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The effect of disgust-related side-effects on symptoms of depression and anxiety in people treated for cancer: a moderated mediation model

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Abstract

As maladaptive disgust responses are linked to mental health problems, and cancer patients may experience heightened disgust as a result of treatments they receive, we explored the associations between disgust-related side-effects and symptoms of depression and anxiety in people treated for cancer. One hundred and thirty two (83 women, $M_{\rm age}$ = 57.48 years) participants answered questions about their treatments, side-effects, disgust responding, and mental health. Experiencing bowel and/or bladder problems, sickness and/or nausea (referred to here as "core" disgust-related side-effects) was significantly related to greater symptoms of depression and borderline increased anxiety. Further, these links were explained by a moderated mediation model, whereby the effects of core disgust side-effects on depression and anxiety were mediated by (physical and behavioural) self-directed disgust, and disgust propensity moderated the effect of core disgust side-effects on self-disgust. These findings stress the importance of emotional responses, like disgust, in psychological adaptation to the side-effects of cancer treatments.

Keywords: anxiety, cancer, depression, disgust, self-disgust, side-effects

Introduction

Cancer is a major public health problem and the leading cause of death in England and Wales (Office for National Statistics, 2015). The mental health of cancer patients has significant effects on their overall quality of life (e.g., Skarstein, Aass, Fosså, Skovlund, & Dahl, 2000), responses to treatment (e.g., DiMatteo, Lepper, & Croghan, 2000), and survival times (e.g., Falagas et al., 2007). While estimates vary, of the most common mental health problems, the prevalence of depression and anxiety in people with cancer has been conservatively estimated to be around twice that of the general population (Hinz et al., 2010; Walker et al., 2014). Common mental health problems, such as depression and anxiety, can be experienced as a consequence of cancer treatment regimens (e.g., Spiegel, 1997), as well as affecting how well cancer patients respond to them (DiMatteo et al., 2000).

Depression and anxiety in cancer patients are associated with decreased treatment adherence (e.g., Arrieta et al., 2013), greater healthcare costs and longer hospital stays (e.g., Hosaka, Aoki, Watanabe, Okuyama, & Kurosawa, 1999), lower chances of survival (e.g., Falagas et al., 2007), and an increased risk of suicide (e.g., Llorente et al., 2005). Given these and other significant consequences of common mental health problems in people with cancer (which may go unrecognised and untreated; Walker et al., 2014), being able to identify which patients may be particularly vulnerable to developing symptoms of depression and anxiety, and which factors may be important in their antecedence, is increasingly important, as is recognising psychological factors to target during psycho-therapeutic interventions.

Increasingly, one's emotional reactions are being understood as important, yet historically underexplored, predictors of mental health outcomes (Whelton & Greenberg, 2005). Experiencing cancer, and cancer treatment, can elicit multiple negative emotional responses (e.g., Kennifer et al., 2009). One prominent, yet under-researched, emotion relevant to the experience of cancer is disgust. Individuals with cancer potentially have to

confront a range of disgust-inducing stimuli, including, but not being limited to, becoming a "diseased" or "contaminating" object (Neal et al., 2007), sickness and nausea (e.g., Carey & Burish, 1988), bowel and bladder problems (e.g., Bauer, Bastian, Gozzi, & Stief, 2009), changes to an idealised body envelope (e.g., Bredin, 1999), and the salience of their own mortality and death (e.g., Goldenberg, Arndt, Hart, & Routledge, 2008). Moreover, the majority of these may not be caused by the cancer itself, but as side-effects of the treatments patients receive. A study in people with a colostomy, for example, demonstrated significant negative relationships between a bowel-related disgust measure and individuals' adjustment to colostomy and their overall life satisfaction (Smith, Loewenstein, Rozin, Sherriff, & Ubel, 2007). Further experimental work has shown that trait and state disgust indices may interact in deterring help-seeking behaviour for bowel-related symptoms (Reynolds, McCambridge, Bissett, & Consedine, 2014).

Maladaptive disgust responding (e.g., experiencing heightened levels of disgust) has been linked to mental health problems, including anxiety (Cisler, Olatunji, & Lohr, 2009) and depression (Alanazi, Powell, & Power, 2015), by a considerable literature (Davey, 2011). Thus, one potential yet underexplored link between the experience of cancer and depressive and/or anxious outcomes is individuals' maladaptive disgust responses, which may be associated with certain side-effects that are typical of cancer treatments (e.g., problems with body waste products, including incontinence, diarrhoea, and vomiting as a result of chemotherapy, or physical deformities as a consequence of surgery).

Disgust is a universal human emotion (Ekman, 1999), whose primary function is to protect us from the risk of disease (Curtis, Aunger, & Rabie, 2004). Through evolutionary exaptation, disgust has expanded from an oral inhibition response (which has adaptive value in protecting an organism from ingesting harmful substances; Rozin & Fallon, 1987), to an emotion that facilitates the avoidance and rejection of wider pathogenic stimuli, broader

threats to our biological fitness (e.g., unfavourable mates), and sociomoral transgressions that desecrate culturally-defined virtues of purity and divinity (e.g., sexual violations; Chapman & Anderson, 2012). The most influential psychological model of disgust to date was proposed by Rozin and Fallon (1987) and distinguishes four categories of disgust elicitors: 1) a "core" set that includes animals, food, and body waste products (e.g., faeces, vomit, urine); 2) an "animal-nature" domain that reminds of us of our base status as animals (e.g., poor hygiene, death, violations of an idealised body envelope); 3) "interpersonal" (or contamination) disgust for contact with other persons; and 4) "moral" disgust elicited by sociomoral violations (Rozin, Haidt, & McCauley, 2008). These theoretical divisions have been supported empirically, by factor analyses differentiating between core, animal-nature, and contamination-based disgusts (e.g., Olatunji, Williams, Lohr, Sawchuk, 2005; Olatunji et al., 2007), and via unique predictive associations with other related constructs, including mental health outcomes (e.g., Olatunji et al., 2007).

