ORIGINAL ARTICLE

An investigation of temperament endophenotype candidates for early emergence of the core cognitive component of eating disorders

S. M. Wilksch* and T. D. Wade

School of Psychology, Flinders University, Adelaide, Australia

Background. This study was designed to investigate potential temperament endophenotypes for clinically significant importance of shape and weight.

Method. Seven temperament risk factors for eating disorders and the Eating Disorder Examination were assessed in 699 female twins aged 12–15 years. Each variable was evaluated against the following endophenotype criteria: associated with illness in the general population; found in non-affected family members at a higher rate than in the general population; and, heritable.

Results. All seven variables were significantly associated with clinically significant importance of shape and weight, while thin-ideal internalization, ineffectiveness, body dissatisfaction and sensitivity to punishment were found at significantly elevated levels in non-affected twins, when controlling for sister's temperament score. These four variables had genetic correlations with importance of shape and weight, ranging from 0.48 to 0.95.

Conclusions. Future research should evaluate the stability of the identified endophenotypes and their utility for predicting significant growth in importance of shape and weight, and also whether different endophenotypes emerge when the importance of weight and shape reaches its peak in adolescents, around 15 to 16 years of age.

Introduction

The importance of weight and shape has been described as the 'core psychopathology' of eating disorders (Cooper & Fairburn, 1993) and is included in the current diagnostic definitions for eating disorders as 'undue influence of body shape or weight on self-evaluation' (APA, 1994). Clinically significant (or threshold) importance of shape and weight has been operationalized as a mean item score of $\geqslant 4$ on the Shape Concern and Weight Concern scales of the Eating Disorder Examination 12th edition (EDE; Fairburn & Cooper, 1993). Robust prospective evidence now exists for the importance of shape and weight in the development and maintenance of eating disorders (Goldfein *et al.* 2000; Gowers & Shore, 2001; Jacobi *et al.* 2004) and in the cognitive model of eating disorders this construct is postulated to lead to the

development of extreme and rigid dietary behaviour that can lead to the development of disordered eating (Fairburn *et al.* 2003).

Whilst this variable represents an important target of both eating disorder prevention and treatment, predictors of this phenotype have received surprisingly little empirical attention. A useful framework for investigating such predictors is a candidate endophenotype approach, where endophenotypes are seen to be expressions of biological markers for a phenotype that meet the following criteria (Gershon & Goldin, 1986): (i) associated with illness in the general population; (ii) a stable, state-independent characteristic (i.e. observable regardless of whether illness is active); (iii) found in non-affected family members at a higher rate than in the general population, and; (iv) heritable. The importance of weight and shape in adults has been previously found to be mainly affected by the environment (Wade & Bulik, 2007), with only 25% of the variance accounted for by additive genetic action, with 95% confidence intervals (CI) of 14–36%. Identifying endophenotypes that are more influenced by genetic action and that share genetic

* Address for correspondence: S. M. Wilksch, Ph.D., School of Psychology, Flinders University, PO Box 2100, Adelaide, SA, 5001, Australia.

(Email: simon.wilksch@flinders.edu.au)

risk factors with the importance of weight and shape may allow for identification of plausible biological mechanisms along the pathway to development of this phenotype.

Another important potential of a candidate endophenotype approach in psychiatry research is a shift from unitary conceptualizations of psychopathology to a more homogeneous view of common features across diagnostic categories (Berrettini, 2005) that will permit refinement of diagnostic criteria (Bulik *et al.* 2007). For example, Hollander *et al.* (2007) have suggested that eating disorders, obsessive–compulsive disorder and autistic spectrum disorders be a joint diagnostic category due to shared personality traits caused by underlying biological and/or environmental triggers, while the Maudsley group is investigating similar representations of cognitive inflexibility in the development of eating disorders (e.g. Treasure *et al.* 2007). It is important that investigations of potential endophenotypes include temperament as this is likely to provide markers more genetically influenced than importance of shape and weight and may provide the greatest value in informing aetiological processes (Lilenfeld *et al.* 1997).

Thus the current study was designed to investigate potential temperament endophenotypes for clinically significant importance of shape and weight. Temperament variables selected to be investigated were those that have received previous empirical support for playing a causal role in the development of disordered eating and are theorized to be premorbid to the development of importance of weight and shape, are likely to be more genetically influenced than importance of shape and weight, and are likely to be relatively stable in the absence of a targeted intervention. Variables meeting these criteria included thin-ideal internalization, ineffectiveness, body dissatisfaction, concern over mistakes, sensitivity to punishment/reward and obsessionality.

