Journal: Biological Psychology Running head: Alexithymia and interoception: problems with the heartbeat counting task Word count: 6169 words Submission date: 7th February 2018 (Revision date: 16th May 2018)

Is alexithymia characterised by impaired interoception? Further evidence, the importance of control variables, and the problems with the Heartbeat Counting Task

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Abstract

Interoception, the perception of one's internal state, is commonly quantified using the heartbeat counting task (HCT) – which is thought to be a measure of cardiac interoceptive sensitivity (accuracy). Interoceptive sensitivity has been associated with a number of clinical traits and aspects of higher order cognition, including emotion processing and decision-making. It has been proposed that alexithymia (difficulties identifying and describing one's own emotions) is associated with impaired interoceptive sensitivity, but new research questions this association. Problematically, much evidence attesting to the absence of this association has been conducted using the HCT, a measure affected by various physiological and psychological factors. Here, we present novel data (N=287) examining the relationship between alexithymia and HCT performance, controlling for a number of potential confounds. Inclusion of these control measures reveals the predicted negative relationship between alexithymia and HCT performance. Results are discussed with regard to difficulties quantifying interoceptive sensitivity using the HCT.

Keywords: alexithymia; heartbeat counting; interoception; interoceptive accuracy; interoceptive sensitivity

1. Introduction

Interoception is generally defined as the ability to perceive one's internal state. Such a seemingly simple definition hides a great deal of uncertainty as to what constitutes an internal signal. For example, some consider proprioception, or perception of external signals that activate interoceptive pathways such as 'affective touch' (e.g. slow stroking of the forearm), to be interoceptive signals, while others do not (Khalsa & Lapidus, 2016). Further, the degree to which interoceptive signals need to be consciously perceived and/or recognised in order for a process to be described as interoceptive has been debated (see Murphy, Brewer, Catmur, & Bird, 2017, for discussion). The wider nature of interoception is also under debate; Garfinkel, Seth, Barrett, Suzuki and Critchley (2015) have proposed extending the notion of interoception by separating it into a tripartite model, whereby three facets of interoceptive 'ability' exist. Under this model, interoceptive sensitivity refers to one's objective accuracy in perceiving interoceptive states. Interoceptive sensitivity is assessed by comparing the degree to which one's perception of one's internal state aligns with objective measures of that internal state. Interoceptive sensibility, on the other hand, describes subjective beliefs about one's own interoceptive states, including the extent to which one perceives oneself to be a) aware of internal signals, and b) accurate at detecting these internal signals. Finally, interoceptive awareness refers to the degree to which one can accurately assess one's own interoceptive sensitivity (a metacognitive ability). However, other models of interoceptive ability have been proposed, with new approaches advocating a distinction between beliefs (self-report) and objective data concerning a) the ability to perceive the internal state of one's body, and b) the propensity to become aware of, and separately to utilise, interoceptive information (Murphy, Catmur, & Bird, 2017).

Several models of higher-order cognition assign a role to interoception, in areas as diverse as emotion processing, learning and decision-making, and the sense of self (e.g., Critchley &

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Nagai, 2012; Critchley & Harrison, 2013; Dunn et al., 2010; Füstös, Gramann, Herbert, & Pollatos, 2013; Quattrocki & Friston, 2014; Seth, 2013). The study of interoception has also been extended into the clinical domain, due to the fact that atypical interoception is thought to characterise a number of physical and psychiatric conditions such as eating disorders, anxiety, depression and Autism Spectrum Disorder (see Barrett & Simmons, 2015; Khalsa & Lapidus, 2016; Murphy, Brewer, et al., 2017). One of the most comprehensive clinically-relevant interoceptive theories was that advanced by Quattrocki and Friston (2014), who suggested an interoceptive deficit was responsible for the symptoms of Autism Spectrum Disorder (henceforth 'autism'). However, the literature on interoception in autism is mixed; while Garfinkel, Tiley, et al. (2016) found that adults with autism demonstrated worse performance on the Heartbeat Counting Task (HCT; Dale & Anderson, 1978; Schandry, 1981), a commonly-used measure in which participants are required to count their heartbeats over a specified interval and their count is compared to an objective measure, they were unimpaired on another measure of cardiac interoceptive sensitivity (the heartbeat discrimination task, in which participants are required to judge whether auditory or visual signals are synchronous with their heartbeat). In addition, Schauder, Mash, Bryant and Cascio (2015) examined HCT performance in autistic¹ children and found them to be unimpaired. In fact, autistic children performed better on the task than typical children over longer durations. Noel, Lytle, Cascio and Wallace, (2018) also found that a small sample of adults with autism performed at a level similar to typical individuals on the HCT.

Brewer, Happé, Cook, & Bird (2015) have argued that the pattern of deficits predicted by Quattrocki and Friston's interoceptive model does not characterise autism, but that some of the deficits instead characterise alexithymia (a sub-clinical condition in which individuals are

¹ The use of the term 'autistic' is endorsed by many individuals with ASD (see Kenny et al., 2016). We therefore use this term as well as language preferred by clinical professionals (e.g., 'individuals with autism') to respect the wishes of autistic individuals and report the study in line with scientific parlance.

poor at identifying and describing their emotions, and have an externally-oriented thinking style (Bagby, Parker, & Taylor, 1994). The possibility of a link between alexithymia and interoceptive sensitivity was supported by the results of an initial study, which found that increased levels of alexithymia were associated with worse performance on the HCT (Herbert, Herbert, & Pollatos, 2011). As approximately 50-60% of individuals with autism also have alexithymia (e.g., Berthoz & Hill, 2005), it is possible that sampling variance with respect to alexithymia within the autistic population explains the inconsistent findings concerning autism and interoception. It may be the case that when samples of autistic individuals are largely comprised of alexithymic individuals, group level deficits are observed, but when the autism sample has a smaller proportion of alexithymic individuals, the autistic group perform as well as a group of typical individuals. This hypothesis was supported in a study in which the impact of autistic and alexithymic traits on the HCT were contrasted. Across two experiments, alexithymia, rather than autism, predicted lower sensitivity to cardiac signals as measured using the HCT (Shah, Hall, Catmur, & Bird, 2016). While the relationship between alexithymia and HCT performance replicated across both experiments, samples were relatively small (N < 50), likely providing an imprecise measure of the true effect size. Indeed, the magnitude of the correlation between alexithymia and performance on the HCT varied considerably across experiments (-.36 and -.64). It is notable, therefore, that the one previous study to examine the relationship between alexithymia and performance on the HCT in a larger (N = 155) non-clinical sample found a correlation of -.37 (Herbert et al., 2011). More recent indirect studies also support this association in a typical population; Bornemann and Singer (2017) demonstrated that 9 months of meditative training had correlated effects on levels of alexithymia and interoceptive sensitivity, such that the reduction in alexithymia following the meditative training was associated with improvements in interoceptive sensitivity measured using the HCT.

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Evidence therefore supports the hypothesis that alexithymia, rather than autism, is associated with poor performance on the HCT and, assuming the HCT is an index of interoceptive sensitivity, that alexithymia is associated with impaired interoceptive sensitivity. However, since these initial studies were published, a small number of papers and conference proceedings have reported a failure to replicate the association between performance on the HCT and alexithymia (e.g. Borhani, Làdavas, Fotopoulou, & Haggard, 2017; Zamariola, Vlemincx, & Luminet, 2018; Palser, Pellicano, Fotopoulou & Kilner, 2017). It is therefore crucial to examine methodological factors that may explain these inconsistent findings, in order to guide both on-going and future studies.

This paper therefore presents novel data on the link between alexithymia and performance on the HCT from a larger sample (N = 287) of adult participants, and scrutinises factors that may impact results across studies. In particular, we examine how the inclusion of control variables influences the effect size (and therefore significance) of the relationship between alexithymia and performance on the HCT. Considering how inclusion of appropriate control variables affects the observed results adds to pre-existing concerns regarding the suitability and validity of the HCT as a measure of interoceptive sensitivity. Despite its popularity, the task has received a significant degree of criticism: evidence suggests that the task may be completed using exteroceptive information alone (Khalsa, Rudrauf, Sandesara, Olshansky, & Tranel, 2009), and that the task may index prior knowledge of resting heart rate rather than interoceptive sensitivity (Brener & Ring, 2016). Previous studies have also detailed a range of psychological and physiological factors which impact on performance on cardiac based measures of interoceptive sensitivity and may determine the degree of performance explained by interoceptive and exteroceptive factors (e.g., blood pressure; see O'Brien, Reid, & Jones, 1998), heart rate variability and resting heart rate (Knapp-Kline & Kline, 2005), body mass index (Rouse, Jones, & Jones, 1988), and beliefs and knowledge of resting heart rate (Brener

& Ring, 2016; Ring, Brener, Knapp, & Mailloux, 2015; Ring & Brener, 1996; Windmann, Schonecke, Fröhlig, & Maldener, 1999). Further criticism centres on the lack of a suitable control task, and inconsistencies in the implementation of the task across studies. All of these factors are discussed below, and the current results contribute to the debate around the validity and reliability of the HCT as a measure of interoceptive sensitivity.

2. Methods

2.1 Participants

299 volunteers took part in this study in exchange for a small honorarium. Participants were recruited via local advertisements and databases of individuals who had expressed an interest in taking part in psychological research. Ethical approval was granted by the local ethics committee. In line with the declaration of Helsinki, all participants gave informed consent and were fully debriefed upon task completion. 12 participants were removed for extreme scores on control variables (see Analysis Strategy) resulting in 287 valid cases (86 Males, $M_{age} = 38.07$ years, $SD_{age} = 21.09$ years, Range = 18-90 years).

2.2 Measures

Data presented here were collected from participants across a period of two years and combined for the purpose of the present analysis. Some participants took part in more than one study using the HCT, therefore duplicate values for participants were removed prior to analyses. During this time period the measures used by our research group have changed. Therefore, for three factors (depression, anxiety, heart rate estimates) the measures utilised differ across participants (detailed below). The method used for the collection of all other variables was the same across all participants.

2.2.1 Alexithymia

Alexithymia was quantified using the Toronto Alexithymia Scale (TAS-20; (Bagby, et al., 1994). This measure includes 20 items, rated on a scale from 1-5, yielding scores between 20 and 100, with higher scores representative of more severe alexithymic traits. In this sample, total scores ranged from 20-82 (M = 45.26, SD = 13.00) with 39 individuals in the sample meeting cut off for alexithymia (≥ 61).