Given its evolutionary grounding as a cornerstone of our "behavioural immune system" (Schaller & Park, 2011), disgust responding has a particular relevance in understanding people's psychological reactions to disease and its treatment, not least significant threats to health, such as cancer (e.g., Reynolds, Consedine, Pizarro, & Bissett, 2013). Of the disgust categories outlined above, "core" (e.g., sickness, incontinence) and "animal-nature" (e.g., hair loss, scarring) disgust-eliciting side-effects are particularly relevant to cancer treatments. An important *mediational* pathway through which the disgust-related side-effects of cancer treatments may lead to increased depression and/or anxiety is via affect-congruent negative appraisals of the self. In particular, cancer patients may come to appraise themselves (or certain self-aspects) as disgusting, which may then lead to heightened symptoms of depression and/or anxiety (e.g., Beck, 1967; Powell, Overton, Simpson, 2013). Self-disgust has been shown to be a significant temporal antecedent of (i.e.,

a consistent vulnerability factor for) depressive symptoms (Overton, Markland, Taggart, Bagshaw, & Simpson, 2008; Powell et al., 2013), and has also been linked to anxious responding (Olatunji, Cox, & Kim, 2015). Furthermore, self-directed disgust has a hypothesised role in chronic physical health conditions, including cancer (Reynolds, McCambridge, & Consedine, 2015). Broadly, people can experience enduring self-disgust toward their physical bodies and/or their behavioural acts (Overton et al., 2008), with independent predictive effects on measures of psychopathology (e.g., Powell et al., 2013).

In addition, one may expect the psychological impact of these disgust-related physical side-effects to be influenced by an individual's underlying proneness to disgust (i.e., "disgust propensity"; van Overveld, de Jong, Peters, Cavanagh, & Davey, 2006), which is a trait individual difference factor that can be measured reliably by self-report (Olatunji, Cisler, Deacon, Connolly, & Lohr, 2007). A recent prospective study by Reynolds, Bissett, and Consedine (2015) in patients with anal incontinence, for example, found that proneness to disgust negatively predicted a number of quality of life indices assessed 3 months later. Thus, one's propensity to be disgusted can be hypothesised to be a critical *moderator* of the effects of disgust-related side-effects on self-directed disgust, and thus psychological health outcomes. In particular, we would expect the effects of disgust-related side-effects on self-disgust, and thus the mediational pathway described above, to be greater for those who were higher in disgust propensity.

In the present paper we tested the above mediation and moderation hypotheses in a community sample who reported having received treatment(s) for cancer. In particular, we sought to explore: i) whether experiencing two types of disgust-related physical side-effects (i.e., "core" and "animal-nature") were positively related to symptoms of depression and anxiety; ii) the degree that physical and/or behavioural self-disgust *mediated* the link between the presence of a disgust-related side-effect and depressive/anxious symptoms; and iii)

whether participants' underlying propensity to disgust significantly *moderated* the impact of experiencing a disgust-related side-effect on self-directed disgust, and thus any indirect effect on depression and anxiety. We predicted that:

- 1) Experiencing a disgust-related physical side-effect ("core" or "animal-nature") would be positively related to symptoms of depression and anxiety.
- 2) Physical and/or behavioural self-disgust would positively mediate the effect of having a disgust-related side-effect on depression and anxiety.
- 3) Trait propensity to disgust would positively moderate the effect of having a disgust-related physical side-effect on self-directed disgust.

Method

Participants

One hundred and thirty two community volunteers (83 women, $M_{\rm age} = 57.48$ years, $SD_{\rm age} = 14.19$ years) who had been treated for a broad range of cancers participated in this study. They were recruited from cancer charities and support groups. Table 1 illustrates study participants' characteristics. No reimbursement was provided for participation.

Measures

Cancer treatment and side-effects.

All participants reported having being treated for cancer. Participants were asked whether they had received a particular treatment (e.g., chemotherapy; coded as 0 = had not received, 1 = had received; see Table 1). They were also asked if their treatment(s) caused any side-effects, and if so to list them as a free-text response. These free-text responses were coded for i) the number of side-effects reported (continuous scale); and ii) whether the participant reported a "core" or "animal-nature" disgust side-effect (0 = no, 1 = yes). The decision to code a side-effect as related to "core" or "animal-nature" disgust was based on the conceptualisation provided by Rozin et al. (2008), and supported empirically by others (e.g.,

Olatunji et al., 2007), that core disgust elicitors are related to food/eating and body waste products, and animal-nature disgust elicitors are related to death, hygiene, and body envelope violations. Thus, a participant was coded as having experienced a "core" disgust-related side-effect if they reported any bowel and/or bladder problems or sickness and/or nausea. An "animal-nature" side-effect was noted if participants reported a change to their physical body envelope (i.e., exterior form), including visible infections. Examples of side-effects that were not coded as disgust-related included fatigue, pain, and motor problems.

Participants' free-text responses were coded independently by two raters, the primary study author and an independent graduate student who was paid for their time. The graduate student was unaffiliated with the current study and blind to its aims, and was directed to code the data using only the definitions given above. Interrater agreement was high, with a two-way absolute average-measures intraclass correlation coefficient (ICC; see Hallgren, 2012) of 1.00, 95% CI [1.00, 1.00], p < .001 (97% agreement), for the number of side-effects; Siegel and Castellan's (1988) kappa (κ) of .98, p < .001 (99% agreement) for "core" disgust side-effects; and $\kappa = 1.00$, p < .001 (100% agreement) for "animal-nature" side-effects. In the case of discrepancies, the independent secondary coding of the data was used for analyses.

Disgust propensity and sensitivity.

Participants' disgust propensity (how easily one is disgusted) and disgust sensitivity (how negatively these disgust experiences are appraised) were measured using the Disgust Propensity and Sensitivity Scale-Revised (DPSS-R; van Overveld et al., 2006). Participants rated their agreement to 16 items on a 5-point Likert scale (1 = never, 5 = always). An example disgust propensity item is "I experience disgust", and an example disgust sensitivity item is "It scares me when I feel nauseous". Disgust sensitivity was included as a control variable in the current analyses. Based on psychometric evaluation of the DPSS-R (Goetz, Cougle, & Lee, 2013; Olatunji et al., 2007), a recommended 10-item solution (six items for

propensity and four for sensitivity) was used for analyses. Both the disgust propensity, α = .78, and disgust sensitivity, α = .83, subscales demonstrated good internal reliability.

Self-disgust.

The Self-Disgust Scale (SDS; Overton et al., 2008) was used to measure participants' trait disgust for the self. For each of 18 items, participants rated their agreement on a 7-point Likert scale (1 = $strongly \ agree$, 7 = $strongly \ disagree$). The scale contains a number of filler items and two 5-item subscales, one measuring physical self-disgust (e.g., "I find myself repulsive") and the other behavioural self-disgust ("I often do things I find revolting"). The internal reliabilities of the two subscales in the current sample were excellent, with $\alpha = .90$ and $\alpha = .83$, respectively.

Depression and anxiety.