Thin-ideal internalization and body dissatisfaction both feature in Stice's dual pathway model of bulimic pathology (Stice, 2001) and prospectively predict growth of bulimic pathology through both negative affect and dieting. Both body dissatisfaction and drive for thinness (seen to be allied to thin-ideal internalization) have been found to be affected significantly by heritability of 59% (95% CI 53–65%) and 51% (95% CI 44–58%) respectively (Keski-Rahkonnen *et al.* 2005). Ineffectiveness has been identified as a risk factor for disordered eating (Jacobi *et al.* 2004), and it could be argued that ineffectiveness is allied with 'core low self-esteem' in the transdiagnostic model of eating disorders (Fairburn *et al.* 2003), hypothesized to contribute to the onset of clinically significant importance of shape and weight. The heritability of self-esteem has been found to be 0.40 (95% CI 0.26–0.54) in 14-year-old girls (Raevuori *et al.* 2007). Perfectionism related to concern over mistakes has been uniquely associated with eating disorders compared with other psychiatric disorders (Bulik *et al.* 2003), and linked prospectively (Tyrka *et al.* 2002) and experimentally (Shafran *et al.* 2004) with the development of eating disorder symptoms. Previous examination of this construct in adults has estimated the heritability of this perfectionism construct to be 0.39 (95% CI 0.27–0.49; Wade & Bulik, 2007). Based on Gray's model of personality (Gray, 1981), sensitivity to punishment has been proposed as an indicator of anxiety, while sensitivity to reward has been suggested as an indicator of impulsivity. Jacobi *et al.* (2004) described anxiety as a non-specific risk factor for eating disorders and patients with eating disorders characterized by bingeing and purging have been consistently found to score higher on novelty-seeking measures, indicating higher levels of impulsivity and excitability (Cassin & von Ranson, 2005). Impulsivity has also been linked to impaired serotonergic responsiveness, which is associated with greater bulimic symptom severity (Jimerson *et al.* 1992; Steiger *et al.* 2001). Both temperaments have been found to be influenced significantly by heritability in young adults (Larsson *et al.* 2006; Nes *et al.* 2007). Finally, premorbid obsessive–compulsive personality disorder has been found to be significantly higher among non-eating-disordered relatives of eating-disorder probands than among relatives of comparison subjects (Lilenfeld *et al.* 1998; Bellodi *et al.* 2001), with the suggestion that eating disorders should be considered as part of the obsessive–compulsive spectrum of disorders. Previous research has indicated that just under half the variation in obessionality is due to genetic influences (Clifford *et al.* 1984).

Thus the current study sought to examine if any of the risk factors described above met criteria for endophenotypes for one of the diagnostic criteria for eating disorders, namely clinically significant importance of shape and weight. We examined adolescents who were, on the whole, asymptomatic for disordered eating as they were too young for risk to be expressed (Wade *et al.* 2008), but of whom 7.9% reported clinical levels of importance of shape and weight. In particular, we examined three of the criteria for endophenotypes, namely association with illness in the general population, levels in non-affected family members, and heritability along with shared genetic risk factors with importance of weight and shape. It was not possible to investigate the second criterion (i.e. stable, state-independent characteristic), given that the current paper describes only the first wave of data of a prospective research project.

Method

Participants

Female–female twins who were registered with the Australian Twin Registry (ATR) and were aged between 12 and 15 years of age and their parents were approached to participate in the present study by the ATR. Of the 719 families approached by the ATR, 411 (57.2%) agreed to participate, 237 (32.9%) said no, and 71 (9.9%) did not reply. Families were then approached by the researchers with self-report questionnaires sent to both parents, including those families where the parents did not live together. At this stage 595 parents returned questionnaires, representing 351 families. When questionnaires were returned from the parents, an adapted version of the EDE was conducted over the telephone with the twins, at separate times and with a different interviewer for each child in the family. Interviews were completed with 699 children, where 349 Twin 1 children participated, 350 Twin 2 children participated, and two Twin 1 children and one Twin 2 child did not participate (two children had cerebral palsy and had difficulties talking and one withdrew), thus representing 348 complete pairs and three incomplete pairs. Parents reported their child's zygosity as follows: 184 monozygotic (MZ) pairs, 158 dizygotic (DZ) pairs, and eight where zygosity was not reported. This sample represents 48.8% of those families who were approached by the ATR. The mean age of the girls at the time of the data collection was 13.96 years (S.D. = 0.80 years), ranging from 12.70 to 16.28 years (two pairs of twins had turned 16 years between the approach and data collection). The total protocol consisted of a self-report questionnaire for the parents and a telephone interview for the twins. Zygosity assignment was based on parental responses to standard questions about physical similarity and confusion of twins by parents, teachers and strangers, methods that give better than 95% agreement with genotyping (Eaves *et al.* 1989). The Flinders University Clinical Research Ethics Committee approved the data collection process and written informed consent from parents and written assent from the twins was obtained after the procedures had been fully explained.

Measures

Eating interview

This protocol has been previously described (Wade *et al.* 2008). The telephone interview consisted of two parts. The first part utilized the EDE (Fairburn & Cooper, 1993) and the second part consisted of questions from various self-report questionnaires. The EDE included questions relating to behavioural features of eating disorders as well as dietary restraint, and eating, shape and weight concern over the last 28 days. In addition, all diagnostic questions addressed a 3-month time-frame. The EDE was slightly modified in line with previous recommendations for use with children (Bryant-Waugh *et al.* 1996) and according to current training protocols for use with children. In order to allow for the assessment of lifetime disorders using DSM-IV diagnostic criteria, the EDE was also revised with insertion of lifetime questions, including the age range during which the behaviour occurred in order to assess the co-occurrence of features, as used previously with an adult sample of twins (Wade *et al.* 2006). Three of these revisions are of relevance to the current study. First, as the EDE does not currently assess fasting (apart from those who have experienced concurrent binge episodes), a 3-month and lifetime version of the 'avoidance of eating' question was included. The threshold criterion was met if the person had gone for ≥8 or waking hours without eating anything in order to influence shape or weight on more than half the days each week for a 3-month period. Second, lifetime occurrence of objective binge eating was assessed with an additional question about whether the behaviour met threshold criteria (at least twice per week for a 3-month period with breaks of no more than 2 weeks). Third, lifetime questions assessing use of self-induced vomiting, laxatives, diuretics and excessive exercise were included, along with questions about threshold frequency. The threshold was defined the same as for objective binge eating with the exception of excessive exercise, which was defined as driven or compulsive exercise that had occurred for at least 1 h 5 days per week for a 3-month period with no breaks of more than 2 weeks. Questions were asked separately for competitive sport and other forms of exercise, where only exercise for weight or shape reasons was included and exercise for fitness or recreation was excluded. Abstinence from these weight-control behaviours for ≥2 weeks was assessed by a follow-up question, as was co-occurrence with any binge eating behaviour.