2.2.2 The Heartbeat Counting Task

As is typical during the HCT, participants were asked to silently count their heartbeats over a series of intervals whilst their heartbeat was objectively recorded using a pulse oximeter. Participants were explicitly instructed not to count seconds or guess; if they could not feel their heartbeat at all, they were asked to give a response of zero. Four durations were examined (25, 35, 45, 100 seconds) with half of participants completing longer durations (28, 38, 48, 103 seconds). As a control, participants were also asked to complete a time estimation task, in which they were asked to count seconds instead of heartbeats. The durations utilised (e.g., 25 vs. 28) were counterbalanced across the time and heartbeat tasks. Across both tasks, the order of durations was counterbalanced across participants, and half of the participants completed the timing task first, while half completed the HCT first. HCT and time estimation accuracy were estimated on a scale from 0 - 400 using the following equation, where higher scores indicate better cardiac/time estimation accuracy: SUM(1 - (|Objective measure participant's estimate/Objective measure)) x 100. In individuals for whom counterbalancing information was available (N = 271) no order effect (HCT vs time estimation first) was observed for performance on the HCT (t(269) = -.559, p > .50). Levene's test indicated that the assumption of equal variances was violated for the time estimation task data (F = 11.845, p=.001), therefore a robust method was utilised to analyse these data (see Field & Wilcox,

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2017). The Yuen (1974) modified t-test revealed no significant difference in time estimation performance as a function of task order (Mdiff = -10.77 [-28.60, 7.06], Yt = -1.19, p > .20).

2.2.3 Additional control measures

As performance on the HCT has been found to be influenced by various physiological and psychological factors (e.g., Brener & Ring, 2016; Khalsa, Rudrauf, Sandesara, et al., 2009; Knapp-Kline & Kline, 2005; O'Brien et al., 1998; Pollatos, Traut-Mattausch, & Schandry, 2009; Rouse et al., 1988; Wittmann, 2013) a number of control measures were employed. These were available for the majority of participants (see Table 1). Body mass index (BMI), systolic blood pressure and knowledge of the 'typical' resting heartrate were collected post-study for all participants. Depression and anxiety were assessed at the same time as alexithymia with these questionnaires completed in a randomised order.

2.2.3.1 Body Mass Index

BMI was calculated using the following equation: $mass(kg)/(height(m))^2$.

2.2.3.2 Systolic blood pressure

Blood pressure was taken using an electronic upper arm monitor (Omron M2) whilst participants were seated. High scores indicate higher systolic blood pressure.

2.2.3.3 Resting heart rate & heart rate variability

Average resting heart rate was taken as a measure of resting heart rate. For some participants all intervals were included, whereas for others the last 60 seconds of the longest duration was utilised. The root mean square of successive differences was used as a measure of heart rate variability (HRV). Higher scores indicate higher resting heart rate or increased heart rate variability.

2.2.3.4 Knowledge of average resting heart rate

After the heartbeat counting task participants were asked to estimate the average person's resting heart rate "how many times do you think the average person's heart beats in

60 seconds when they are at rest?" (see Murphy, Geary, Millgate, Catmur, & Bird, 2017; Murphy et al., 2018). The absolute difference between the participant's estimate and average resting heart rate (reported in large studies of human physiology; 72.26; Agelink et al., 2001; Ramaekers, Ector, Aubert, Rubens, & Van de Werf, 1998) was taken as a measure of accuracy. This was favoured over asking participants to estimate their own heart rate to avoid effects of estimation on the HCT and vice versa. High scores on this variable indicate greater deviation between the participant's estimate and average resting heart rate, and therefore greater inaccuracy.

2.2.3.5 Depression

Depressive traits were measured using either the Beck Depression Inventory (BDI; Beck, Ward, Mendelson, Mock & Erbaugh, 1961; Beck, Steer, & Brown, 1996) or the depression subscale from the Depression, Anxiety and Stress Scale (Lovibond & Lovibond, 1995; Lovibond, 1993). To combine these scores into one variable, scores within the sample reported here were Z-scored and these Z scores were then combined into one variable indexing depressive traits. As such, high scores indicate greater depressive traits.

2.2.3.6 Anxiety

Anxiety was measured using either the anxiety subscale from the Depression, Anxiety and Stress Scale (Lovibond & Lovibond, 1995; Lovibond, 1993) or the trait anxiety subscale from the State-Trait Anxiety Inventory (Spielberger, Gorsuch, & Lushene, 1970). To combine these scores into one variable, scores within the sample reported here were Z-scored and these Z scores were then combined into one variable indexing anxious traits. As such, high scores indicate greater anxiety.

2.3 Analysis Strategy

Initially, the zero-order correlation between alexithymia and score on the HCT is reported (ignoring performance on the time estimation task and without accounting for any physiological or psychological control variables). Zero-order correlations between all variables are also reported, as well as partial correlations between alexithymia and HCT performance controlling for each control variable separately. We then report the results of a series of multiple regressions in which control measures are successively added (see Supplementary Results). These analyses are included for illustrative purposes only and demonstrate how the results change with each added control variable. Whilst directional predictions can be made for all variables, results of two-tailed statistical tests are reported for all analyses. Univariate outliers more than three times the interquartile range were removed. This resulted in exclusion of one outlier on the basis of BMI, four on the basis of HRV, two on their knowledge of the average heart rate, four for extreme depression scores, and one on the basis of systolic blood pressure. The inclusion of these 12 individuals, however, did not alter the pattern observed in the final model reported in Table 2. For each regression, we report the predictor values and number of participants, and the standardised Beta, t value, significance, and partial correlation coefficient for alexithymia in each regression model (see Supplementary Results; Table S1), and the same values for each predictor variable in the full regression model. It is the full, final, regression model that tests the association between alexithymia and performance on the HCT after controlling for all relevant variables (Table 2).

3. Results and Discussion

3.1 Results

The zero-order correlation between alexithymia and HCT performance, ignoring performance on the control task and failing to account for any control variable, was not

significant (r(285) = -.079, p=.182). Table 1 presents the descriptive statistics for all measured variables, the zero-order correlations between all variables, and the partial correlation between HCT performance and alexithymia controlling for each variable separately.

The models, and relevant values for alexithymia, for a series of multiple regression models in which alexithymia and an increasing number of control variables are used to predict performance on the HCT are reported in the Supplementary Results (with each predictor entered in the order that maintained maximum statistical power; Table S1). It can be seen that inclusion of the various control variables changes the observed effect size from r(partial) = -.079 to -.193. Importantly, however, when the full range of control measures was included (Table 2), alexithymia was a significant predictor of performance on the HCT. The same pattern was observed when missing values were replaced to retain power (see Supplementary Results; Table S3). Moreover, whilst the residuals from these models were normally distributed (see Supplementary Results: Figure S1), to confirm the final model reported in Table 2 the data were analysed using a robust regression procedure (Field & Wilcox, 2017) implemented in Matlab (2014) with the default weighting function employed. This analysis confirmed the same pattern of results; alexithymia was a significant predictor of poor performance (p = .021), and accurate knowledge of resting heartrate and male gender were both predictors of good performance (p=.048 and p=.003 respectively).

These results highlight a potential reason for the observed variance in the effect size relating HCT performance to alexithymia across studies: failure to appropriately control for the various non-interoceptive factors that influence performance on the HCT will influence the observed effect size. We now turn to other potential reasons why one may see variance in the observed effect size across studies.

3.2 The Heartbeat Counting Task is a Poor Measure of Interoceptive Sensitivity

The HCT is commonly used as a measure of interoceptive sensitivity as it is very quick, cheap, and easy to administer, but it is generally recognised as having substantial problems. Approximately 40% of typical individuals report no conscious awareness of their heartbeat at all (Khalsa, Rudrauf, Sandesara et al., 2009), making this task unsuitable for examining interoception at lower ranges of ability. Perhaps most problematic, however, is that heartbeat may be perceived via (exteroceptive) touch receptors due to the vibration of the chest wall (Khalsa, Rudrauf, Sandesara et al., 2009). The extent to which the heartbeat may be perceived exteroceptively² depends on factors such as the percentage of body fat (Rouse et al., 1988), systolic blood pressure (O'Brien et al., 1998), resting heart rate, and heart rate variability (Knapp-Kline & Kline, 2005). This is clearly of concern when it comes to comparisons across studies; even if the relationship between interoception and alexithymia is perfectly fixed and unchanging, one may observe large variation in the size of the relationship between performance on the HCT and alexithymia (or any other variable to which HCT performance is related) depending on the particular physical characteristics of the sample tested, and the ratio of interoceptive to exteroceptive information participants were using to perform the task. This is further complicated by the fact that some of these factors may themselves be associated with alexithymia (or autism) (e.g., rates of alexithymia are higher in obese individuals; Pinna et al., 2011). This highlights the need to control for these factors when using the HCT, as failure to do so renders the results extremely hard to interpret. Indeed, in the current data depending on which physical parameters (e.g., BMI, HRV, Systolic blood pressure, resting heart rate) are controlled for, the observed r value for the correlation between the HCT and alexithymia varies between -.079 and -.167, and in this

² Whilst these factors are likely to impact upon the perception of cardiac signals via exteroceptive mechanisms, it is important to note that in the absence of data it remains a possibility that these physiological factors also contribute towards individual differences in interoceptive perception.

sample controlling for systolic blood pressure and heart rate variability alone resulted in a significant relationship between alexithymia and HCT performance (Table 1).Whilst the inclusion of all control variables only had a modest influence on the effect size of the relationship between alexithymia and HCT performance, the importance of controlling for these factors may be greater in clinical populations characterised by ill-health (and thus, higher BMI, Systolic blood pressure and greater HRV and resting heart rate; e.g., Hert et al., 2011), or at certain stages of development (e.g., knowledge of resting heartrate and beliefs may differ substantially within children or adolescents).

Procedural differences in the way the task is administered can also contribute to the discrepant findings across studies. A time estimation task is often used alongside the HCT to control for nonspecific factors that may influence HCT performance such as motivation, fatigue, etc. (e.g., Ainley, Brass, & Tsakiris, 2014; Murphy, Geary, et al., 2017; Shah et al., 2016), and inclusion of this task may be especially crucial for autistic or alexithymic individuals, who may feel anxious or distracted during any experimental task. Failure to include a control task therefore means that any studies reporting the presence or absence of a correlation with the HCT are extremely difficult to interpret - any of these nonspecific factors may artificially inflate the relationship between HCT performance and alexithymia, or mask a real association. Indeed, in the present data, time estimation remained a significant predictor of performance on the HCT in the final model. It is worth noting, however, that although the time estimation task controls for many nonspecific factors, it does not control for differences in detection thresholds relating to decision parameters. For example, if those with autism or alexithymia require more sensory evidence (regardless of whether this is interoceptive or exteroceptive) than neurotypical individuals in order to decide an event has occurred, this factor would affect the HCT but not the time estimation task. To control for factors such as

these, a control task in which exteroceptive stimuli must be counted (matched for detectability with heartbeats at the population level) may be preferable.