Participants' levels of anxiety and depression were measured using the Hospital Anxiety and Depression Scale (HADS; Zigmond & Snaith, 1983). The scale was developed for use amongst hospital inpatients and has been previously validated in community patients with cancer (e.g., Smith et al., 2002). For 14 items (seven for each subscale) participants rated their current agreement on a 4-point scale (0-3, with reversed labels). An example item from the anxiety subscale is "I get sudden feelings of panic", and an example from the depression subscale is "I feel as if I am slowed down". In the current sample, the Cronbach's alpha coefficients for the anxiety, $\alpha = .88$, and depression, $\alpha = .84$, subscales were good.

To maintain consistency with prior research on disgust in the context of depression (e.g., Overton et al., 2008; Powell et al., 2013), and to account for potential criticisms of the sensitivity of the HADS depression subscale (see Luckett et al., 2010), a second measure of depression was included in the surveys, the 7-item depression subscale of the short-form Depression, Anxiety, and Stress Scales (DASS–21; Lovibond & Lovibond, 1995). For each item (e.g., "I felt that life was meaningless") participants indicated how much it had applied

to them over the previous week on a 4-point Likert scale ($0 = did \ not \ apply \ to \ me \ at \ all$, $3 = applied \ to \ me \ very \ much, \ or \ most \ of \ the \ time$). Summative scores were multiplied by two to make them comparable with the extended form of the DASS (Lovibond & Lovibond, 1995). The internal consistency of this scale was excellent, $\alpha = .94$.

Control variables.

In our path analyses we controlled for observed variables that we expected, a priori, to have a significant relationship with symptoms of depression and anxiety, including gender, age, years since cancer diagnosis, number of side-effects reported, disgust sensitivity, and whether the participant reported a current medical diagnosis of depression (0 = no, 1 = yes) or anxiety (0 = no, 1 = yes). Bivariate correlations showed that all control variables were significantly related to at least two of the three outcome variables (rs = .18 - .46). Analyses without the inclusion of control variables produced the same (stronger) pattern of results.

Procedure

Ethical approval was granted by the host research institution prior to data collection. All procedures were in accordance with the ethical standards of the institutional research committee and with the 1964 Helsinki declaration and its later amendments. Informed consent was obtained from all individual participants included in the study. As part of a larger survey into psychological responses to cancer, cancer charities and support groups were approached with a link to an online survey. Volunteers completed the measures listed above online in a counterbalanced order and were fully debriefed.

Data Analysis

Missing data were minimal, with a single missing value on the depression subscale of the DASS (Lovibond & Lovibond, 1995). To minimise data loss, we imputed this value at the mean of available data. Following descriptive and correlational analyses on SPSS v. 21 (IBM Corp., Armonk, NY, US), path analysis on AMOS v. 22 (IBM Corp., Armonk, NY,

US) was used to model the hypothesised relationships between the variables. Path analysis has several advantages over standard multiple regression, including the estimation of direct and indirect effects (through mediating variables) simultaneously; the ability to model multiple endogenous (i.e., dependent) variables at the same time, allowing one to account for their interdependence caused by extraneous variables (by correlating their error terms); and the calculation of multiple measures of fit to the data. Further, it is a less resource-intensive technique than structural equation modelling (SEM), which typically requires a larger sample, due to the inclusion of latent variables (Wolf, Harrington, Clark, & Miller, 2013).

As recommended by Hayes (e.g., Hayes, 2009; Hayes & Scharkow, 2013), bias-corrected percentile bootstrapping was used to produce robust confidence intervals and standard errors (and hence probability values) for all estimates, including direct and indirect effects, removing any restrictions on the underlying sampling distribution. Bootstrapping provides a non-parametric robust alternative to traditional parametric estimates, when those estimates may be biased due, for example, to the violation of parametric assumptions and/or a restricted sample size (Fox, 2008). Ten thousand resamples were used for the bootstrapped estimates (Mallinckrodt, Abraham, Wei, & Russell, 2006). The Bollen and Stine (1992) bootstrap adjusted *p*-value was interpreted to assess model fit based on the chi-square statistic, along with the CFI and RMSEA. After fitting the model, to obtain separate bootstrapped estimates of specific indirect effects through a single mediator (not provided by default in AMOS), the path from the predictor to the alternative mediator was constrained to zero, as was the correlation between the two mediators (MacKinnon, 2008). To reduce potential multicollinearity associated with the inclusion of an interaction term, all continuous predictors (and mediators) were centred prior to analysis.

Results

Descriptives and Correlations

Initial bivariate correlations and descriptive data for the primary study variables are presented in Table 2. One hundred and nineteen participants (90%) reported at least one sideeffect from their treatment(s). The average number of side-effects reported per participant was 3.36 (SD = 2.60). Forty-five (34.1%) participants reported a "core" disgust side-effect, and 55 (41.7%) an "animal-nature" disgust side-effect, from their treatment(s). The results from the HADS are broadly similar to those reported for other heterogeneous cancer samples (e.g., Smith et al., 2002), with higher scores for anxiety than depression. In a large varied cancer sample, Smith et al. (2002) reported a mean of 4.38 for depression and 6.05 for anxiety, with 33.3% of participants presenting with clinically-relevant symptoms of anxiety and 19.8% with clinically-relevant symptoms of depression (Zigmond & Snaith, 1983). Using the same criteria (scores ≥ 8 indicating potentially clinically-relevant symptoms; Zigmond & Snaith, 1983), 19.7% of our sample presented with current indicative symptoms of depression, but approximately 50.0% presented with indicative symptoms of anxiety. The DASS has been used less in cancer samples (Luckett et al., 2010), but indicated a potentially higher level of depressive symptoms in the participants over the previous week. Using the criteria of Lovibond and Lovibond (1995), 37.9% presented with mild-to-severe symptoms of depression using the DASS (scores ≥ 10), suggesting it may be a more sensitive measure of, at least, mild depressive states (Luckett et al., 2010).

Partial support was found for prediction (1); total number of side-effects reported was significantly bivariately-related to symptoms of depression (HADS), r = .19, p < .05, and depression (DASS), r = .18, p < .05, but not anxiety, r = .14, p = .111. Reporting a core disgust side-effect was significantly positively related to symptoms of depression (HADS), $r_{\rm pb} = .23$, p < .01, and depression (DASS), $r_{\rm pb} = .27$, p < .01, and had a borderline significant relationship with anxiety, $r_{\rm pb} = .16$, p < .10. However, reporting an animal-nature side-effect was not related to depression (HADS), $r_{\rm pb} = -.03$, p = .716, depression (DASS), $r_{\rm pb} = -.11$, p = .11, p = .11

= .219, or anxiety, $r_{\rm pb}$ = -.06, p = .532, nor was it significantly related to any of the measured disgust traits. Accordingly, only the core disgust-related side-effect variable was included in the path analyses below. Experiencing a core disgust side-effect was significantly positively related to physical, $r_{\rm pb}$ = .20, p < .05, and behavioural, $r_{\rm pb}$ = .22, p < .05, self-disgust, and borderline significantly related to disgust sensitivity, $r_{\rm pb}$ = .16, p < .10. All the proposed mediating and outcome variables were significantly related (p < .001).