Given the difficulty of assessing lifetime importance of weight and shape, these diagnostic items were assessed for the previous 3 months and then during any lifetime objective binge eating behaviour or anorexia nervosa symptoms.

Weight and height of the twins were reported by both the mother and father separately. These reports were highly correlated and so the mother's report was used in the current report or the father's report if the mother's was missing. The percentage weight for height ratio was used in the current study as it is considered to be more accurate for children than body

Table 1. *Summary and description of the self-report measures used in the analyses*

Variable	Description	Cronbach's α	Reference
Thin-ideal internalization	Multi-dimensional Media Influence Scale, six items	0.89	Cusumano & Thompson (2001)
Ineffectiveness	Eating Disorder Inventory, 10 items	0.088	Garner *et al.* (1983)
Body dissatisfaction	Eating Disorder Inventory, eight items	0.91	Garner *et al.* (1983)
Concern over mistakes	Multi-dimensional Perfectionism Scale, nine items	0.87	Frost *et al.* (1990)
Sensitivity to punishment	Sensitivity to Punishment and Sensitivity to Reward Questionnaire, five items	0.74	Torrubia *et al.* (2001)
Sensitivity to reward	Sensitivity to Punishment and Sensitivity to Reward Questionnaire, five items	0.64	Torrubia *et al.* (2001)
Obsessionality	Vancouver Obsessional Compulsive Inventory – Just Right scale, 12 items	0.85	Thordarson *et al.* (2004)

mass index. As suggested by Cole *et al.* (2007), underweight was considered to be $\leqslant 80\%$ weight for height, while overweight was considered to be $\geqslant 120\%$ weight for height.

Temperament

The self-report questionnaires were also asked in the interview and are described in Table 1. The internal consistency for each scale was generally adequate; however, the five-item Sensitivity to Reward scale (Torrubia *et al.* 2001) was not highly reliable ($\alpha = 0.64$). It should be noted that the Sensitivity to Punishment and Sensitivity to Reward Questionnaire was abbreviated from its original 48-item version to a 10-item version in order to reduce the burden of measure completion. The selected items were those that have been previously found to load most strongly on the sensitivity to punishment (five items) and sensitivity to reward (five items) factors (Torrubia *et al.* 2001).

The only deviation from prescribed scoring procedures of any measure was for the Eating Disorder Inventory scales (Ineffectiveness and Body Dissatisfaction) where a system of 1 (always) to 6 (never) was used. The original scoring for eating disorder populations was not used (0, 0, 0, 1, 2, 3) as this was neither expected to be a sensitive enough measure of change in a population expected to be less extreme with respect to eating disorder behaviour and psychopathology nor to have acceptable internal consistency (Schoemaker *et al.* 1994; van Strien & Ouwens, 2003).

Interviewers

All interviewers were postgraduate clinical psychology trainees ($n = 16$) who had been trained in use of the EDE. Each of the interviews was taped and

corrective feedback was provided until the interviewer had acquired the skills required to complete the interview independently. Throughout the interviewing process monthly group meetings were held to discuss the interview process in order to ensure interview fidelity. Independent ratings by the second author (T.D.W.) of a subsample of 20 randomly chosen interviews (five from each of the four age groups, i.e. 12, 13, 14 and 15–16 years) yielded inter-rater reliability statistics (intra-class correlation coefficients) of the four subscales (last 4 weeks) as follows: 1.00 (dietary restraint); 0.96 (eating concern); 0.98 (weight concern); 0.99 (shape concern).

Statistical analyses

Descriptive data

All analyses were conducted using SPSS (version 15; SPSS Inc., Chicago, IL, USA). Associations between threshold shape and weight importance, eating disorder behaviours and weight status were investigated using generalized estimating equations (GEE), where results are presented as odds ratios (OR) with 95% CIs. Given that Twin 1 and Twin 2 are correlated observations, GEE corrects for within-family shared variance that is inherent in family research.

Endophenotype criteria

To investigate whether the measured temperament variables were candidate endophenotypes for threshold importance of shape and weight, three different analyses were conducted in order to assess the three endophenotype criteria. First, GEE were used to investigate if the within-twin temperament variables were associated with threshold importance of shape and weight, where results are presented as OR with 95% CIs. Second, logistic regression was used to

Table 2. *Frequency of lifetime disordered eating behaviours, weight status and their association with clinically significant importance of shape and weight in twins*

Is this threshold behaviour present?	Importance of shape and weight		OR (95% CI)	p
	$\geqslant 4$ (n=56) Yes (%)	<4 (n=643) Yes (%)		
Any threshold eating disorder behaviour	15 (26.8)	16 (2.5)	14.21 (6.41–31.46)[a]	<0.01
Objective bulimic episodes	1 (1.7)	2 (0.3)	6.40 (0.61–66.83)	0.12
Fasting (avoidance of eating)	8 (14.2)	6 (0.9)	17.62 (6.06–51.19)[a]	<0.01
Excessive exercise	12 (21.4)	9 (1.3)	18.30 (7.07–47.41)[a]	<0.01
Weight status				
Weight for height $\leqslant 80\%$	1 (1.7)	27 (4.2)	0.51 (0.09–3.02)	0.46
Weight for height $\geqslant 120\%$	25 (44.6)	75 (11.7)	6.11 (3.32–11.24)[a]	<0.01

OR, Odds ratio; CI, confidence interval.