Several other factors have been shown to significantly impact upon the results obtained using the HCT. For example, one factor relates to the effect of knowledge of one's own, or the average person's, heart rate. Indeed, a body of evidence demonstrates that manipulating participants' beliefs about one's own resting heart rate alters heartbeat counting estimates in the HCT (Ring et al., 2015; Ring & Brener, 1996; Windmann et al., 1999). Likewise, accurate knowledge of average heart rate correlates with improved performance on the HCT (Murphy, Geary, et al., 2017; Murphy et al., 2018), as replicated in the current data, and it may do so via at least two routes. The first may be considered interoceptive (depending on one's definition of interoception); knowledge of the frequency of one's heartbeat may allow the heartbeat signal to be distinguished from other interoceptive signals and therefore accurately recognised. The second is not interoceptive and may be evidenced by an interaction with an important procedural factor; whether participants are encouraged to guess if they cannot feel their heartbeat. The particular instructions given to participants in the HCT are rarely reported but an informal survey suggests that participants are often instructed to guess (or 'estimate') if they cannot feel a heartbeat as per early descriptions of the task (Schandry, 1981; see Brener & Ring, 2016). This instruction is not universal however; the same informal survey also found that participants were sometimes (less frequently) instructed to report zero heartbeats if they felt no heartbeats. If participants are instructed to guess (or if they do so regardless of the instruction not to guess) then a sensible strategy is to estimate the duration of the interval over which one is required to count one's heartbeats and use the knowledge of one's resting heart rate (or the average resting heart rate) to arrive at an estimate of the number of heartbeats. It is therefore crucial to control for the accuracy of participants' knowledge concerning their own or the typical resting heart rate, and to (at least)

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use the time estimation task as a control. It is also important to report the instructions given to participants with respect to guessing. Arguably, it is more appropriate to instruct participants to report zero heartbeats if they can feel zero heartbeats than asking them to make an arbitrary guess. Importantly, these factors may differentially impact upon typical and clinical groups; using a modified version of the HCT, Khalsa et al. (2017) demonstrated that instructions relating to guessing had a significant impact on performance, and that this impact was significantly different in a clinical group (those with Substance Use Disorder) and a typical control group.

Importantly, even if all of the limitations listed above are taken into account and appropriately controlled for, it is still unclear whether the HCT is a measure of interoceptive sensitivity. Due to the lack of work controlling for all potential confounds we cannot currently be sure whether, when administered properly, it is an adequate test of interoceptive sensitivity. It is therefore particularly problematic to relate individual differences in HCT performance to psychological variables or performance on cognitive tasks. Importantly, these difficulties are problematic not only for relating HCT performance to alexithymia, but apply to any individual difference measure. For example, the present data demonstrate that the documented decrease in interoceptive sensitivity with advancing age (Khalsa, Rudrauf, & Tranel, 2009; Murphy, Geary, et al., 2017) is absent when a number of control variables are taken into account. Likewise, well-evidenced relationships between anxiety and interoceptive sensitivity, and depression and interoceptive sensitivity (e.g., Pollatos et al., 2009; see section 3.3) were not found in the present sample. Thus, whilst the following sections focus on the relationship between alexithymia and HCT performance, it is important to acknowledge that the limitations described above are applicable to all research using the HCT, and so are also likely to impact the debate concerning interoception in autism.

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3.3 The Relationship Between Anxiety, Depression, Alexithymia and Autism

As noted, it has been argued that poor interoception may give rise to the characteristic features of autism (Quattrocki & Friston, 2014), but we have proposed that it is alexithymia, not autism, that is characterised by impaired interoception (Brewer et al., 2015). It is well known that individuals with high levels of autistic or alexithymic traits have an increased likelihood of suffering from increased levels of depression and anxiety (e.g., Ghaziuddin, Ghaziuddin, & Greden, 2002; Gillott & Standen, 2007; Hendryx, Haviland, & Shaw, 1991; Honkalampi, Hintikka, Tanskanen, Lehtonen, & Viinamäki, 2000). Interestingly, symptoms of both anxiety and depression have been associated with atypical interoception (almost exclusively measured using the HCT), whereby increased symptoms of anxiety are associated with increased interoceptive sensitivity (Ehlers & Breuer, 1996; Ehlers, Mayou, Sprigings, & Birkhead, 2000; Pineles & Mineka, 2005; Roth et al., 1992; Wald & Taylor, 2005; White, Brown, Somers, & Barlow, 2006; Willem Van der Does, Antony, Ehlers, & Barsky, 2000; Zoellner & Craske, 1999), and increased symptoms of depression are associated with reduced interoceptive sensitivity (Pollatos et al., 2009); note that it is also possible that the relationship between depression symptom severity and interoception is non-linear (Dunn, Dalgleish, Ogilvie, & Lawrence, 2007).

The implications of this relationship for measuring the interoceptive ability (interoceptive sensitivity, sensibility or awareness) of individuals with alexithymia or autism are clear; without accounting for depression and anxiety (either by matching alexithymic/autistic and control groups on levels of depression and anxiety, or through controlling for anxiety and depression statistically³) one simply cannot measure the relationship between alexithymia or autism and interoceptive ability. If the group of interest

³Note that one must be aware of issues with collinearity when relying on statistical control of effects of depression and anxiety (or any other highly correlated variables). Matched groups are therefore the preferred method.

is more depressed than the control group, this may artificially inflate any deleterious effect of the condition of interest on interoception, whereas the opposite will be true if the group of interest is more anxious than the control group. It is therefore very difficult to conclude anything about the relationship between autism or alexithymia and interoception from studies that have not controlled for these factors.

3.4 The relationship between Alexithymia and the HCT

Above, we suggested that the relationship between alexithymia and interoceptive sensitivity may be incredibly reliable in actuality, but very unreliable when interoceptive sensitivity is measured using the HCT due to the many problems associated with using the HCT as a measure of interoceptive sensitivity. Of course, it is also true that measurement of alexithymia may be unreliable or invalid. Likewise, it remains a possibility that there may be multiple routes to alexithymia and that sometimes alexithymia may be observed in the absence of interoceptive impairment. Whilst these issues are beyond the scope of discussion of the current results, we have commented on these issues in the Supplementary Discussion.

Beyond problems with the measurement of alexithymia, or multiple types of alexithymia (some being associated with interoceptive impairment and some not), a further explanation for the variability in the strength of the association between alexithymia and performance on the HCT could be that there is no association, and that findings of an association are false positives. If true, it is still unclear what should be concluded. As detailed above, even if all appropriate physiological and psychological factors are controlled for, and an adequate control task employed, it is still not clear that the HCT is actually a measure of interoceptive sensitivity. This is important as relationships between alexithymia and interoceptive sensitivity in the domains of taste (Murphy, Catmur, et al., 2017), muscular effort (Murphy, Catmur, et al., 2017; Van Der Cruijsen, Murphy, Crone & Bird, in prep.), temperature (Borhani et al., 2017) and physiological arousal (Gaigg et al., 2016). Furthermore, several studies have reported a relationship between alexithymia and selfreported interoceptive sensibility (Betka et al., 2018; Brewer, Cook, & Bird, 2016; Longarzo et al., 2015; Zamariola et al., 2018) or the objectively-measured propensity to utilise interoceptive information in the respiratory domain (Murphy, Catmur, et al., 2017; Zhang, Murphy, Bird & Lau, in prep. (adolescent females only)). If these relationships are not false positives, and yet the previously observed (and currently observed) relationship between HCT performance and alexithymia is, then the logical conclusion is that either the HCT is not a (good) measure of interoception, or that interoception fractionates, such that performance in some interoceptive domains (and dimensions; e.g., self-report vs objective measures) is predicted by alexithymia, and some not. The possible fractionation of interoceptive ability is currently a matter of debate; whilst the perception of cardiac signals has received much research attention, with the HCT commonly employed (Dale & Anderson, 1978; Schandry, 1981), in recent years a number of novel measures of interoception have been developed to assess interoceptive ability across multiple domains (e.g., respiratory). In part, these research efforts have been driven by concerns over the validity of cardiac measures of interoception (Khalsa et al., 2009), and by the need to test the assumption of a unitary interoceptive ability. Whilst some studies support a unitary view, for example moderate correlations have been reported between measures of gastric and cardiac interoception (Herbert, Muth, Pollatos, & Herbert, 2012; Whitehead & Drescher, 1980), others find no relationship across domains (e.g., respiratory and cardiac; Ehlers & Breuer, 1992; Garfinkel, Manassei, et al., 2016; Pollatos, Herbert, Mai, & Kammer, 2016; Steptoe & Vögele, 1992). Likewise, the HCT is often uncorrelated with self-report measures of interoception (Garfinkel et al., 2015). It is worth noting, however, that even if interoceptive ability does fractionate according to the

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signal to be perceived and along the dimension measured (e.g., self-report (sensibility) vs. objective accuracy (sensitivity)), it is still the case that performance on the HCT only correlates modestly with the second-most commonly used test of objective interoceptive sensitivity, which also tests the ability to perceive cardiac signals (the Heartbeat Discrimination Task; Whitehead, Drescher, Heiman, & Blackwell, 1977) with reports of small (e.g., Garfinkel et al., 2015) or absent (e.g., Ring & Brener, 2018) relationships between performance on these two tasks. If the two 'gold-standard' measures of cardiac interoceptive sensitivity correlate only modestly (if at all), then it becomes problematic to conclude anything about cardiac interoceptive sensitivity from either test. Indeed, recent work assessing the test-retest reliability of the HCT suggests that scores at one time point correlate only moderately (r = .6) with scores just two months later i.e. only 36% of the variance in test scores at Time 2 are predicted by Time 1⁴ (Ferentzi, Drew, Tihanyi, & Köteles, 2018). Assuming interoceptive sensitivity does not actually fluctuate over 2 months to this degree, then one may find very different estimates of the true correlation between alexithymia (or autism) and cardiac interoceptive ability due to the unreliability of the HCT.