Path Models

Model 1 (mediation).

We estimated two path models to test predictions (2) and (3). The first was a mediation model, with core disgust side-effects and disgust propensity as the exogenous predictors, physical and behavioural self-disgust as hypothesised mediators, and depression (HADS), depression (DASS), and anxiety as outcomes (Figure 1). In this model, regression weights on the disgust propensity*core side-effect interaction term were constrained to zero. The model fit the data reasonably well, $\chi^2(5) = 9.45$, p = .174; CFI = 0.99; RMSEA = 0.08, 90% CI [0.00, 0.16], p = .210. All path estimates and associated maximum likelihood and bootstrap SEs/CIs are presented in Table 3. Reporting a core disgust side-effect significantly predicted behavioural, β_{a_2} = .17, p < .05, but not physical, β_{a_1} = .14, p = .119, self-disgust. Physical self-disgust significantly predicted depression (HADS), β_{b_1} = .57, p < .001, depression (DASS), β_{b_2} = .42, p < .01, and anxiety, β_{b_3} = .27, p < .05. Similarly, behavioural self-disgust significantly predicted depression (DASS), β_{b_5} = .27, p < .05, and anxiety, β_{b_6} = .23, p < .05, but not current depression as measured by the HADS, β_{b_4} = .06, p = .566. The direct effects of experiencing a core disgust side-effect on depression (HADS), β_{c_1} = .07, p = .287, depression (DASS), β_{c_2} = .11, p = .129, and anxiety, β_{c_3} = -.01, p = .997, were not significant. The indirect effect of core disgust side-effects through physical self-

disgust was borderline significant for depression (DASS), $\beta_{a_1b_2}$ = .06, p < .10, and anxiety,

 $\beta_{a_1b_3}$ = .04, p < .10, but not significant for depression (HADS), $\beta_{a_1b_1}$ = .08, p = .105, while the indirect effect of core disgust side-effects via behavioural self-disgust was statistically significant for depression (DASS), $\beta_{a_2b_5}$ = .05, p < .05, and anxiety, $\beta_{a_2b_6}$ = .04, p < .05, but not depression (HADS), $\beta_{a_2b_4}$ = .01, p = .383. The total indirect effect of core disgust side-effects through self-disgust was borderline significant for depression (HADS), $\beta_{a_{1,2}b_{1,4}}$ = .09, p < .10, depression (DASS), $\beta_{a_{1,2}b_{2,5}}$ = .11, p < .10, and anxiety, $\beta_{a_{1,2}b_{3,6}}$ = .08, p < .10.

Model 2 (moderated mediation).

Second, we estimated a moderated mediation model, by removing the parameter constraints on the disgust propensity*core disgust-related side-effect interaction term (Figure 2). This model fit the data very well, $\chi^2(3) = 2.82$, p = .530, CFI = 1.00, RMSEA = 0.00, 90% CI [0.00, 0.14], p = .555, showing a significant improvement in overall model fit over the constrained model, $\Delta \chi^2(2) = 6.63$, p < .05. The interaction term significantly positively predicted behavioural self-disgust, $\beta_{a_6} = .25$, p < .05, and borderline-significantly predicted physical self-disgust, $\beta_{a_5} = .18$, p < .10. To clarify the nature of the moderating effect, the effects of experiencing a core disgust-related side effect on (physical and behavioural) self-disgust were estimated at three levels of disgust propensity (-1.5D, M, +1.5D). As shown in Figure 3, having a core disgust side-effect significantly predicted behavioural self-disgust at high, $\beta_{a_2} = .36$, p < .05, but not low or moderate levels of disgust propensity. This pattern was the same for physical self-disgust. Tests of mediation (model 1) at differing levels of disgust propensity reflected the above; self-disgust significantly mediated the relationship between core disgust side-effects and depression and anxiety only when disgust propensity was high, providing evidence of moderated mediation (see Table 4).

Discussion

In this paper we explored the relationship between disgust-related physical sideeffects of cancer treatments and individuals' symptoms of depression and anxiety. Our first prediction (1), that disgust-related side-effects would be positively related to mental health symptoms, received partial support. Reporting a "core" (e.g., incontinence), but not "animalnature" (e.g., hair loss) disgust side-effect (vs. having no or any other type of side-effect) was positively related to symptoms of depression and anxiety. There are at least three reasons why disgust-related side-effects may be especially associated with states of depression and/or anxiety. First, disgust has been implicated in the phenomenology of depressed mood, which is theorised to be a combination of sadness and self-disgust (Alanazi et al., 2015; Power & Dalgleish, 2008). Second, disgust is socioculturally-determined (Rozin et al., 2008), and in experiencing disgust there is an implicit appraisal that the same stimulus would be disgusting to others. Individuals experiencing self-disgust think that they are repulsive to other people (Powell, Overton, & Simpson, 2014). Disgust causes a negative interpretation bias (Davey, Bickerstaffe, & MacDonald, 2006), motivates avoidance (Rozin & Fallon, 1987), and is associated with stigma (Park, Schaller, & Crandall, 2007). Vartanian (2010), for example, found disgust to be a stronger predictor of obesity stigma in undergraduate students than attributions of control or demographic factors (e.g., body mass index). Thus, disgust-related side-effects may increase the perceived (and actual) prejudicial discrimination patients receive from others, and/or increase social isolation (Powell et al., 2014), leading to heightened anxiety and/or depression (e.g., Else-Quest, LoConte, Schiller, & Hyde, 2009). Third, disgust is linked to other negative self-conscious emotional states (e.g., shame; Alanazi et al., 2015; Powell et al., 2014), and dysfunctional thought processes (Overton et al., 2008), which are linked to depression and anxiety (e.g., Gilbert, 2000).

It is unclear, however, why the animal-nature disgust side-effects did not show the same pattern of results as core disgust side-effects in this sample. It may be that there is a greater range of side-effects in this grouping (e.g., "ulceration" to "facial reconstruction"), than in core disgust side-effects, which constitute a more narrowly-defined class of elicitors

(i.e., bodily waste products), thus obscuring any effects at the aggregate level. Further work with larger samples may be able to explore this heterogeneity in greater detail. Further, it may be that the animal-nature category suffered from a lack of validity (i.e., our sample of cancer patients did not actually find these side-effects disgusting). Certain animal-nature side-effects (e.g., hair loss) may be more accepted socially, particularly in the context of cancer, than core disgust side-effects (e.g., incontinence; Reeve, 2015). Furthermore, disgust is heterogeneous (Simpson, Carter, Anthony, & Overton, 2006); core disgust stimuli (e.g., faeces) are considered some of the most salient and universal disgust-provoking objects in existence (Rozin et al., 2008). They elicit disgust in almost all adults, while animal-nature disgust shows greater variability (Haidt, 2015). This factor too may contribute to the stronger impact of core (vs. animal-nature) disgust side-effects on mental health.