[a] Significant predictors of threshold importance of shape and weight.

investigate whether Twin 2 scores for temperament variables were associated with threshold importance of shape and weight in Twin 1, where results are presented as OR with 95% CIs. For this and subsequent investigations, where discordant threshold importance of shape and weight was present across Twin 1 and Twin 2, the data were reorganized such that all twins who had high importance of shape or weight were assigned as Twin 1. The second investigation was repeated but this time with the addition of controlling for Twin 1 scores on the respective temperament variables, in order to identify the most robust predictors.

Heritability was investigated by a comparison of correlations between Twin 1 and Twin 2 of each MZ and each DZ pair, derived using Maximum Likelihood Estimations in Mx (Neale, 1997). In addition, univariate twin models were tested for each construct, reporting only the full model that included additive genetic influences (A), shared environment (C) and non-shared environment (E), as we did not have the power to distinguish between the full and sub-models. Additionally, a multivariate Cholesky decomposition analysis was used as a means of calculating the genetic correlations between the five variables. These genetic correlations represent the extent to which the same genes contribute to the observed phenotypic correlation between the variables.

Results

Descriptives

Of the 696 twins for which importance of shape and weight data were available, a total of 55 twins (7.9%) recorded threshold importance of shape and weight. Of these, 43 were discordant for threshold importance

of shape and weight (i.e. only one twin from the pair had a mean item score $\geqslant 4$ for the previous 3 months, while the other twin reported low importance of shape and weight), six twin pairs were concordant for high importance of shape and weight, and 299 twin pairs were concordant for low importance of weight and shape. All of the concordant pairs were MZ, while of the discordant pairs, 19 were MZ, 23 were DZ and one did not have zygosity reported.

GEE was used to investigate the association between threshold importance of shape and weight and lifetime disordered eating behaviours (see Table 2). No lifetime diagnostic threshold use of laxatives or diuretics was reported, while only one participant with threshold importance of shape and weight reported threshold levels of self-induced vomiting. Generally, the frequency of disordered eating behaviours was higher in those with threshold importance of shape and these associations were significant for fasting and excessive exercise. The proportion of females with any threshold lifetime eating disorder behaviour was considerably more common in the threshold importance of shape and weight group (26.8%) than in the non-threshold group (2.5%). Almost half (44.6%) of the females with threshold importance of shape and weight met our criteria for being overweight, compared with 11.7% in the non-threshold group, while this pattern of findings was reversed for the underweight criterion.

Within-twin associations between temperament variables and threshold importance of shape and weight

When examining the first endophenotype criterion (i.e. whether an association exists between candidate

Table 3. *Temperament associations with clinically significant importance of shape and weight in twins*

Independent variable	Importance of shape and weight		OR (95% CI)	*p*
	⩾4 (*n*=56) Mean (s.d.)	<4 (*n*=643) Mean (s.d.)		
Thin-ideal internalization	2.44 (0.95)	1.55 (0.64)	3.53 (2.58–4.84)[a]	<0.01
Ineffectiveness	2.68 (0.96)	1.82 (0.69)	3.07 (2.20–4.28)[a]	<0.01
Body dissatisfaction	3.87 (1.03)	2.30 (1.03)	3.51 (2.70–4.58)[a]	<0.01
Concern over mistakes	1.92 (0.41)	1.66 (0.43)	4.08 (2.18–7.62)[a]	<0.01
Sensitivity to punishment	2.72 (0.60)	2.24 (0.53)	5.29 (2.89–9.70)[a]	<0.01
Sensitivity to reward	2.32 (0.38)	2.17 (0.33)	3.39 (1.46–7.87)[a]	0.01
Obsessionality	0.93 (0.69)	0.62 (0.54)	2.19 (1.50–3.21)[a]	<0.01

s.d., Standard deviation; OR, odds ratio; CI, confidence interval.

[a] Significant predictors of threshold importance of shape and weight.

variables and threshold importance of shape and weight concern), GEE analyses revealed that all seven temperament variables were significantly positively associated with threshold importance of shape and weight. That is, higher levels of each of the temperament variables were associated with increased likelihood of threshold shape and weight concern. Results are summarized in Table 3.

Cross-twin associations between Twin 2 temperament variables and threshold importance of shape and weight

When examining the second endophenotype criterion using Twin 2 temperament scores as predictors, logistic regressions revealed five of the seven temperament variables were significantly positively associated with threshold importance of shape and weight in Twin 1. These findings are presented in Table 4, where it can be seen that sensitivity to reward and obsessionality approached significance (respective *p* values of 0.06 and 0.05). When Twin 1 scores were entered as a covariate and then Twin 2 temperament scores were entered, thin-ideal internalization, ineffectiveness, body dissatisfaction and sensitivity to punishment remained significant predictors of threshold importance of shape and weight in Twin 1 (see Table 4).

Heritability

In order to examine heritability, the MZ and DZ twin pair correlations, along with 95% CI, were examined for importance of weight and shape and the four variables identified in Table 4. In addition, the full ACE univariate models were examined and are reported in Table 5. It can be seen that heritability only accounted for a small amount of variance of the importance of weight and shape whilst the other four variables had two to five times as much variance accounted for by heritability. The correlation matrix for the additive genetic risk factors showed that all four temperament variables had genetic correlations with importance of weight and shape, ranging from 0.48 (ineffectiveness) to 0.95 (thin-ideal internalization).