3.5 Limitations

Despite the relevance of this evidence for our understanding of the relationship between alexithymia and interoception it is important to acknowledge certain limitations. First, although a number of control variables were present in the current dataset, no measure of autistic traits was included. Indeed, given relationships between alexithymia and autism (Berthoz & Hill, 2005), it is important that future research assesses the relationship between alexithymia, autism and interoceptive sensitivity using appropriate measures of interoception,

⁴Note that this study did not use a control task - therefore it is possible, in principle, that .6 is an over- (or under-) estimate of the reliability of the task.

coupled with the inclusion of necessary control variables. These considerations are important also for any future research examining the contribution of alexithymia to interoceptive difficulties in other conditions characterised by increased rates of alexithymia and poor interoceptive sensitivity e.g. eating disorders (Cochrane, Brewerton, Wilson, & Hodges, 1993; Klabunde, Acheson, Boutelle, Matthews, & Kaye, 2013; Pollatos et al., 2008) and shizophrenia (Ardizzi et al., 2016; Henry, Bailey, von Hippel, Rendell, & Lane, 2010). Second, although control variables were present for a number of individuals, data were not available for all participants. Despite these shortcomings, the present data underscore the need for future research to consider the appropriateness of the HCT as a measure of interoceptive sensitivity, and the use of appropriate control variables.

3.6 Conclusions

In conclusion, although the HCT is the most commonly used measure of interoceptive sensitivity, results from studies using this task are extremely difficult to interpret, even when those studies control for most, or all, of the factors identified here. Indeed, given the number of limitations of the task, it is unclear why it is used so frequently. Previous concerns over poor correlations with other measures of interoception (self-reported and objective, including other tests purporting to measure cardiac interoceptive sensitivity) and possible exteroceptive compensation strategies may be further exacerbated by the inconsistencies in administering the task. Currently, researchers do not provide consistent instructions, utilise the same control tasks, or account for the same set of variables known to have an impact on performance. The current results demonstrate that these factors can substantially influence the relationship between alexithymia and HCT performance, and presumably would also substantially influence the association between autism and HCT performance or indeed the relationship between any factor related to HCT performance. In particular, knowledge of the average heart

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rate may have a substantial effect on HCT performance. These results highlight that those studies that fail to use an adequate control task, or account for the full range of relevant psychological and physiological factors, may provide a very inaccurate estimate of relationships with HCT performance.

With respect to the relationship between alexithymia and performance on the HCT, previous findings of a significant relationship were replicated, such that those individuals with higher levels of alexithymia performed worse on the HCT task. It should be noted, however, that the size of the effect observed in these data was smaller than that observed in previous studies (\cong .20 versus \cong .35 in previous studies). To our knowledge, the studies failing to replicate this association have not accounted for the complete set of relevant control variables or included an appropriate control task. Given the current results, it is unlikely that these studies can provide an accurate assessment of the relationship between alexithymia and HCT performance.

In order to further our understanding of the relationship between interoception and alexithymia, it is essential to include a large number of individuals who score at least above the cut-off for alexithymia when investigating the relationship between alexithymia and interoception, and to control for co-occurring traits such as depression and anxiety. More broadly, for the study of interoception, it is clearly crucial to reduce the discrepancies across studies in the HCT methodology, as well as utilising alternative measures of interoception.

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Acknowledgments

JM was supported by a doctoral studentship from the Economic and Social Research

Council [1599941; ES/J500057/1]. GB was supported by the Baily Thomas Trust.

Data declaration

A subsample of the HCT data presented in this paper has previously been reported in the following papers, however the relationship between alexithymia and HCT performance controlling for all possible confounds has not been previously reported.

- Murphy, J., Millgate, E., Geary, H., Ichijo, E., Coll, M. P., Brewer, R., Catmur, C., & Bird, G. (2018). Knowledge of resting heart rate mediates the relationship between intelligence and the heartbeat counting task. *Biological Psychology*. 133, 1-3.
- Murphy, J., Geary, H., Millgate, E., Catmur, C., & Bird, G. (2017). Direct and indirect effects of age on interoception across the adult lifespan. *Psychonomic Bulletin and Review*, 1-10.
- Murphy, J., Millgate, E., Geary, H., Catmur, C., & Bird, G. (*Under review*). No effect of age on emotion recognition after accounting for cognitive factors and depression.

References

- Agelink, M. W., Malessa, R., Baumann, B., Majewski, T., Akila, F., Zeit, T., & Ziegler, D. (2001). Standardized tests of heart rate variability: normal ranges obtained from 309 healthy humans, and effects of age, gender, and heart rate. *Clinical Autonomic Research*, 11(2), 99–108. doi:10.1007/BF02322053
- Ainley, V., Brass, M., & Tsakiris, M. (2014). Heartfelt imitation: high interoceptive awareness is linked to greater automatic imitation. *Neuropsychologia*, 60, 21–28. doi:10.1016/j.neuropsychologia.2014.05.010
- Ardizzi, M., Ambrosecchia, M., Buratta, L., Ferri, F., Peciccia, M., Donnari, S., ... Gallese,
 V. (2016). Interoception and positive symptoms in schizophrenia. *Frontiers in Human Neuroscience*, 10, 379. doi:10.3389/fnhum.2016.00379
- Bagby, R. M., Parker, J. D., & Taylor, G. J. (1994). The twenty-item Toronto Alexithymia Scale--I. Item selection and cross-validation of the factor structure. *Journal of Psychosomatic Research*, 38(1), 23–32. doi:10.1016/0022-3999(94)90005-1
- Bagby, R. M., Taylor, G. J., & Parker, J. D. (1994). The Twenty-item Toronto Alexithymia Scale--II. Convergent, discriminant, and concurrent validity. *Journal of Psychosomatic Research*, 38(1), 33–40.
- Barrett, L. F., & Simmons, W. K. (2015). Interoceptive predictions in the brain. Nature Reviews. Neuroscience, 16(7), 419–429. doi:10.1038/nrn3950
- Beck, A. T., Steer, R. A., & Brown, G. K. (1996). Beck depression inventory-II. Autsin, TX.
- Beck, A.T., Ward, C, H., Mendelson, M., Mock, J., Erbaugh, J. (1961). "An inventory for measuring depression". Arch. Gen. Psychiatry. 4 (6): 561–71.
- Berthoz, S., & Hill, E. L. (2005). The validity of using self-reports to assess emotion regulation abilities in adults with autism spectrum disorder. *European Psychiatry*, 20(3), 291–298. doi:10.1016/j.eurpsy.2004.06.013

Psychiatry Research, 208(3), 257-264. doi:10.1016/j.psychres.2013.05.014

- Betka, S., Pfeifer, G., Garfinkel, S., Prins, H., Bond, R., Sequeira, H., ... Critchley, H. (2018).
 How Do Self-Assessment of Alexithymia and Sensitivity to Bodily Sensations Relate to Alcohol Consumption? *Alcoholism, Clinical and Experimental Research*, 42(1), 81–88. doi:10.1111/acer.13542
- Borhani, K., Làdavas, E., Fotopoulou, A., & Haggard, P. (2017). Lacking warmth": Alexithymia trait is related to warm-specific thermal somatosensory processing. *Biological Psychology*, 128, 132–140. doi:10.1016/j.biopsycho.2017.07.012
- Bornemann, B., & Singer, T. (2017). Taking time to feel our body: Steady increases in heartbeat perception accuracy and decreases in alexithymia over 9 months of contemplative mental training. *Psychophysiology*, 54(3), 469–482. doi:10.1111/psyp.12790
- Brener, J., & Ring, C. (2016). Towards a psychophysics of interoceptive processes: the measurement of heartbeat detection. *Phil. Trans. R. Soc. B*, 371(1708), 20160015.
- Brewer, R., Cook, R., & Bird, G. (2016). Alexithymia: a general deficit of interoception. *Royal Society Open Science*, *3*(10), 150664. doi:10.1098/rsos.150664

- Brewer, R., Happé, F., Cook, R., & Bird, G. (2015). Commentary on "Autism, oxytocin and interoception": Alexithymia, not Autism Spectrum Disorders, is the consequence of interoceptive failure. *Neuroscience and Biobehavioral Reviews*, 56, 348–353. doi:10.1016/j.neubiorev.2015.07.006
- Cochrane, C. E., Brewerton, T. D., Wilson, D. B., & Hodges, E. L. (1993). Alexithymia in the eating disorders. *The International Journal of Eating Disorders*, *14*(2), 219–222.
- Critchley, H D, & Nagai, Y. (2012). How Emotions Are Shaped by Bodily States. *Emotion Review*, 4(2), 163–168. doi:10.1177/1754073911430132
- Critchley, Hugo D, & Harrison, N. A. (2013). Visceral influences on brain and behavior. *Neuron*, 77(4), 624–638. doi:10.1016/j.neuron.2013.02.008
- Dale, A., & Anderson, D. (1978). Information variables in voluntary control and classical conditioning of heart rate: Field dependence and heart-rate perception. *Perceptual and Motor Skills*.
- Dunn, B. D., Dalgleish, T., Ogilvie, A. D., & Lawrence, A. D. (2007). Heartbeat perception in depression. *Behaviour Research and Therapy*, 45(8), 1921–1930. doi:10.1016/j.brat.2006.09.008
- Dunn, B. D., Galton, H. C., Morgan, R., Evans, D., Oliver, C., Meyer, M., ... Dalgleish, T. (2010). Listening to your heart. How interoception shapes emotion experience and intuitive decision making. *Psychological Science*, 21(12), 1835–1844. doi:10.1177/0956797610389191
- Ehlers, A., & Breuer, P. (1992). Increased cardiac awareness in panic disorder. *Journal of Abnormal Psychology*, *101*(3), 371–382. doi:10.1037/0021-843X.101.3.371
- Ehlers, A., & Breuer, P. (1996). How good are patients with panic disorder at perceiving their heartbeats?. *Biological psychology*, *42*(1-2), 165-182.
- Ehlers, A., Mayou, R. A., Sprigings, D. C., & Birkhead, J. (2000). Psychological and perceptual factors associated with arrhythmias and benign palpitations. *Psychosomatic Medicine*, 62(5), 693–702.
- Ferentzi, E., Drew, R., Tihanyi, B. T., & Köteles, F. (2018). Interoceptive accuracy and body awareness - Temporal and longitudinal associations in a non-clinical sample. *Physiology & Behavior*, 184, 100–107. doi:10.1016/j.physbeh.2017.11.015
- Field, A. P., & Wilcox, R. R. (2017). Robust statistical methods: a primer for clinical psychology and experimental psychopathology researchers. *Behaviour research and therapy*, 98, 19-38.
- Füstös, J., Gramann, K., Herbert, B. M., & Pollatos, O. (2013). On the embodiment of emotion regulation: interoceptive awareness facilitates reappraisal. *Social Cognitive* and Affective Neuroscience, 8(8), 911–917. doi:10.1093/scan/nss089
- Gaigg, S. B., Cornell, A. S., & Bird, G. (2016). The psychophysiological mechanisms of alexithymia in autism spectrum disorder. *Autism: The International Journal of Research and Practice*. doi:10.1177/1362361316667062
- Garfinkel, S. N., Manassei, M. F., Hamilton-Fletcher, G., In den Bosch, Y., Critchley, H. D.,
 & Engels, M. (2016). Interoceptive dimensions across cardiac and respiratory axes.
 Philosophical Transactions of the Royal Society of London. Series B, Biological Sciences, 371(1708). doi:10.1098/rstb.2016.0014