In general, predictions (2) and (3) were supported through a moderated mediation model (Figure 2). Regarding (2), the impact of core disgust side-effects on depression and anxiety were fully explained by indirect effects through self-directed disgust, significantly predicting two out of three outcome variables (DASS depression and HADS anxiety). The specific indirect effects were larger through behavioural than physical self-disgust, presumably representing the nature of the core disgust category as measured in this study (i.e., problems with bodily waste products, including incontinence and sickness, typically constitute disgusting *behaviours* rather than necessitate alterations to the physical self). That a significant mediation effect was observed for only one of the depression indices (DASS) included in the study, and not the HADS depression subscale (although this effect was borderline), raises questions about measurement. First, the HADS asked about current (i.e., "in-the-moment") symptoms of depression, while the DASS measures symptoms over the *previous week*, which may help to explain this discrepancy. Second, the HADS has been criticised as having a reduced sensitivity to minor depression (Luckett et al., 2010). Thus, the

DASS may be a more sensitive instrument than the HADS, yet its use in cancer samples remains limited. Of course, the way self-report measures such as this may translate to clinical outcomes should be interpreted with caution.

Supporting prediction (3), disgust proneness positively moderated the effect of experiencing core disgust side-effects on self-disgust. This is consistent with prior work in the field, showing a potentially detrimental interactive effect between individuals' disgust proneness and exposure to one's core bodily disgust elicitors on psychological well-being (Reynolds, Bissett et al., 2015; Smith et al., 2007). This significant interaction also supports the contention that it is the disgust(ing) aspect of these physical side-effects that contributes to the development of trait self-disgust, and thereby symptoms of depression and anxiety.

As well as depression (Overton et al., 2008) and anxiety (Olatunji et al., 2015), self-disgust has been shown to be involved in a number of, often comorbid, mental health issues, including eating (Moncrieff-Boyd, Allen, Byrne, & Nunn, 2014) and personality (Abdul-Hamid, Denman, & Dudas, 2014) disorders. A model of enduring, problematic self-disgust as an "emotion schema" was outlined by Powell, Simpson, and Overton (2015a). Powell et al. (2015a) described self-disgust as a lasting disgust-based cognitive-affective orientation toward (an aspect of) the self, composed of interacting state and higher-order trait components, the latter of which are largely cognitive and persist over time (i.e., "my body is revolting"). Importantly, self-disgust has been shown to be relatively temporally stable (Powell et al., 2013), and thus is likely to be particularly detrimental based on its schematic, top-down influence on information processing (Powell et al., 2015a). Self-disgust is formed either during the acquisition of disgust, as one learns that certain self-aspects fit within an emerging repertoire of disgust elicitors, or through a dramatic change in the self (e.g., via physical trauma) that is then appraised as disgusting (Powell et al., 2015a). The latter would explain self-disgust as a consequence of disgust-related side-effects of cancer treatment.

The findings from this study have implications for understanding patients' psychological adaptation to cancer treatments, and they add to a literature on the importance of addressing emotional factors in the aetiology of mental health problems (Greenberg, 2008). This work suggests that the core disgust side-effects possible in cancer treatments may be particularly deleterious to psychological wellbeing through increases in (behavioural) self-directed disgust. There are two potential points for intervention. First, it is possible to identify, by measuring disgust proneness, which patients may particularly suffer as a result of these side-effects, and to monitor and treat them accordingly (Reynolds, Bissett et al., 2015). There is evidence that cognitive reappraisal (vs. e.g., affective suppression) may be a useful strategy for the psychological regulation of disgust, and that it can be primed in the laboratory (e.g., Goldin, McRae, Ramel, & Gross, 2008; Gross, 1998). Second, reducing the enduring self-disgust associated with core disgust side-effects may disrupt the link with depressive and anxious outcomes. Recent experimental work has shown that the self-affirmation of valued character traits may be a promising tool for reducing in-the-moment feelings of self-directed disgust (Powell, Simpson, & Overton, 2015b). Paul Gilbert's "compassion-focused therapy" is another approach formulated to work with people with high levels of self-criticism, hatred, and disgust (Gilbert, 2015). Nevertheless, research on the effective regulation and treatment of disgust is in its infancy and there are plentiful opportunities for future work in this area; what is critical, in this context, is that the emotional (i.e., disgust) component forms the primary target for clinical intervention (Whelton & Greenberg, 2005).

This study is limited by its cross-sectional design. While mediation normally requires a temporal lag between observed variables, a strong theoretical justification can be made for the proposed directionality of the current model. In particular, treatment side-effects can be viewed as primarily exogenous to self-disgust (i.e., they are more likely to cause self-disgust than the reverse; a pathway that is moderated by disgust proneness). In turn, previous

longitudinal work has shown that self-disgust (as measured by the SDS; Overton et al., 2008) significantly predicts depressive symptoms (as measured by the DASS; Lovibond & Lovibond, 1995) – and not the reverse – over 6- and 12-month periods (Powell et al., 2013). Given the strong relationship observed between symptoms of depression (DASS) and anxiety, r = .62, p < .001, we assume this directional pattern to be analogous for anxiety. Nevertheless, reverse effects to those hypothesised are possible; the side-effects measure is subjective and limited by recollection, so it is plausible that people who had higher levels of self-disgust at data collection were more likely to recall disgust-related side-effects. Further, disgust propensity may be affected by self-disgust (and mental health) as much as the reverse. Experimental designs (i.e., focusing on reducing disgust) are necessary to establish causation and test between the above alternative explanations.

This study is also limited by its modest sample size. However, this sample exhibited very similar levels of depressive symptoms (HADS) to a larger heterogeneous cancer sample (e.g., Smith et al., 2002), suggesting a degree of representativeness. While all cell sizes for the dummy side-effect variables were above a commonly-accepted minimum (30 cases), the reduction in cell sizes that would occur by further subsampling by side-effect type restricts the utility of such an analysis (Gordon, 2010). The study is also limited by our aggregated, "top-level" analysis of physical side-effects. While we control for some important factors (e.g., years since cancer diagnosis), we do not have available deeper information on participants' side-effects, such as their temporal permanency or recency, which may be key moderators of the effects observed. Nevertheless, the fact that an effect is obtained at the "top-level" of these events (i.e., experiencing a core disgust side-effect or not) is impressive and speaks to the enduring power of acquired disgust, which is particularly resistant to temporal extinction (Olatunji, Forsyth, & Cherian, 2007). Future work may explore how the

above (and other) conditioning variables influence the impact of core disgust side-effects and how patients' disgust appraisals change as a result.