Discussion

This study sought to investigate if any of the seven measured temperament risk factors for disordered eating met criteria for being an endophenotype for clinically significant importance of shape and weight. This was done by examining three of the criteria for endophenotypes (Gershon & Goldin, 1986), namely: association with illness in the general population; elevated levels in non-affected family members compared with the general population; and heritability. Whilst family studies have been used previously to investigate the importance of shape and weight (e.g. Reichborn-Kjennerud *et al*. 2004), these studies have been with adults rather than adolescents and thus the current study provides useful information about importance of shape and weight closer to the time at which it starts to emerge.

Threshold importance of shape and weight was chosen for examination as it is considered to be an important predictor of the development of later disordered eating behaviours. Whilst the cross-sectional design of our study could not address causality, we did find, consistent with previous research, a positive relationship between the presence of clinically significant importance of shape and weight and disordered eating behaviours (Gowers & Shore, 2001). Indeed, 26.8% of the 56 participants with threshold

Table 4. *Twin 2 temperament associations with clinically significant importance of shape and weight in Twin 1, both with and without controlling for Twin 1 scores*

	Twin 2 descriptives		Twin 2 associations with Twin 1 importance of shape and weight					
	Twin 2 means (s.d.) by Twin 1 importance of shape and weight scores		Not controlling for Twin 1 temperament score		Controlling for Twin 1 temperament score			
					Twin 1		Twin 2	
Twin 2 temperament	Twin 1 ⩾4 (n=48)	Twin 1 <4 (n=299)	OR (95% CI; W)	p	OR (95% CI)	p	OR (95% CI; W)	p
Thin-ideal internalization	2.00 (0.84)	1.51 (0.63)	2.35 (1.60–3.46; 18.9)[a]	<0.01	3.70 (2.42–5.67)[a]	<0.01	1.86 (1.21–2.84; 8.2)[a]	<0.01
Ineffectiveness	2.25 (0.87)	1.75 (0.66)	2.27 (1.56–3.31; 18.2)[a]	<0.01	2.46 (1.66–3.64)[a]	<0.01	1.64 (1.07–2.50; 5.2)[a]	0.02
Body dissatisfaction	3.15 (1.14)	2.22 (0.97)	2.31 (1.70–3.13; 28.6)[a]	<0.01	2.87 (2.04–4.02)[a]	<0.01	1.64 (1.14–2.37; 7.0)[a]	0.01
Concern over mistakes	1.82 (0.51)	1.65 (0.40)	2.62 (1.25–5.47; 6.54)[a]	<0.01	2.54 (1.23–5.27)[a]	0.01	2.11 (0.98–4.54; 3.7)	0.06
Sensitivity to punishment	2.58 (0.56)	2.24 (0.50)	3.54 (1.92–6.56; 16.2)[a]	<0.01	2.98 (1.57–5.68)[a]	<0.01	2.39 (1.24–4.58; 6.9)[a]	0.01
Sensitivity to reward	2.25 (0.32)	2.15 (0.35)	2.32 (0.96–5.58; 3.5)	0.06	2.71 (1.03–7.12)[a]	0.04	1.97 (0.79–4.90; 2.1)	0.15
Obsessionality	0.78 (0.63)	0.62 (0.54)	1.63 (0.99–2.66; 3.7)	0.05	1.89 (1.14–3.14)[a]	0.01	1.44 (0.86–2.41; 1.9)	0.17

s.d., Standard deviation; OR, odds ratio; CI, confidence interval; W, Wald statistic.

[a] Significant predictors of threshold importance of shape and weight.

Table 5. *Candidate endophenotype twin pair correlations for MZ and DZ twins*

| | | | Parameter estimates ×100 (95% CI) for full (ACE) univariate model[a] | | | Cholesky decomposition: correlations between genetic risk factors for each variable | | | | |
Independent variable	MZ correlation (95% CI)	DZ correlation (95% CI)	A	C	E	(1)	(2)	(3)	(4)	(5)
(1) Importance of weight/shape[b]	0.42 (0.29–0.53)	0.29 (0.15–0.44)	15 (0–48)	23 (0–43)	62 (52–73)	1.00	0.95	0.48	0.80	0.52
(2) Thin-ideal internalization	0.37 (0.24–0.49)	0.19 (0.04–0.34)	30 (0–54)	13 (0–41)	57 (46–69)	–	1.00	0.73	0.79	0.61
(3) Ineffectiveness	0.46 (0.34–0.57)	0.23 (0.07–0.37)	28 (0–51)	12 (0–39)	60 (49–72)	–	–	1.00	0.51	0.64
(4) Body dissatisfaction	0.56 (0.46–0.65)	0.36 (0.22–0.49)	35 (6–64)	23 (0–47)	42 (34–52)	–	–	–	1.00	0.90
(5) Sensitivity to punishment	0.44 (0.32–0.55)	0.24 (0.09–0.38)	71 (52–78)	0 (0–15)	29 (22–37)	–	–	–	–	1.00

MZ, Monozygotic; DZ, dizygotic; CI, confidence interval; A, additive genetic variance; E, non-shared environmental variance; C, shared environmental variance.

[a] Given lack of power to discriminate between the full model and the sub-models, only the parameter estimates for the full model are presented for each variable.