- Garfinkel, S. N., Seth, A. K., Barrett, A. B., Suzuki, K., & Critchley, H. D. (2015). Knowing your own heart: distinguishing interoceptive accuracy from interoceptive awareness. *Biological Psychology*, 104, 65–74. doi:10.1016/j.biopsycho.2014.11.004
- Garfinkel, S. N., Tiley, C., O'Keeffe, S., Harrison, N. A., Seth, A. K., & Critchley, H. D. (2016). Discrepancies between dimensions of interoception in autism: Implications for emotion and anxiety. *Biological Psychology*, *114*, 117–126. doi:10.1016/j.biopsycho.2015.12.003
- Ghaziuddin, M., Ghaziuddin, N., & Greden, J. (2002). Depression in persons with autism: implications for research and clinical care. *Journal of Autism and Developmental Disorders*, 32(4), 299–306.
- Gillott, A., & Standen, P. J. (2007). Levels of anxiety and sources of stress in adults with autism. *Journal of Intellectual Disabilities : JOID*, 11(4), 359–370. doi:10.1177/1744629507083585
- Hendryx, M. S., Haviland, M. G., & Shaw, D. G. (1991). Dimensions of alexithymia and their relationships to anxiety and depression. *Journal of personality assessment*, 56(2), 227-237.
- Henry, J. D., Bailey, P. E., von Hippel, C., Rendell, P. G., & Lane, A. (2010). Alexithymia in schizophrenia. *Journal of Clinical and Experimental Neuropsychology*, 32(8), 890– 897. doi:10.1080/13803391003596462
- Herbert, B. M., Herbert, C., & Pollatos, O. (2011). On the relationship between interoceptive awareness and alexithymia: is interoceptive awareness related to emotional awareness? *Journal of Personality*, 79(5), 1149–1175. doi:10.1111/j.1467-6494.2011.00717.x
- Herbert, B. M., Muth, E. R., Pollatos, O., & Herbert, C. (2012). Interoception across modalities: on the relationship between cardiac awareness and the sensitivity for gastric functions. *Plos One*, 7(5), e36646. doi:10.1371/journal.pone.0036646
- Hert, M., Correll, C. U., Bobes, J., Cetkovich Bakmas, M., Cohen, Asai, I., ... & Newcomer, J. W. (2011). Physical illness in patients with severe mental disorders. I. Prevalence, impact of medications and disparities in health care. *World psychiatry*, 10(1), 52-77.
- Honkalampi, K., Hintikka, J., Tanskanen, A., Lehtonen, J., & Viinamäki, H. (2000).
 Depression is strongly associated with alexithymia in the general population. *Journal* of Psychosomatic Research, 48(1), 99–104. doi:10.1016/S0022-3999(99)00083-5
- Kenny, L., Hattersley, C., Molins, B., Buckley, C., Povey, C., & Pellicano, E. (2016). Which terms should be used to describe autism? Perspectives from the UK autism community. *Autism: The International Journal of Research and Practice*, 20(4), 442– 462. doi:10.1177/1362361315588200
- Khalsa, S. S., Kuplicki, R., Hung-wen, Y., Clary, K., Feinstein, J, S., Simmons, K., Paulus, M, P. (2017) Interoceptive dysfunction across psychiatric disorders: preliminary results from a large-scale naturalistic study. The American College of Neuropsychopharmacology (ACNP) conference. (December, 2017).
- Khalsa, S S, Rudrauf, D., Sandesara, C., Olshansky, B., & Tranel, D. (2009). Bolus isoproterenol infusions provide a reliable method for assessing interoceptive

awareness. *International Journal of Psychophysiology*, 72(1), 34–45. doi:10.1016/j.ijpsycho.2008.08.010

- Khalsa, Sahib S, & Lapidus, R. C. (2016). Can Interoception Improve the Pragmatic Search for Biomarkers in Psychiatry? *Frontiers in Psychiatry*, 7, 121. doi:10.3389/fpsyt.2016.00121
- Khalsa, Sahib S, Rudrauf, D., & Tranel, D. (2009). Interoceptive awareness declines with age. *Psychophysiology*, *46*(6), 1130–1136. doi:10.1111/j.1469-8986.2009.00859.x
- Klabunde, M., Acheson, D. T., Boutelle, K. N., Matthews, S. C., & Kaye, W. H. (2013).
 Interoceptive sensitivity deficits in women recovered from bulimia nervosa. *Eating Behaviors*, 14(4), 488–492. doi:10.1016/j.eatbeh.2013.08.002
- Knapp-Kline, K., & Kline, J. P. (2005). Heart rate, heart rate variability, and heartbeat detection with the method of constant stimuli: slow and steady wins the race. *Biological Psychology*, 69(3), 387–396. doi:10.1016/j.biopsycho.2004.09.002
- Longarzo, M., D'Olimpio, F., Chiavazzo, A., Santangelo, G., Trojano, L., & Grossi, D. (2015). The relationships between interoception and alexithymic trait. The Self-Awareness Questionnaire in healthy subjects. *Frontiers in Psychology*, 6, 1149. doi:10.3389/fpsyg.2015.01149
- Lovibond, P. F., & Lovibond, S. H. (1995). The structure of negative emotional states: comparison of the Depression Anxiety Stress Scales (DASS) with the Beck Depression and Anxiety Inventories. *Behaviour Research and Therapy*, 33(3), 335– 343. doi:10.1016/0005-7967(94)00075-U
- Lovibond, S. H. (1993). Lovibond PF (Eds.): Manual for the Depression Anxiety Stress Scales (DASS): Psychology Foundation Monograph.
- Murphy, J., Brewer, R., Catmur, C., & Bird, G. (2017). Interoception and psychopathology: A developmental neuroscience perspective. *Developmental Cognitive Neuroscience*, 23, 45–56. doi:10.1016/j.dcn.2016.12.006
- Murphy, J., Catmur, C., & Bird, G. (2017). Alexithymia is associated with a multidomain, multidimensional failure of interoception: evidence from novel tests. *Journal of Experimental Psychology. General.* doi:10.1037/xge0000366
- Murphy, J., Geary, H., Millgate, E., Catmur, C., & Bird, G. (2017). Direct and indirect effects of age on interoceptive accuracy and awareness across the adult lifespan. *Psychonomic Bulletin & Review*, 1–10. doi:10.3758/s13423-017-1339-z
- Murphy, J., Millgate, E., Geary, H., Ichijo, E., Coll, M.-P., Brewer, R., ... Bird, G. (2018).
 Knowledge of resting heart rate mediates the relationship between intelligence and the heartbeat counting task. *Biological Psychology*. doi:10.1016/j.biopsycho.2018.01.012
- Noel, J.-P., Lytle, M., Cascio, C., & Wallace, M. T. (2018). Disrupted integration of exteroceptive and interoceptive signaling in autism spectrum disorder. *Autism Research : Official Journal of the International Society for Autism Research*, 11(1), 194–205. doi:10.1002/aur.1880
- O'Brien, W. H., Reid, G. J., & Jones, K. R. (1998). Differences in heartbeat awareness among males with higher and lower levels of systolic blood pressure. *International Journal of Psychophysiology*, 29(1), 53–63. doi:10.1016/S0167-8760(98)00004-X

- Palser, E, R., Pellicano, E., Fotopoulou, A., Kilner, J, M., (2017). Interoceptive sensibility not accuracy is linked to childhood alexithymia. Alexithymia Day, London (September, 2017).
- Pineles, S. L., & Mineka, S. (2005). Attentional biases to internal and external sources of potential threat in social anxiety. *Journal of Abnormal Psychology*, 114(2), 314–318. doi:10.1037/0021-843X.114.2.314
- Pinna, F., Lai, L., Pirarba, S., Orrù, W., Velluzzi, F., Loviselli, A., & Carpiniello, B. (2011). Obesity, alexithymia and psychopathology: a case-control study. *Eating and Weight Disorders*, 16(3), e164–70. doi:10.3275/7509
- Pollatos, O., Herbert, B. M., Mai, S., & Kammer, T. (2016). Changes in interoceptive processes following brain stimulation. *Philosophical Transactions of the Royal Society of London. Series B, Biological Sciences*, 371(1708). doi:10.1098/rstb.2016.0016
- Pollatos, O., Kurz, A.-L., Albrecht, J., Schreder, T., Kleemann, A. M., Schöpf, V., ... Schandry, R. (2008). Reduced perception of bodily signals in anorexia nervosa. *Eating Behaviors*, 9(4), 381–388. doi:10.1016/j.eatbeh.2008.02.001
- Pollatos, O., Traut-Mattausch, E., & Schandry, R. (2009). Differential effects of anxiety and depression on interoceptive accuracy. *Depression and Anxiety*, 26(2), 167–173. doi:10.1002/da.20504
- Quattrocki, E., & Friston, K. (2014). Autism, oxytocin and interoception. *Neuroscience and Biobehavioral Reviews*, 47, 410–430. doi:10.1016/j.neubiorev.2014.09.012
- Ramaekers, D., Ector, H., Aubert, A. E., Rubens, A., & Van de Werf, F. (1998). Heart rate variability and heart rate in healthy volunteers. Is the female autonomic nervous system cardioprotective? *European Heart Journal*, 19(9), 1334–1341. doi:10.1053/euhj.1998.1084
- Ring, C., & Brener, J. (2018). Heartbeat counting is unrelated to heartbeat detection: A comparison of methods to quantify interoception. *Psychophysiology*, e13084.
- Ring, C., & Brener, J. (1996). Influence of beliefs about heart rate and actual heart rate on heartbeat counting. *Psychophysiology*, 33(5), 541–546. doi:10.1111/j.1469-8986.1996.tb02430.x
- Ring, C., Brener, J., Knapp, K., & Mailloux, J. (2015). Effects of heartbeat feedback on beliefs about heart rate and heartbeat counting: a cautionary tale about interoceptive awareness. *Biological Psychology*, 104, 193–198. doi:10.1016/j.biopsycho.2014.12.010
- Roth, W. T., Margraf, J., Ehlers, A., Haddad, J. M., Maddock, R. J., Agras, W. S., & Taylor, C. B. (1992). Imipramine and alprazolam effects on stress test reactivity in panic disorder. *Biological Psychiatry*, 31(1), 35–51.
- Rouse, C. H., Jones, G. E., & Jones, K. R. (1988). The effect of body composition and gender on cardiac awareness. *Psychophysiology*, 25(4), 400–407. doi:10.1111/j.1469-8986.1988.tb01876.x
- Schandry, R. (1981). Heart Beat Perception and Emotional Experience. *Psychophysiology*, *18*(4), 483–488. doi:10.1111/j.1469-8986.1981.tb02486.x