This study was the first to explore quantitatively the effects of disgust-related physical side-effects of cancer treatment on mental health outcomes. The findings suggest that disgust matters; people who had experienced a core disgust side-effect (vs. no or any other kind of side-effect) exhibited higher levels of depression and anxiety. Furthermore, this association was explained entirely by increased self-directed disgust, as moderated by trait disgust proneness. Taken together, these findings stress the importance of emotional factors (i.e., disgust) in psychological adaptation to the side-effects of cancer treatment. They suggest that disgust-targeted interventions may be useful in reducing self-disgust and/or disgust proneness in individuals exposed to relevant side-effects in order to improve mental health outcomes.

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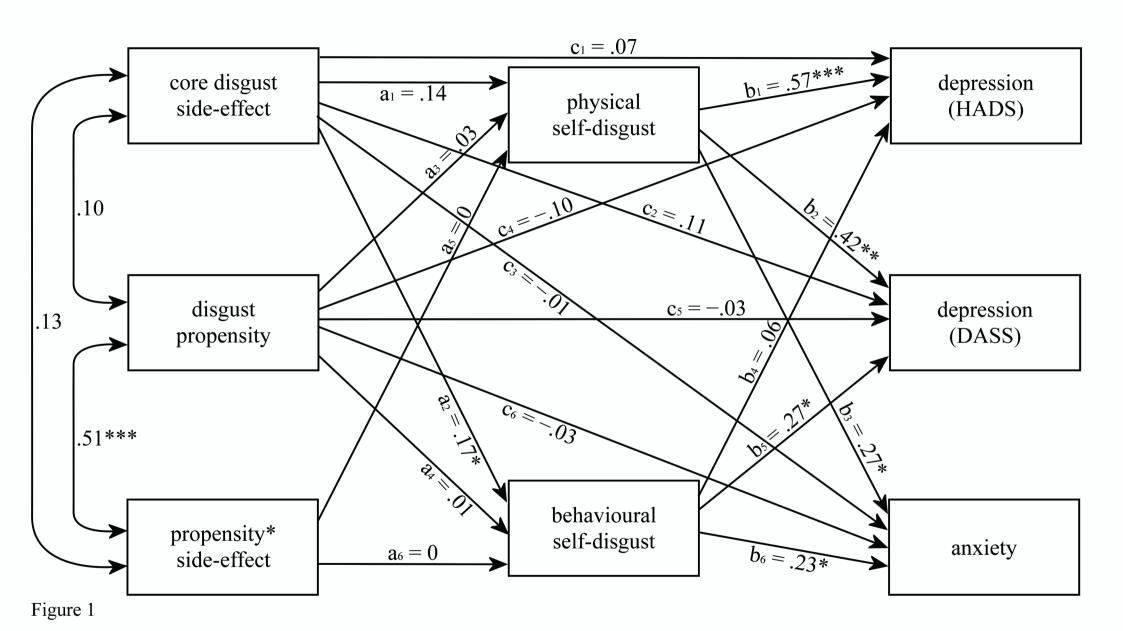
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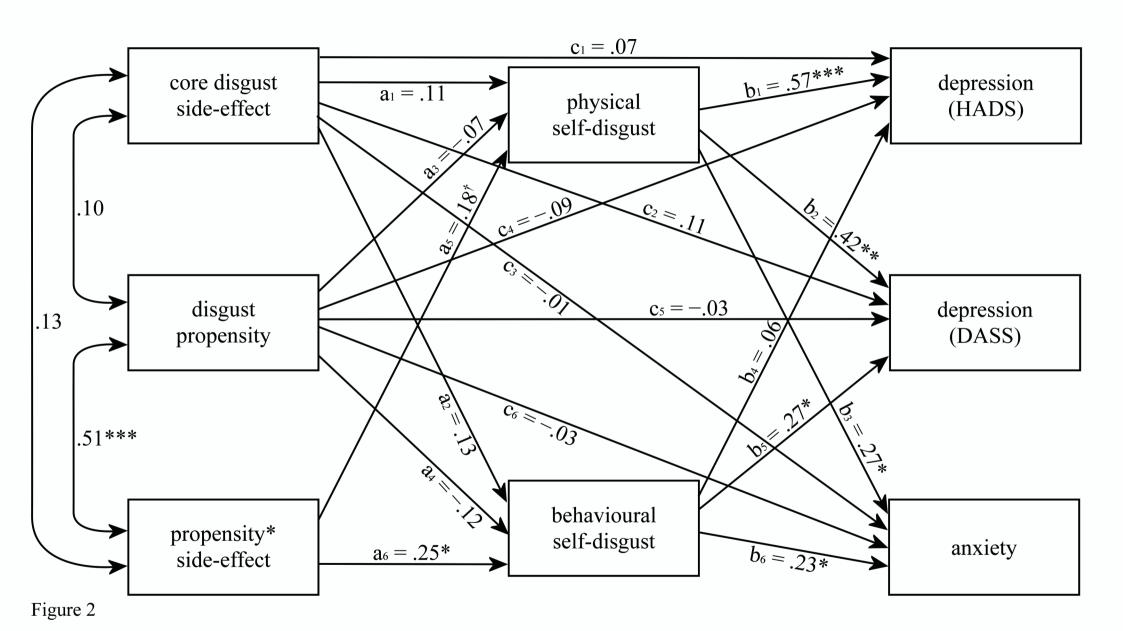
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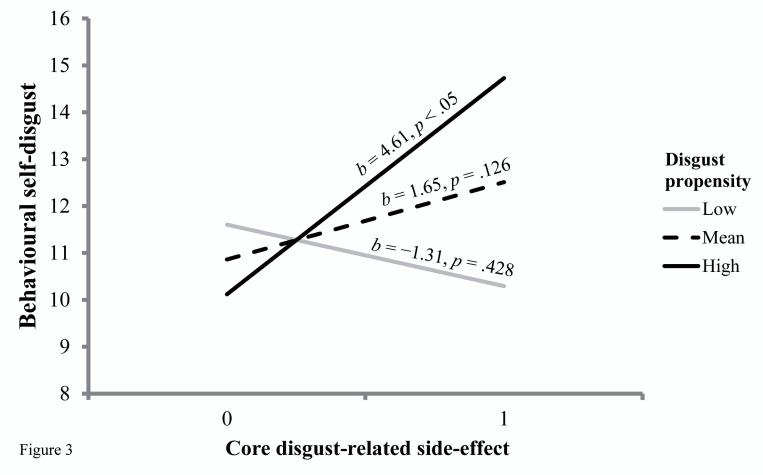