[b] Continuous measure.

importance of shape and weight met threshold criteria for at least one lifetime disordered eating behaviour, compared with just 2.5% of participants who did not meet threshold importance of shape and weight. Investigations of the weight status of those with threshold importance of shape and weight revealed almost half (44.6%) met our criteria for being over-weight, which is consistent with other findings that a history of being overweight in childhood (Micali *et al.* 2007) is a risk factor for later disordered eating, in particular of a bulimic phenotype. Conversely, there was a non-significant trend for a higher proportion of the non-threshold importance of shape and weight group meeting criteria for being underweight (4.2%) compared with those in the threshold group (1.7%). In sum, descriptive analyses in the current study provide further support for importance of shape and weight as having a strong association with disordered eating behaviours (Cooper & Goodyer, 1997; Goldfein *et al.* 2000).

The first endophenotype criterion we investigated was whether the temperament variables were associated with illness; in this case, presence of threshold importance of shape and weight. Findings indicated each of the candidate endophenotypes measured was significantly associated with threshold importance of shape and weight in the current study. The second criterion assessed was whether the temperament variables were found at an elevated level in non-affected family members (i.e. twins without threshold importance of shape and weight) compared with the general population. Our most rigorous test of non-affected family members showed four significant predictors of threshold importance of shape and weight in the affected twin – namely, thin-ideal internalization, ineffectiveness, body dissatisfaction and sensitivity to punishment. These Twin 2 variables predicted variance in Twin 1 importance of shape and weight, even when controlling for Twin 1 scores on these variables, suggesting that these variables are found at a significantly elevated level in non-affected family members compared with the general population. These findings provide support for conceptualizations of the development and maintenance of disordered eating that include one or more of these variables (e.g. Stice, 2001). They suggest that prevention and early intervention efforts that target one or more of these variables are likely to be of value and indeed some such prevention programmes, particularly those targeting thin-ideal internalization, have had success at reducing eating disorder risk (e.g. Wade *et al.* 2003). On a clinical level, these findings are also a reminder of the potentially important role of anxiety (represented by sensitivity to punishment) as a worthwhile target in the treatment of eating disorders. The third endophenotype criterion

evaluated was that of heritability. The amount of variance explained by genetic influences ranged from 28% (ineffectiveness) to 71% (sensitivity to punishment). A far lower proportion of variance was explained by genetic influences for importance of shape and weight (15%), suggesting this construct is more environmentally influenced, consistent with previous findings (Wade & Bulik, 2007). The heritability findings of the current study provide support for differentiation of the importance of shape and weight construct from the temperament variables investigated. It should be noted that one of the limitations with evaluating temperament candidate endophenotypes is that temperaments are genetically complex constructs where influences are difficult to simplify. However, the endophenotype criteria do provide a useful indication of the essential building blocks on the pathway to the development of clinically significant importance of shape and weight.

In summary, of the four endophenotypes identified, two related to general temperament relating to ineffectiveness and anxiety, and the other two related to body-specific temperament. This conceptualization is consistent with the model proposed by Peter Slade (Slade, 1982) where premorbid personal ineffectiveness and its associated anxiety, when allied with weight sensitivity triggered by adolescence, can result in coping strategies that focus on the importance of controlling weight, shape and eating in order to manufacture some sense of personal success.

Limitations

The results of the current study should be interpreted in the context of four limitations. First, the use of an abbreviated Sensitivity to Punishment and Sensitivity to Reward Questionnaire (Torrubia *et al.* 2001) may not be as psychometrically robust as the full scales and thus findings related to these scales should be interpreted with some caution. Second, only one dimension of perfectionism, concern over mistakes, was measured in the current study. Given that this candidate endophenotype did approach significance in our most robust test, it would have been of interest to have measured other perfectionism constructs such as personal standards, which has been found to be associated with disturbed eating behaviour (Shafran *et al.* 2006). Third, due to the cross-sectional nature of the current study, we were unable to investigate all of the previously stated criteria for an endophenotype (Gershon & Goldin, 1986) and thus future investigations will need to establish if the temperament endophenotypes identified in the current study meet the criterion of being a stable, state-independent characteristic. A further limitation is our moderate response

rate (48.8%), which is commensurate with other large Australian twin studies (Wade *et al.* 2006). A previous study of Australian twins using interviews focused only on eating indicated that response was not biased by previous eating problems (Wade *et al.* 1997).

Future direction and final comments

The identification of endophenotypes for the development of clinically significant importance of shape and weight, previously described as the 'core psychopathology' of eating disorders (Cooper & Fairburn, 1993), is a valuable endeavour given that endophenotypes are particularly useful in deconstructing complex disorders (Bulik *et al.* 2007) and that importance of shape and weight is considered to be an important diagnostic criterion across different eating disorders. The issue of complexity goes to the heart of the eating disorder field, where up to 30 variables have been reported in the literature as possible risk factors (Jacobi *et al.* 2004), and increased success in developing parsimonious formulations and models that indicate more helpful therapeutic directions are urgently required (Fairburn & Cooper, 2007).

To this end, the current study has identified four temperament endophenotypes for the development of clinically significant importance of shape and weight, namely, thin-ideal internalization, ineffectiveness, body dissatisfaction and sensitivity to punishment (anxiety). Future research should evaluate the utility of these endophenotypes for predicting significant growth in importance of shape and weight, and also whether different endophenotypes emerge when the importance of weight and shape reaches its peak in adolescents, around the age of 15–16 years (Cooper & Goodyer, 1997).

Acknowledgements

Declaration of Interest

None.