- Schauder, K. B., Mash, L. E., Bryant, L. K., & Cascio, C. J. (2015). Interoceptive ability and body awareness in autism spectrum disorder. *Journal of Experimental Child Psychology*, 131, 193–200. doi:10.1016/j.jecp.2014.11.002
- Seth, A. K. (2013). Interoceptive inference, emotion, and the embodied self. *Trends in Cognitive Sciences*, 17(11), 565–573. doi:10.1016/j.tics.2013.09.007
- Shah, P., Hall, R., Catmur, C., & Bird, G. (2016). Alexithymia, not autism, is associated with impaired interoception. *Cortex*, *81*, 215–220. doi:10.1016/j.cortex.2016.03.021
- Spielberger, C. D., Gorsuch, R. L., & Lushene, R. E. (1970). Manual for the state-trait anxiety inventory.
- Steptoe, A., & Vögele, C. (1992). Individual differences in the perception of bodily sensations: the role of trait anxiety and coping style. *Behaviour Research and Therapy*, 30(6), 597–607. doi:10.1016/0005-7967(92)90005-2
- Van Der Cruijsen, R., Murphy, J., Crone., E & Bird., G (in preparation). Alexithymia and the perception of muscular effort in adolescents.
- Wald, J., & Taylor, S. (2005). Interoceptive exposure therapy combined with trauma-related exposure therapy for post-traumatic stress disorder: a case report. *Cognitive Behaviour Therapy*, 34(1), 34–40. doi:10.1080/16506070510010648
- White, K. S., Brown, T. A., Somers, T. J., & Barlow, D. H. (2006). Avoidance behavior in panic disorder: the moderating influence of perceived control. *Behaviour Research* and Therapy, 44(1), 147–157. doi:10.1016/j.brat.2005.07.009
- Whitehead, W. E., & Drescher, V. M. (1980). Perception of Gastric Contractions and Self-Control of Gastric Motility. *Psychophysiology*, 17(6), 552–558. doi:10.1111/j.1469-8986.1980.tb02296.x
- Whitehead, W. E., Drescher, V. M., Heiman, P., & Blackwell, B. (1977). Relation of heart rate control to heartbeat perception. *Biofeedback and Self-regulation*, 2(4), 371–392. doi:10.1007/BF00998623
- Willem Van der Does, A. J., Antony, M. M., Ehlers, A., & Barsky, A. J. (2000). Heartbeat perception in panic disorder: a reanalysis. *Behaviour Research and Therapy*, 38(1), 47–62. doi:10.1016/S0005-7967(98)00184-3
- Windmann, S., Schonecke, O. W., Fröhlig, G., & Maldener, G. (1999). Dissociating beliefs about heart rates and actual heart rates in patients with cardiac pacemakers. *Psychophysiology*, 36(3), 339–342. doi:10.1017/S0048577299980381 Wittmann,
- M. (2013). The inner sense of time: how the brain creates a representation of duration. *Nature Reviews. Neuroscience*, 14(3), 217–223. doi:10.1038/nrn3452
- Yuen, K. K. (1974). 2-sample trimmed t for unequal population variances. *Biometrika*, *61*(1), 165-170.
- Zamariola, G., Vlemincx, E., Luminet, O., & Corneille, O. (2018). Relationship between interoceptive accuracy, interoceptive sensibility, and alexithymia. *Personality and Individual Differences*, *125*, 14-20.
- Zoellner, L. A., & Craske, M. G. (1999). Interoceptive accuracy and panic. *Behaviour Research and Therapy*, *37*(12), 1141–1158.
- Zhang, Y., Murphy, J., Bird, G., & Lau., J. (in preparation). Alexithymia is associated with poor interoception in adolescent females.

Table 1.

	Descriptives						Zero-order correlations							Partial Correlation	
Variable	Ν	Mean	SD	1	2	3	4	5	6	7	8	9	10	11	
1. Age	287	38.07	21.09	1											090
2. HCT	287	192.77	117.94	173**	1										-
3. Time	287	292.20	61.08	004	.163**	1									087
4. Knowledge	209	12.37	11.61	042	137*	.025	1								125
5. TAS-20	287	45.26	13.00	056	079	.045	025	1							-
6. Depression	286	-0.08	0.80	008	009	.025	.211**	.478**	1						083
7. Anxiety	287	-0.07	0.97	188**	016	002	.123	.498**	.694**	1					082
8. BMI	280	23.24	4.55	.348**	108	003	.212**	025	.081	039	1				098
9. Systolic BP	183	124.04	18.64	.595**	136	107	.055	.011	.012	048	.367**	1			169*
10. HRV	180	5.64	1.73	331**	.016	098	053	.036	.021	017	072	141	1		179*
11. Mean HR	268	73.65	12.66	247**	094	074	067	.099	.139*	.110	117	023	- .148 [*]	1	084

*denotes significant at p < .05, **denotes significant at p < .01

Model parameters	Т	р	β	F	df	р	Partial correlation TAS-20 HCT
Overall Model				2.297	11, 162	.012	
TAS-20	-2.486	.014	222				192
Age	901	.369	094				
Gender	3.000	.003	.235				
Time	1.299	.196	.099				
Anxiety	357	.722	039				
Depression	.807	.421	.090				
BMI	571	.569	048				
Mean HR	793	.429	063				
Knowledge	-2.026	.044	162				
Systolic BP	955	.341	092				
HRV	937	.350	077				

Table 2. Final Model

Table legends

Table 1. Provides the descriptive statistics and correlations between all measured variables. Partial correlations between alexithymia and HCT performance controlling for each variable are also reported. TAS-20 = Alexithymia scores, HCT = heartbeat counting scores, Age = age in years, Gender = 0 = females, 1 = males, Time = scores on the time estimation task, BMI = body mass index, Knowledge = inaccuracy of estimates regarding the average resting heart rate (see text for details), HRV = heart rate variability, Depression = Z-score depression scores, Anxiety = Z-score anxiety scores (see text for details), Systolic BP = Systolic blood pressure, Mean HR = the participant's average resting heart rate.

Table 2. Depicts the results of the final regression analysis predicting scores on the HCT from alexithymia after the inclusion of all control variables (for each step please see supplementary results). As can be seen, after controlling for all variables a significant relationship between alexithymia and HCT performance was observed. TAS-20 = Alexithymia scores, Age = age in years, Gender = 0 = females, 1 = males, Time = scores on the time estimation task, BMI = body mass index, Knowledge = inaccuracy of estimates regarding the average resting heart rate (see text for details), HRV = heart rate variability, Depression = Z-score depression scores, Anxiety = Z-score anxiety scores (see text for details), Systolic BP = Systolic blood pressure, Mean HR = the participant's average resting heart rate.

Supplementary Results

Table S1.

Model parameters	t	р	β	F	df	Р	Partial correlation TAS-20 HCT
Overall Model	-1 526	128	- 089	5.588	2, 284	.004	- 090
Age	-3.056	.002	178				090
Overall Model				6.051	3,283	.001	
TAS-20	-1.762	.079	102				104
Age	-3.373	.001	196				
Gender	2.598	.010	.151				
Overall Model				6.099	4,282	<.001	
TAS-20	-1.844	.066	106				109
Age	-3.340	.001	193				
Gender	2.084	.038	.123				
Time	2.435	.016	.142				
Overall Model				4.868	5, 281	<.001	
TAS-20	-1.677	.095	112				100
Age	-3.253	.001	191				
Gender	2.087	.038	.124				
Time	2.432	.016	.142				
Anxiety	.171	.864	.012				
Overall Model				4.043	6, 279	.001	
TAS-20	-1.769	.078	121		-		105
Age	-3.251	.001	194				
Gender	2.038	.043	.121				
Time	2.439	.015	.143				
Anxiety	357	.722	030				

Depression	.794	.428	.066				
Overall Model				3.715	7, 271	.001	
TAS-20	-2.097	.037	146				126
Age	-2.596	.010	165				
Gender	2.352	.019	.144				
Time	2.179	.030	.130				
Anxiety	479	.632	041				
Depression	1.061	.290	.090				
BMI	-1.331	.184	084				
Overall Model				3.444	8, 253	.001	
TAS-20	-2.109	.036	152				131
Age	-3.018	.003	202				
Gender	2.282	.023	.143				
Time	1.831	.068	.113				
Anxiety	515	.607	047				
Depression	1.239	.216	.110				
BMI	630	.529	041				
Mean HR	-1.850	.065	117				
Overall Model				3.277	9, 191	.001	
TAS-20	-2.484	.014	208				177
Age	-2.332	.021	175				
Gender	2.637	.009	.187				
Time	1.872	.063	.129				
Anxiety	585	.559	059				
Depression	1.459	.146	.148				
BMI	688	.492	052				
Mean HR	-1.607	.110	112				
Knowledge	-2.477	.014	180				

	0						
Overall Model				2.440	10, 163	.010	
TAS-20	-2.516	.013	225				193
Age	601	.549	059				
Gender	2.905	.004	.225				
Time	1.488	.139	.112				
Anxiety	223	.824	024				
Depression	.657	.512	.072				
BMI	577	.565	048				
Mean HR	586	.559	045				
Knowledge	-1.927	.056	153				
Systolic BP	-1.031	.304	099				
Overall Model				2.297	11, 162	.012	
Overall Model TAS-20	-2.486	.014	222	2.297	11, 162	.012	192
Overall Model TAS-20 Age	-2.486 901	.014 .369	222 094	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender	-2.486 901 3.000	.014 .369 .003	222 094 .235	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time	-2.486 901 3.000 1.299	.014 .369 .003 .196	222 094 .235 .099	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety	-2.486 901 3.000 1.299 357	.014 .369 .003 .196 .722	222 094 .235 .099 039	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety Depression	-2.486 901 3.000 1.299 357 .807	.014 .369 .003 .196 .722 .421	222 094 .235 .099 039 .090	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety Depression BMI	-2.486 901 3.000 1.299 357 .807 571	.014 .369 .003 .196 .722 .421 .569	222 094 .235 .099 039 .090 048	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety Depression BMI Mean HR	-2.486 901 3.000 1.299 357 .807 571 793	.014 .369 .003 .196 .722 .421 .569 .429	222 094 .235 .099 039 .090 048 063	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety Depression BMI Mean HR Knowledge	-2.486 901 3.000 1.299 357 .807 571 793 -2.026	.014 .369 .003 .196 .722 .421 .569 .429 .044	222 094 .235 .099 039 .090 048 063 162	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety Depression BMI Mean HR Knowledge Systolic BP	-2.486 901 3.000 1.299 357 .807 571 793 -2.026 955	.014 .369 .003 .196 .722 .421 .569 .429 .044 .341	222 094 .235 .099 039 .090 048 063 162 092	2.297	11, 162	.012	192
Overall Model TAS-20 Age Gender Time Anxiety Depression BMI Mean HR Knowledge Systolic BP HRV	-2.486 901 3.000 1.299 357 .807 571 793 -2.026 955 937	.014 .369 .003 .196 .722 .421 .569 .429 .044 .341 .350	222 094 .235 .099 039 .090 048 063 162 092 077	2.297	11, 162	.012	192

Table S2.