Figure Captions

Figure 1. Mediation model explaining the effect of "core" disgust side-effects on symptoms of depression and anxiety in people treated for cancer. Self-disgust significantly mediated the effect of experiencing a core disgust side-effect on symptoms of mental health, with significant indirect effects through behavioural self-disgust on depression (DASS), $\beta_{a_2b_5}$ = .05, p < .05, and anxiety, $\beta_{a_2b_6}$ = .04, p < .05; and borderline-significant indirect effects through physical self-disgust on depression (DASS), $\beta_{a_1b_2}$ = .06, p < .10, and anxiety, $\beta_{a_1b_3}$ = .04, p < .10. Regression coefficients associated with the interaction term (a_{5-6}) were constrained to zero. Control variables and error terms are omitted for clarity. Estimates on the endogenous variables were conditioned on: gender, age, years since diagnosis, number of side-effects reported, disgust sensitivity, and medical diagnoses of depression or anxiety. Error terms for the pair of mediators (physical and behavioural self-disgust) were correlated, as were the error terms for the three outcome variables (depression and anxiety). All estimates are standardised betas (β). Significance levels were determined based on bootstrapped CIs (10000 resamples). A chi-square test with Bollen-Stine bootstrap indicated adequate model fit, $\chi^2(5) = 9.45$, p = .174; CFI = 0.99; RMSEA = 0.08, p = .210. *p < .05. **p < .01. ***p < .001.

Figure 2. Moderated mediation model explaining the effect of "core" disgust side-effects on symptoms of depression and anxiety in people treated for cancer. Propensity to disgust significantly positively moderated the effect of experiencing a core disgust side-effect on behavioural self-disgust, β_{a_6} = .25, p < .05, and had a borderline-significant moderation effect on physical self-disgust, β_{a_5} = .18, p < .10. Control variables and error terms are omitted for clarity. Estimates on the endogenous variables were conditioned on: gender, age, years since diagnosis, number of side-effects reported, disgust sensitivity, and medical diagnoses of depression or anxiety. Error terms for the pair of mediators (physical and behavioural self-disgust) were correlated, as were the error terms for the three outcome variables (depression and anxiety). All estimates are standardised betas (β). Significance levels were determined based on bootstrapped CIs (10000 resamples). A chi-square test with Bollen-Stine bootstrap indicated excellent model fit, $\chi^2(3) = 2.82$, p = .530, CFI = 1.00, RMSEA = 0.00, 90% CI [0.00, 0.14], p = .555. $^{\dagger}p$ < .10. * $^{\dagger}p$ < .05. ** $^{\dagger}p$ < .01. ** $^{\dagger}p$ < .001.

Figure 3. Simple slopes analysis explaining the effect of experiencing a "core" disgust-related side-effect on levels of behavioural self-disgust at three levels of underlying disgust propensity (-1 SD = "low"; M = "mean"; & +1 SD = "high"). Experiencing a core disgust side-effect significantly predicted greater behavioural self-disgust only when disgust propensity was high. This pattern of results is the same for physical self-disgust (not shown).

Table 1

Participant characteristics

Variable	
Gender	49 men (37.1%); 83 women (62.9%)
Age	Range = 19-85; $M = 57.48$; $SD = 14.19$
Years since diagnosis	Range = 0-21; $M = 5.42$; $SD = 4.87$
Primary cancer diagnosis ^a	1 blood (0.7%); 3 bone (2.1%); 10 brain (7.1%); 38 breast (27.1%); 6 colorectal
	(4.3%); 24 gynaecological (17.1%); 29 head and neck (20.7%); 4 lung (2.9%); 17
	prostate (12.1%); 4 skin (2.9%); 1 stomach (0.7%); 1 testicular (0.7%); 2
	unspecified (1.4%)
Treatment ^a	6 biological therapy (4.5%); 59 chemotherapy (44.7%); 41 hormonal therapy
	(31.1%); 78 radiotherapy (59.1%); 1 stem cell/bone marrow transplant (0.8%); 107
	surgery (81.1%)
Diagnosed as depressed	12 yes (9.1%); 120 no (90.9%)
Diagnosed as anxious	14 yes (10.6%); 118 no (89.4%)

Note. N = 132. ^aAbsolute values greater than N because five participants reported more than one type of primary cancer and the majority of participants had received multiple treatments for their cancer.

Table 2

Means (M), standard deviations (SD), range, and inter-correlations of study variables

	1	2	3	4	5	6	7	8	9	10
1. Number of side-effects										
2. Core disgust side-effect	.42***	_								
3. Animal nature side-effect	.53***	.14	_							
4. Physical self-disgust	.20*	.20*	.05	_						
5. Behavioural self-disgust	.16 [†]	.22*	.02	.78***	_					
6. Disgust propensity	.10	.10	.07	.28**	.24**	_				
7. Disgust sensitivity	.23**	$.16^{\dagger}$.14	.38***	.36***	.65***				
8. Anxiety	.14	$.16^{\dagger}$	06	.60***	.58***	.30***	.46***	_		
9. Depression (HADS)	.19*	.23**	03	.72***	.60***	.27**	.40***	.66***		
10. Depression (DASS)	.18*	.27**	11	.70***	.66***	.29**	.40***	.62***	.77***	_
Range	0-12	0-1	0-1	5-35	5-31	6-30	4-20	0-19	0-21	0-42
M	3.36	0.34	0.42	13.04	11.12	14.33	8.51	7.52	4.88	8.93
SD	2.60	0.48	0.49	7.66	6.08	3.88	3.65	4.71	4.03	10.34

Note. N = 132. Correlations represent Pearson's r, pointbiseral (r_{pb}) , or phi (r_{ϕ}) coefficients. HADS = Hospital Anxiety and Depression Scale; DASS = Depression Anxiety and Stress Scales. $^{\dagger}p < .10. *p < .05. **p < .01.$ ***p < .001.