References

APA (1994). *Diagnostic and Statistical Manual of Mental Disorders*, 4th edn. American Psychiatric Publishing, Inc.: Washington, DC.

Bellodi L, Cavallini MC, Bertelli S, Chiapparino D, Riboldi C, Smeraldi E (2001). Morbidity risk for obsessive-compulsive spectrum disorders in first-degree relatives of patients with eating disorders. *American Journal of Psychiatry* **158**, 563–569.

Berrettini WH (2005). Genetic bases for endophenotypes in psychiatric disorders. *Dialogues in Clinical Neuroscience* **7**, 95–101.

Bryant-Waugh RJ, Cooper PJ, Taylor CL, Lask BD (1996). The use of the eating disorder examination with children: a pilot study. *International Journal of Eating Disorders* **19**, 391–397.

Bulik CM, Hebebrand J, Keski-Rahkonen A, Klump KL, Reichborn-Kjennerud T, Mazzeo S E, Wade TD (2007). Genetic epidemiology, endophenotypes, and eating disorder classification. *International Journal of Eating Disorders* **40** (Suppl.), S52–S60.

Bulik CM, Tozzi F, Anderson C, Mazzeo SE, Aggen S, Sullivan PF (2003). The relation between eating disorders and components of perfectionism. *American Journal of Psychiatry* **160**, 366–368.

Cassin SE, von Ranson KM (2005). Personality and eating disorders: a decade in review. *Clinical Psychology Review* **25**, 895–916.

Clifford CA, Murray RM, Fulker DW (1984). Genetic and environmental influences on obsessional traits and symptoms. *Psychological Medicine* **14**, 791–800.

Cole TJ, Flegal KM, Nicholls D, Jackson AA (2007). Body mass index cut offs to define thinness in children and adolescents: international survey. *British Medical Journal* **335**, 1–12.

Cooper PJ, Fairburn CG (1993). Confusion over the core psychopathology of bulimia nervosa. *International Journal of Eating Disorders* **13**, 385–389.

Cooper PJ, Goodyer I (1997). Prevalence and significance of weight and shape concerns in girls aged 11–16 years. *British Journal of Psychiatry* **171**, 542–544.

Cusumano DL, Thompson J (2001). Media influence and body image in 8–11 year-old boys and girls: a preliminary report on Multidimensional Media Influence Scale. *International Journal of Eating Disorders* **29**, 37–44.

Eaves LJ, Eysenck HJ, Martin NG, Jardine R, Heath AC, Feingold L, Young PA, Kendler KS (1989). *Genes, Culture and Personality: An Empirical Approach*. Oxford University Press: Oxford.

Fairburn CG, Cooper Z (1993). The Eating Disorder Examination, 12th edn. In *Binge Eating: Nature, Assessment, and Treatment* (ed. C. G. Fairburn and G. T. Wilson), pp. 317–360. Guilford Press: New York.

Fairburn CG, Cooper Z (2007). Thinking afresh about the classification of eating disorders. *International Journal of Eating Disorders* **40**, S107–S110.

Fairburn CG, Cooper Z, Shafran R (2003). Cognitive behaviour therapy for eating disorders: a 'transdiagnostic'

theory and treatment. *Behaviour Research and Therapy* **41**, 509–528.

Frost RO, Marten P, Lahart C, Rosenblate R (1990). The dimensions of perfectionism. *Cognitive Therapy and Research* **14**, 449–468.

Garner DM, Olmstead MP, Polivy J (1983). Development and validation of a multidimensional Eating Disorder Inventory for anorexia nervosa and bulimia nervosa. *International Journal of Eating Disorders* **2**, 15–34.

Gershon ES, Goldin LR (1986). Clinical methods in psychiatric genetics: I. Robustness of genetic marker investigative strategies. *Acta Psychiatrica Scandinavica* **74**, 113–118.

Goldfein JA, Walsh B, Midlarsky E (2000). Influence of shape and weight on self-evaluation in bulimia nervosa. *International Journal of Eating Disorders* **27**, 435–445.

Gowers SG, Shore A (2001). Development of weight and shape concerns in the aetiology of eating disorders. *British Journal of Psychiatry* **179**, 236–242.

Gray JA (1981). A model for personality. In *A Critique of Eysenck's Theory of Personality* (ed. H. J. Eysenck), pp. 246–276. Springer: New York.

Hollander E, Kim S, Khanna S, Pallanti S (2007). Obsessive-compulsive disorder and obsessive-compulsive spectrum disorders: diagnostic and dimensional issues. *CNS Spectrums* **12** (Suppl. 3), 5–13.

Jacobi C, Hayward C, de Zwaan M, Kraemer H C, Agras W (2004). Coming to terms with risk factors for eating disorders: application of risk terminology and suggestions for a general taxonomy. *Psychological Bulletin* **130**, 19–65.

Jimerson DC, Lesem MD, Kaye WH, Brewerton TD (1992). Low serotonin and dopamine metabolite concentrations in cerebrospinal fluid from bulimic patients with frequent binge episodes. *Archives of General Psychiatry* **49**, 132–138.

Keski-Rahkonnen A, Bulik CM, Neale BM, Rose RJ, Rissanen A, Kaprio J (2005). Body dissatisfaction and drive for thinness in young adult twins. *International Journal of Eating Disorders* **37**, 188–199.

Larsson H, Lichtenstein P, Larsson J (2006). Genetic contributions to the development of ADHD subtypes from childhood to adolescence. *Journal of the American Academy of Child and Adolescent Psychiatry* **45**, 973–981.