Model parameters	t	р	β	F	df	р	Partial TAS-HCT
Overall Model				2.693	11, 133	.004	
TAS-20 (Grouped)	-2.442	.016	230				207
Age	-1.390	.167	162				
Gender	3.064	.003	.260				
Time	.892	.374	.075				
Anxiety	212	.832	025				
Depression	.510	.611	.063				
BMI	778	.438	069				
Mean HR	-1.889	.061	169				
Knowledge	-2.270	.025	192				
Systolic BP	743	.459	078				
HRV	-1.370	.173	123				

Table S3. Final Model (with mean replacement)

Model parameters	t	р	β	F	df	р	Partial TAS-HCT
Overall Model				3.876	11, 275	<.001	
TAS-20	-2.119	.035	143				127
Age	-3.524	.000	261				
Gender	2.534	.012	.151				
Time	2.057	.041	.120				
Anxiety	684	.494	058				
Depression	1.822	.069	.153				
BMI	832	.406	052				
Mean HR	-2.641	.009	160				
Knowledge	-2.962	.003	175				
Systolic BP	.102	.919	.007				
HRV	-1.450	.148	088				





Standardized Residual

Table and Figure legends

Table S1. Shows the results of a series of regression analyses predicting scores on the HCT from alexithymia after the sequential entry of each control variable. As can be seen, the relationship between alexithymia and performance on the HCT varied greatly depending on which control variables were entered, and the relationship between specific control variables and the HCT also varied depending on which other control variables were entered. TAS-20 = Alexithymia scores, Age = age in years, Gender = 0 = females, 1 = males, Time = scores on the time estimation task, BMI = body mass index, Knowledge = inaccuracy of estimates regarding the average resting heart rate (see main text for details), HRV = heart rate variability, Depression = Z-score depression scores, Anxiety = Z-score anxiety scores (see main text for details), Systolic BP = Systolic blood pressure, Mean HR = the participant's average resting heart rate.

Table S2. Depicts the results of a regression analysis predicting scores on the HCT from alexithymia (grouped by cut-off scores) after the entry of each control variable. As can be seen, the relationship between alexithymia and performance on the HCT excluding individuals with borderline scores (between 50-60) was greater than when these individuals were included. TAS-20 = Alexithymia scores (0 = low alexithymia (<50), 1 = high alexithymia (>60), Age = age in years, Gender = 0 = females, 1 = males, Time = scores on the time estimation task, BMI = body mass index, Knowledge = inaccuracy of estimates regarding the average resting heart rate (see text for details), HRV = heart rate variability, Depression = Z-score depression scores, Anxiety = Z-score anxiety scores (see text for details), Systolic BP = Systolic blood pressure, Mean HR = the participant's average resting heart rate.

Table S3. Depicts the results of the final regression analysis predicting scores on the HCT from alexithymia after the inclusion of all control variables but with mean replacement (i.e., missing values were replaced with the mean value). TAS-20 = Alexithymia scores, Age = age in years, Gender = 0 = females, 1 = males, Time = scores on the time estimation task, BMI = body mass index, Knowledge = inaccuracy of estimates regarding the average resting heart rate (see text for details), HRV = heart rate variability, Depression = Z-score depression scores, Anxiety = Z-score anxiety scores (see text for details), Systolic BP = Systolic blood pressure, Mean HR = the participant's average resting heart rate.

Figure S1. Depicts the residual plots for each regression analysis predicting HCT performance. Panels a) – i) refer to Table S1 (Supplementary Results). Panel j) refers to Table S1 (Supplementary Results) and Table 2 (Main Text). Panel k) refers to Table S2 (Supplementary Results). Panel I) refers to Table S3 (Supplementary Results). Predictors: a) Age, TAS-20. b) Age, TAS-20, Gender. c) Age, TAS-20, Gender, Time. d) Age, TAS-20, Gender, Time, Anxiety. e) Age, TAS-20, Gender, Time, Anxiety, Depression. f) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI. g) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR. h) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge. i) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP. j) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV. k) Age, TAS-20 (grouped), Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV I) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV I) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV I) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV I) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV I) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV I) Age, TAS-20, Gender, Time, Anxiety, Depression, BMI, Mean HR, Knowledge, Systolic BP, HRV (with mean replacement).

Supplementary Discussion

S1. The Measurement of Alexithymia is Problematic

Alexithymia is typically measured using the TAS-20, a measure with very good psychometric properties concerning its reliability and factor structure (e.g., Bagby, Parker, et al., 1994; Bagby, Taylor, & Parker, 1994; Parker, Bagby, Taylor, Endler, & Schmitz, 1993), including in clinical groups (e.g. Bagby, Taylor, et al., 1994; Berthoz & Hill, 2005; Loas et al., 2001; Moriguchi et al., 2007). Use of the TAS-20 to measure alexithymia has potential limitations, however, and these are briefly explored here, along with other factors relating to its use. First, it could be argued that the use of a self-report tool to assess difficulties with emotional self-awareness is problematic; individuals with very severe alexithymia may be unaware that they have difficulties and report low levels of alexithymia. Although a logical possibility, there is little empirical support for this position, perhaps due to the difficulties involved in validating objective measures of alexithymia. For example, while correlations between the TAS-20 and observer measures of alexithymia are statistically significant but modest (e.g., Bagby, Taylor, et al., 1994; Lichev et al., 2014; Meganck, Vanheule, Desmet, & Inslegers, 2010), it is unclear whether the two measures should necessarily correlate highly, and whether one would expect self- or observer-report to be more accurate. One other major (although less frequently used) self-report measure of alexithymia is the Bermond-Vorst Alexithymia questionnaire (BVAQ; Vorst & Bermond, 2001), and correlations between the TAS-20 and the BVAQ are relatively strong (~.60; Morera, Culhane, Watson, & Skewes, 2005; Vorst & Bermond, 2001). Perhaps better evidence for the validity of these self-report measures of alexithymia comes from a series of studies demonstrating a 'decoupling' between objective and subjective measures of emotional arousal in those with self-reported high alexithymia, such that individuals reporting alexithymic traits are less able to accurately

report their degree of arousal (an expected consequence of alexithymia; e.g., Connelly & Denney, 2007; Eastabrook, Lanteigne, & Hollenstein, 2013; Gaigg, Cornell, & Bird, 2016; Newton & Contrada, 1994; Pollatos et al., 2011; Stone & Nielson, 2001).

It should be noted however, that both the TAS-20 and BVAQ have a number of subscales. The TAS-20, for example, has three subscales: Difficulties Identifying Feelings (DIF), Difficulties Describing Feelings (DDF), and Externally Oriented Thinking (EOT). Typically, studies find scores on the DIF and DDF subscales are correlated within samples, but the relationship between these factors and EOT, and the reliability of EOT, is far weaker (e.g., Kooiman, Spinhoven, & Trijsburg, 2002). Conceptually, it is possible to speculate that the DIF and DDF subscales should be better associated with interoceptive sensitivity, while EOT scores might instead be associated with the propensity to become aware of, and use, interoceptive information. Regardless of the accuracy of this conjecture, given the fact that two of the TAS-20 subscales are more highly correlated with each other than the third, and that the subscales do not contribute equally to total scores (although the TAS-20 total score is a sum of all item scores, there are 7 items contributing to the DIF score, 5 items for the DDF score, and 8 items for the EOT score), it makes sense that total scores may be less reliable in terms of their association with interoception than subscale scores. This is because a given TAS-20 total score may reflect very different subscales scores across individuals. This highlights the requirement for studies of interoception in those with alexithymia to include a large proportion of individuals who have high alexithymia scores (at least above the cut-off score of 61 on the TAS-20). These individuals will be comparable to each other as they will have high scores on all three subscales of the TAS-20. Indeed, in the current data, when one groups and includes only those individuals scoring above the clinically-relevant cut-off for high alexithymia and below the cut-off for moderate alexithymia (<50) then the association between alexithymia and HCT performance is even stronger (Table S2 Supplementary

Results; *r*(partial) =.207). In addition to these potential problems when measuring alexithymia in adults, measurement of alexithymia may be especially problematic in children. Although a version of the TAS-20 has been developed for children (Rieffe, Oosterveld, & Terwogt, 2006) the questionnaire provides an unreliable estimate of some subscales (Loas, Braun, Delhaye, & Linkowski, 2017; Rieffe et al., 2006), and its factor structure has been questioned (Rieffe et al., 2006). Some studies have also failed to find a significant correlation between child and parent report measures of children's alexithymia (Griffin, Lombardo, & Auyeung, 2016). This discrepancy between parent and child reports could suggest that children are inaccurate reporters of their own alexithymic traits, but as stated above it is unclear to what extent we would expect a strong correlation between these reports, and whether parent report should necessarily be considered more accurate.

S2. Multiple Routes to Alexithymia

One potential reason for variance in the size of the correlation between alexithymia and HCT performance across studies is that the population of individuals with alexithymia is itself not homogeneous. Alexithymia may be the end-product of atypical functioning in any one of a number of neurocognitive systems. If there are two or more 'routes' to alexithymia, only one of which arises from, or is associated with, impaired interoception, then (assuming the HCT is a measure of interoceptive sensitivity), one may expect the association between alexithymia and HCT performance to vary according to the proportion of alexithymic individuals in a particular sample who develop alexithymia as a result of an interoceptionrelated impairment.

Although under-investigated, there exist several pieces of evidence that support the idea that there may be multiple routes to alexithymia, in addition to an interoceptive deficit. Two candidate routes are language impairment, and impaired executive function. With respect to language, there is evidence of disrupted language development, and weak verbal abilities, in developmental and acquired cases of alexithymia, respectively. For example, longitudinal studies show that problems with early language development at age 1 and 5 years are predictive of alexithymia in adulthood (Karukivi et al., 2012; Kokkonen et al., 2003). In addition, in populations with acquired alexithymia, damage to language regions in the inferior frontal gyrus are predictive of alexithymia (Hobson et al., 2018), and alexithymia in patients with traumatic brain injury (TBI) correlates negatively with verbal abilities (Lamberty & Holt, 1995; Wood & Williams, 2007).