Table 3

Path estimates for the mediation model (Figure 1)

Path	Estim	Estimates		E of b	Bootstrap	95% CIs <i>b</i>	Bootstrap 95% CIs β	
	b	β	ML	Bootstrap	Lower	Upper	Lower	Upper
Direct pa	ath estimates							
a_1	2.27	.14	1.34	1.49	-0.57	5.28	04	.33
a_2	2.19*	.17	1.11	1.15	0.03	4.52	.00	.35
a_3	0.06	.03	0.20	0.22	-0.37	0.47	19	.23
a_4	0.01	.01	0.17	0.19	-0.36	0.38	24	.23
b_1	0.30***	.57	0.05	0.06	0.20	0.41	.37	.76
b_2	0.56**	.42	0.14	0.18	0.18	0.90	.14	.67
b_3	0.17*	.27	0.07	0.07	0.03	0.29	.05	.47
b_4	0.04	.06	0.06	0.07	-0.10	0.17	15	.26
b_5	0.47*	.27	0.16	0.22	0.04	0.88	.02	.52
b_6	0.18*	.23	0.08	0.08	0.03	0.33	.04	.43
c_1	0.56	.07	0.53	0.57	-0.51	1.73	06	.20
c_2	2.29	.11	1.42	1.61	-0.70	5.67	03	.25
\mathbf{c}_3	-0.05	01	0.69	0.72	-1.40	1.41	14	.15
c_4	-0.10	09	0.08	0.08	-0.27	0.06	26	.06
\mathbf{c}_5	-0.07	03	0.21	0.20	-0.48	0.32	17	.13
c_6	-0.03	03	0.10	0.10	-0.24	0.15	19	.13
Indirect 1	path estimates	(mediation)						
a_1b_1	0.68	.08		0.49	-0.14	1.80	02	.21
a_1b_2	1.27^{\dagger}	.06		0.98	-0.12	3.94	01	.19
a_1b_3	0.38^{\dagger}	.04		0.31	-0.04	1.25	00	.13
a_2b_4	0.08	.01		0.17	-0.17	0.59	02	.07
a_2b_5	1.02*	.05		0.79	0.03	3.35	.00	.16
a_2b_6	0.39*	.04		0.27	0.03	1.20	.00	.12
a_3b_1	0.02	.02		0.07	-0.11	0.15	11	.14
a_3b_2	0.03	.01		0.13	-0.22	0.30	09	.12
a_3b_3	0.01	.01		0.04	-0.06	0.10	05	.08
a_4b_4	0.00	.00		0.01	-0.03	0.04	03	.03
a_4b_5	0.00	.00		0.10	-0.21	0.21	09	.08
a_4b_6	0.00	.00		0.04	-0.07	0.08	06	.06
$a_{1,2}b_{1,4}$	0.76^{\dagger}	.09		0.50	-0.13	1.84	02	.21
$a_{1,2}b_{2,5}$	2.30^{\dagger}	.11		1.32	-0.16	.5.05	01	.22
$a_{1,2}b_{3,6}$	0.77^{\dagger}	.08		0.44	-0.02	1.73	00	.17
$a_{1,2}b_{1,4}$	0.02	.02		0.07	-0.12	0.16	12	.15
$a_{1,2}b_{2,5}$	0.04	.01		0.20	-0.36	0.44	14	.16
$a_{1,2}b_{3,6}$	0.01	.01		0.07	-0.12	0.14	10	.11

Note. N = 132. ML = Maximum likelihood estimation. Continuous predictor variables centred. Probability values determined on bootstrapped CIs (10000 resamples). $^{\dagger}p < .10. *p < .05. **p < .01. ***p < .001$.

Table 4

Additional path estimates for the moderated mediation model (Figure 2)

Path	Estimates		SI	E of b	Bootstrap	95% CIs <i>b</i>	Bootstrap 95% CIs β		
	ь	β	ML	Bootstrap	Lower	Upper	Lower	Upper	
Direct (mo	oderation) pa	th estimates							
a_5	0.70^{\dagger}	.18	0.36	0.40	-0.11 1.45		02	.38	
a_6	0.76*	.25	0.29	0.35	0.05	1.41	.02	.47	
Indirect pa	ath estimates	(mediation)	at three leve	els of disgust p	ropensity				
Moderate	disgust prop	ensity (M)							
a_1b_1	0.53	.06		0.47	-0.27	1.63	03	.18	
a_1b_2	1.00	.05		0.93	-0.33	3.56	02	.17	
a_1b_3	0.30	.03		0.29	-0.10	1.12	01	.12	
a_2b_4	0.06	.01		0.14	-0.12	0.51	01	.06	
a_2b_5	0.77^{\dagger}	.04		0.69	-0.07	2.86	00	.14	
a_2b_6	0.29^{\dagger}	.03		0.24	-0.02	1.00	00	.10	
$a_{1,2}b_{1,4}$	0.59	.07		0.48	-0.27	1.61	03	.19	
$a_{1,2}b_{2,5}$	1.77	.08		1.25	-0.61	4.30	03	.19	
$a_{1,2}b_{3,6}$	0.59	.06		0.42	-0.18	1.47	02	.15	
High disg	ust propensi	ty (+ 1 <i>SD</i>)							
a_1b_1	1.34*	.16		0.69	0.14	2.91	.02	.33	
a_1b_2	2.51*	.12		1.50	0.34	6.53	.02	.31	
a_1b_3	0.75*	.08		0.47	0.08	2.04	.01	.21	
a_2b_4	0.17	.02		0.34	-0.41	1.03	05	.13	
a_2b_5	2.15*	.11		1.38	0.25	6.00	.01	.29	
a_2b_6	0.82*	.09		0.49	0.13	2.19	.01	.23	
$a_{1,2}b_{1,4}$	1.51*	.18		0.73	0.22	3.15	.02	.35	
$a_{1,2}b_{2,5}$	4.66*	.21		2.03	0.96	8.94	.04	.40	
$a_{1,2}b_{3,6}$	1.57*	.16		0.68	0.35	3.07	.03	.31	
Low disg	ust propensit	ty (- 1 <i>SD</i>)							
a_1b_1	-0.27	03		0.65	-1.56	1.04	19	.13	
a_1b_2	-0.51	03		1.27	-3.48	0.55	17	.08	
a_1b_3	-0.15	02		0.39	-1.07	1.73	11	.06	
a_2b_4	-0.05	01		0.15	-0.63	0.12	08	.01	
a_2b_5	-0.61	03		0.88	-2.99	0.72	15	.04	
a_2b_6	-0.23	02		0.33	-1.15	0.25	12	.03	
$a_{1,2}b_{1,4}$	-0.32	04		0.70	-1.74	1.04	20	.12	
$a_{1,2}b_{2,5}$	-1.13	05		1.88	-5.11	2.42	23	.11	
$a_{1,2}b_{3,6}$	-0.39	04		0.62	-1.65	0.82	17	.08	

Note. N = 132. ML = Maximum likelihood estimation. Continuous predictor variables centred. Probability values determined on bootstrapped CIs (10000 resamples). $^{\dagger}p < .10. *p < .05. **p < .01. ***p < .001$.