Lilenfeld LR, Kaye WH, Greeno CG, Merikangas KR, Plotnicov K, Pollice C, Rao R, Strober M, Bulik CM, Nagy L (1997). Psychiatric disorders in women with bulimia nervosa and their first-degree relatives: effects of comorbid substance dependence. *International Journal of Eating Disorders* **22**, 253–264.

Lilenfeld LR, Kaye WH, Greeno CG, Merikangas KR, Plotnicov K, Pollice C, Rao R, Strober M, Bulik CM, Nagy L (1998). A controlled family study of anorexia nervosa and bulimia nervosa: psychiatric disorders in first-degree relatives and effects of proband comorbidity. *Archives of General Psychiatry* **55**, 603–610.

Micali N, Holliday J, Karwautz A, Haidvogl M, Wagner G, Fernandez-Aranda F, Badia A, Gimenez L, Solano R, Brecelj-Anderluh M, Mohan R, Collier D, Treasure JL (2007). Childhood eating and weight in eating disorders: a multi-centre European study of affected women and their unaffected sisters. *Psychotherapy and Psychosomatics* **76**, 234–241.

Neale MC (1997). *Mx: Statistical Modeling*, 2nd edn. Medical College of Virginia, Department of Psychiatry: Richmond, VA.

Nes RB, Roysamb E, Reichborn-Kjennerud T, Harris JR, Tambs K (2007). Symptoms of anxiety and depression in young adults: genetic and environmental influences on stability. *Twin Research and Human Genetics* **10**, 450–461.

Raevuori A, Dick DM, Keski-Rahkonnen A, Pulkkinen L, Rose RJ, Rissanen A, Kaprio J, Viken RJ, Silventoinen K (2007). Genetic and environmental factors affecting self-esteem from age 14 to 17: a longitudinal study of Finnish twins. *Psychological Medicine* **37**, 1625–1633.

Reichborn-Kjennerud T, Bulik CM, Kendler KS, Roysamb E, Tambs K, Torgersen S, Harris JR (2004). Undue influence of weight on self-evaluation: a population-based twin study of gender differences. *International Journal of Eating Disorders* **35**, 123–132.

Schoemaker C, van Strien T, van der Staak C (1994). Validation of the Eating Disorders Inventory in a nonclinical population using transformed and untransformed responses. *International Journal of Eating Disorders* **15**, 387–393.

Shafran R, Lee M, Fairburn CG (2004). Clinical perfectionism: a case report. *Behavioural and Cognitive Psychotherapy* **32**, 353–357.

Shafran R, Lee M, Payne E, Fairburn CG (2006). The impact of manipulating personal standards on eating attitudes and behaviour. *Behaviour Research and Therapy* **44**, 897–906.

Slade PD (1982). Towards a functional analysis of anorexia nervosa and bulimia nervosa. *British Journal of Clinical Psychology* **21**, 167–179.

Steiger H, Gauvin L, Israel M, Koerner N, Ng Ying Kin N, Paris J, Young SN (2001). Association of serotonin and cortisol indices with childhood abuse in bulimia nervosa. *Archives of General Psychiatry* **58**, 837–843.

Stice E (2001). A prospective test of the dual-pathway model of bulimic pathology: mediating effects of dieting and negative affect. *Journal of Abnormal Psychology* **110**, 124–135.

Thordarson DS, Radomsky AS, Rachman S, Shafran R, Sawchuk CN, Hakstian A (2004). The Vancouver Obsessional Compulsive Inventory (VOCI). *Behaviour Research and Therapy* **42**, 1289–1314.

Torrubia R, Avila C, Molto J, Caseras X (2001). The Sensitivity to Punishment and Sensitivity to Reward Questionnaire (SPSRQ) as a measure of Gray's anxiety and impulsivity dimensions. *Personality and Individual Differences* **31**, 837–862.

Treasure J, Lopez C, Roberts M (2007). Endophenotypes in eating disorders: moving toward ecologically based diagnosis and treatment focussed on pathophysiology. *Pediatric Health* **1**, 171–181.

Tyrka AR, Waldron I, Graber JA, Brooks-Gunn J (2002). Prospective predictors of the onset of anorexic and

bulimic syndromes. *International Journal of Eating Disorders* **32**, 282–290.

van Strien T, Ouwens M (2003). Validation of the Dutch EDI-2 in one clinical and two nonclinical populations. *European Journal of Psychological Assessment* **19**, 66–84.

Wade T, Tiggemann M, Martin NG, Heath AC (1997). Characteristics of interview refusers: women who decline to participate in interviews relating to eating. *International Journal of Eating Disorders* **22**, 95–99.

Wade TD, Bulik CM (2007). Shared genetic and environmental risk factors between undue influence of body shape and weight on self-evaluation and dimensions of perfectionism. *Psychological Medicine* **37**, 635–644.

Wade TD, Byrne S, Bryant-Waugh R (2008). The Eating Disorder Examination: norms and construct validity with young and middle adolescent girls. *International Journal of Eating Disorders*. Published online: 23 April 2008. doi:10.1002/eat.20526.

Wade TD, Crosby RD, Martin NG (2006). Use of latent profile analysis to identify eating disorder phenotypes in an adult Australian twin cohort. *Archives of General Psychiatry* **63**, 1377–1384.

Wade TD, Davidson S, O'Dea JA (2003). A preliminary controlled evaluation of a school-based media literacy program and self-esteem program for reducing eating disorder risk factors. *International Journal of Eating Disorders* **33**, 371–383.