology. Nevertheless, in the absence of longitudinal and experimental data we cannot prove this here (Hill, 1965). Furthermore, a large part of between-study heterogeneity in outcome could not be explained.

The mechanisms that may underlie the observed associations remain elusive. Moreover, the current meta-analytical approach cannot shed much light on the underlying processes. Yet, the outcomes do underline the importance of further elucidating the processes linking CM and EDs, with a particular focus on the associations with bingeing and purging behavior in the context of the enhanced negative affect (i.e. depression, suicidality and anxiety) and co-morbidity in EDs patients with reported CM. Clinically, it is important to elucidate whether similar or specific processes are at stake in patients with a history of CM compared to patients who do not report this (i.e. maladaptive emotion regulation styles (i.e. distorted cognitive schemas of self and body dissatisfaction, and dissociation), and whether this is further dependent on the type of CM a patient has been exposed to. Body dissatisfaction and dissociation may, for example, be specifically at stake in patients with histories of sexual CM (Dunkley *et al.* 2010; Muehlenkamp *et al.* 2011;

Duarte *et al.* 2016; Preti *et al.* 2016), while emotion dysregulation in general may be a process that is particularly prominent in individuals with a history of emotional CM (Michopoulos *et al.* 2015; Moulton *et al.* 2015; Racine & Wildes, 2015). In addition, research on dysregulations in stress-sensitive neurobiological systems, including the HPA-axis and serotonin and dopamine systems is also needed to further elucidate the key neurobiological processes underlying the link between CM and EDs (Kaye *et al.* 2013a, b; Nemeroff, 2016).

Strengths and limitations

In addition to the use of meta-analytical techniques, this study has as strength that it is based on large sample sizes of diagnosed ED subjects and controls and the identification of the type and severity of CM. In addition, unlike previous meta-analyses, rates of CM were also examined in a non-ED psychiatric comparison groups and dose-response associations were assessed at the predictor and outcome level.

A limitation of our work is that prospective studies were not available and mediators of the associations of interest could not be assessed by means of meta-analysis. Moreover, counting as limitation is that we were not able to model the effect of CM exposure on ED course (e.g. duration of illness). A factor that might limit the generalizability of our findings is that most studies solely included women. Hence our results may not generalize to men with EDs. Furthermore, our findings may not generalize to the child and adolescent ED population.

Clinical implications

Our results may have clinical implications. CM exposure obviously conveys important information on who is at risk for developing ED although between-study and between-subject variability in risk-estimates exist. Public health efforts will do well by prioritizing a reduction of CM prevalence and as such decrease the burden caused by the EDs (Fairburn & Harrison, 2003; Arcelus *et al.* 2011; Mischoulon *et al.* 2011; Mitchell *et al.* 2012; Smink *et al.* 2013; Yao *et al.* 2016) by depression (Nanni *et al.* 2012), substance abuse, suicide, and anxiety (Norman *et al.* 2012; Mason *et al.* 2014; Yao *et al.* 2016). Moreover, the importance of CM exposure should also be addressed in the clinical setting. As shown above, reported CM is indicative of a more severe clinical profile. Moreover, one study that we know of accounted for the impact of CM on ED treatment outcome and found that ED patients exposed to CM improve less as compared to those who were not exposed (Vrabel *et al.* 2010; Norman *et al.* 2012), a phenomenon that is well-documented

525 in the treatment of depression (Nanni *et al.* 2012).
 526 Standard assessment of CM exposure and additional
 527 treatment interventions for those reporting CM, such
 528 as enhancing emotion regulation strategies and/or
 529 focused trauma treatment may increase the rather
 530 low treatment responses in ED patients.

531 Conclusions

532 We performed the most comprehensive meta-analysis
 533 of the association between CM exposure and EDs to
 534 date. We found that CM, regardless of whether it is
 535 emotional, sexual and/or physical in nature, was
 536 strongly associated with the presence of all types EDs
 537 and severity parameters that characterize these illnesses,
 538 often in a dose dependent manner. Notwithstanding
 539 strong and consistent associations, unexplained
 540 between-study heterogeneity in outcomes remained
 541 for most of the associations that we observed. This
 542 implies between-study and probably between-subject
 543 variability in risk estimates that need to be explained
 544 in future work on this important topic.

545 Supplementary material

546 The supplementary material for this article can be
 547 found at <https://doi.org/10.1017/S0033291716003561>.

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556 Declaration of Interest

557 None.

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