With respect to executive function, it is also the case that executive dysfunction is predictive of alexithymia in patients with traumatic brain injury (TBI; Henry, Phillips, Crawford, Theodorou, & Summers, 2006; though see Wood & Williams (2007) for evidence that such an association may be driven by verbal measures of executive function). Unlike in TBI, in Parkinson's Disease alexithymia is related to visuo-spatial and non-verbal executive abilities, and does not appear to be related to be related to verbal abilities (Bogdanova & Cronin-Golomb, 2013; Costa, Peppe, Carlesimo, Salamone, & Caltagirone, 2007). In comparison, in those suffering from neurodegeneration due to Human Immunodeficiency Virus (HIV), alexithymia is associated with visuo-spatial abilities and general executive function ability (Bogdanova, Díaz-Santos, & Cronin-Golomb, 2010; McIntosh et al., 2014). In these apparently rather different populations, the association between alexithymia and executive ability may represent shared dependence upon frontal brain systems, regions which are disrupted in both Parkinson's Disease and HIV.

Such evidence is indicative of multiple routes to alexithymia (possibly interoceptive, linguistic and executive), and also demonstrates that different populations may be comprised primarily of alexithymic individuals with a specific pattern of deficits. For example, although such studies are yet to be conducted, it may be the case that individuals with developmental language disorder exhibit high levels of alexithymia in the presence of perfectly intact interoceptive ability. Similarly, as the relationship between executive functioning and alexithymia has only been observed thus far in those with neurological damage, executive functioning may not predict alexithymia in the typical population. Taken together, this evidence underscores the argument that alexithymia may arise through multiple routes. This possibility further complicates examination of the relationship between alexithymia and interoception (regardless of the interoceptive measure employed), suggesting that even if alexithymia and interoceptive sensitivity is associated in some populations, it is plausible that in other populations alexithymia may occur in the absence of interoceptive difficulty.

References

- Bagby, R. M., Parker, J. D., & Taylor, G. J. (1994). The twenty-item Toronto Alexithymia Scale--I. Item selection and cross-validation of the factor structure. *Journal of Psychosomatic Research*, 38(1), 23–32. doi:10.1016/0022-3999(94)90005-1
- Bagby, R. M., Taylor, G. J., & Parker, J. D. (1994). The Twenty-item Toronto Alexithymia Scale--II. Convergent, discriminant, and concurrent validity. *Journal of Psychosomatic Research*, 38(1), 33–40.
- Berthoz, S., & Hill, E. L. (2005). The validity of using self-reports to assess emotion regulation abilities in adults with autism spectrum disorder. *European Psychiatry*, 20(3), 291–298. doi:10.1016/j.eurpsy.2004.06.013
- Bogdanova, Y., & Cronin-Golomb, A. (2013). Alexithymia and apathy in Parkinson's disease: neurocognitive correlates. *Behavioural Neurology*, *27*(4), 535–545. doi:10.3233/BEN-129021
- Bogdanova, Y., Díaz-Santos, M., & Cronin-Golomb, A. (2010). Neurocognitive correlates of alexithymia in asymptomatic individuals with HIV. *Neuropsychologia*, 48(5), 1295– 1304. doi:10.1016/j.neuropsychologia.2009.12.033
- Connelly, M., & Denney, D. R. (2007). Regulation of emotions during experimental stress in alexithymia. *Journal of Psychosomatic Research*, 62(6), 649–656. doi:10.1016/j.jpsychores.2006.12.008
- Costa, A., Peppe, A., Carlesimo, G. A., Salamone, G., & Caltagirone, C. (2007). Neuropsychological correlates of alexithymia in Parkinson's disease. *Journal of the International Neuropsychological Society*, 13(6), 980–992. doi:10.1017/S1355617707071329
- Eastabrook, J. M., Lanteigne, D. M., & Hollenstein, T. (2013). Decoupling between physiological, self-reported, and expressed emotional responses in alexithymia. *Personality and Individual Differences*, 55(8), 978–982. doi:10.1016/j.paid.2013.08.001
- Gaigg, S. B., Cornell, A. S., & Bird, G. (2016). The psychophysiological mechanisms of alexithymia in autism spectrum disorder. *Autism: The International Journal of Research and Practice*. doi:10.1177/1362361316667062
- Griffin, C., Lombardo, M. V., & Auyeung, B. (2016). Alexithymia in children with and without autism spectrum disorders. *Autism Research : Official Journal of the International Society for Autism Research*, 9(7), 773–780. doi:10.1002/aur.1569
- Henry, J. D., Phillips, L. H., Crawford, J. R., Theodorou, G., & Summers, F. (2006).
 Cognitive and psychosocial correlates of alexithymia following traumatic brain injury.
 Neuropsychologia, 44(1), 62–72. doi:10.1016/j.neuropsychologia.2005.04.011
- Hobson, H., Hogeveen, J., Brewer, R., Catmur, C., Gordon, B., Krueger, F., ... Grafman, J. (2018). Language and alexithymia: Evidence for the role of the inferior frontal gyrus in acquired alexithymia. *Neuropsychologia*. doi:10.1016/j.neuropsychologia.2017.12.037
- Karukivi, M., Joukamaa, M., Hautala, L., Kaleva, O., Haapasalo-Pesu, K.-M., Liuksila, P.-R., & Saarijärvi, S. (2012). Deficit in speech development at the age of 5 years predicts

alexithymia in late-adolescent males. *Comprehensive Psychiatry*, 53(1), 54–62. doi:10.1016/j.comppsych.2011.01.012

- Kokkonen, P., Veijola, J., Karvonen, J. T., Läksy, K., Jokelainen, J., Järvelin, M.-R., & Joukamaa, M. (2003). Ability to speak at the age of 1 year and alexithymia 30 years later. *Journal of Psychosomatic Research*, 54(5), 491–495.
- Kooiman, C. G., Spinhoven, P., & Trijsburg, R. W. (2002). The assessment of alexithymia: a critical review of the literature and a psychometric study of the Toronto Alexithymia Scale-20. *Journal of Psychosomatic Research*, 53(6), 1083–1090.
- Lamberty, G. J., & Holt, C. S. (1995). Evidence for a verbal deficit in alexithymia. *The Journal of Neuropsychiatry and Clinical Neurosciences*, 7(3), 320–324. doi:10.1176/jnp.7.3.320
- Lichev, V., Rufer, M., Rosenberg, N., Ihme, K., Grabe, H.-J., Kugel, H., ... Suslow, T. (2014). Assessing alexithymia and emotional awareness: relations between measures in a German non-clinical sample. *Comprehensive Psychiatry*, 55(4), 952–959. doi:10.1016/j.comppsych.2013.12.013
- Loas, G, Braun, S., Delhaye, M., & Linkowski, P. (2017). The measurement of alexithymia in children and adolescents: Psychometric properties of the Alexithymia Questionnaire for Children and the twenty-item Toronto Alexithymia Scale in different non-clinical and clinical samples of children and adolescents. *Plos One*, *12*(5), e0177982. doi:10.1371/journal.pone.0177982
- Loas, G, Corcos, M., Stephan, P., Pellet, J., Bizouard, P., Venisse, J. L., ... Réseau (2001). Factorial structure of the 20-item Toronto Alexithymia Scale: confirmatory factorial analyses in nonclinical and clinical samples. *Journal of Psychosomatic Research*, 50(5), 255–261.
- McIntosh, R. C., Ironson, G., Antoni, M., Kumar, M., Fletcher, M. A., & Schneiderman, N. (2014). Alexithymia is linked to neurocognitive, psychological, neuroendocrine, and immune dysfunction in persons living with HIV. *Brain, Behavior, and Immunity*, 36, 165–175. doi:10.1016/j.bbi.2013.10.024
- Meganck, R., Vanheule, S., Desmet, M., & Inslegers, R. (2010). The Observer Alexithymia Scale: a reliable and valid alternative for alexithymia measurement?. *Journal of personality assessment, 92*(2), 175-185.
- Morera, O. F., Culhane, S. E., Watson, P. J., & Skewes, M. C. (2005). Assessing the reliability and validity of the Bermond-Vorst Alexithymia Questionnaire among U.S. Anglo and U.S. Hispanic samples. *Journal of Psychosomatic Research*, 58(3), 289–298. doi:10.1016/j.jpsychores.2004.09.001
- Moriguchi, Y., Maeda, M., Igarashi, T., Ishikawa, T., Shoji, M., Kubo, C., & Komaki, G. (2007). Age and gender effect on alexithymia in large, Japanese community and clinical samples: a cross-validation study of the Toronto Alexithymia Scale (TAS-20). *BioPsychoSocial Medicine*, 1, 7. doi:10.1186/1751-0759-1-7
- Newton, T. L., & Contrada, R. J. (1994). Alexithymia and repression: contrasting emotionfocused coping styles. *Psychosomatic Medicine*, 56(5), 457–462.
- Parker, J. D. A., Bagby, R., Taylor, G. J., Endler, N. S., & Schmitz, P. (1993). Factorial validity of the 20-item Toronto Alexithymia Scale. *European Journal of Personality*, 7(4), 221–232. doi:10.1002/per.2410070403

- Pollatos, O., Werner, N. S., Duschek, S., Schandry, R., Matthias, E., Traut-Mattausch, E., & Herbert, B. M. (2011). Differential effects of alexithymia subscales on autonomic reactivity and anxiety during social stress. *Journal of Psychosomatic Research*, 70(6), 525–533. doi:10.1016/j.jpsychores.2010.12.003
- Rieffe, C., Oosterveld, P., & Terwogt, M. M. (2006). An alexithymia questionnaire for children: Factorial and concurrent validation results. *Personality and Individual Differences*, 40(1), 123–133. doi:10.1016/j.paid.2005.05.013
- Stone, L. A., & Nielson, K. A. (2001). Intact physiological response to arousal with impaired emotional recognition in alexithymia. *Psychotherapy and Psychosomatics*, 70(2), 92– 102. doi:10.1159/000056232
- Vorst, H. C. M., & Bermond, B. (2001). Validity and reliability of the Bermond–Vorst alexithymia questionnaire. *Personality and Individual Differences*, 30, 413-434.
- Wood, R. L., & Williams, C. (2007). Neuropsychological correlates of organic alexithymia. Journal of the International Neuropsychological Society, 13(3), 471–479. doi:10.1017/S1355617707070518