Contextual influence of reinforcement learning performance in Anorexia Nervosa

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Abstract

Objective: This study utilized a within-subject design to examine whether individuals with 15 restrictive anorexia nervosa (R-AN; n=40) perform similarly to healthy controls (HCs; n=16 45) and healthy controls at risk of eating disorders (RI; n=36) in a reinforcement learning 17 (RL) tasks. Specifically, we aimed to determine if RL performance is comparable between 18 groups for disorder-unrelated choices, but significantly impaired for disorder-related choices. 19 Method: RL performance was assessed using a Probabilistic Reversal Learning (PRL) task, 20 where participants were asked to perform disorder-related choices or disorder-unrelated 21 choices. Results: R-AN individuals demonstrated lower learning rates for disorder-related 22 decisions, while their performance on neutral decisions was comparable to participants with 23 Bulimia Nervosa, Healthy Controls (HCs), and HCs at risk of eating disorders. Additionally, only AN patients exhibited reduced learning rates for outcome-irrelevant food-related 25 decisions in reward-based learning, as opposed to food-unrelated decisions. **Discussion:** Impaired RL task performance in individuals with AN may be attributed to external factors 27 rather than compromised learning mechanisms. These findings indicate that AN may significantly impact the cognitive processing of food-related information, even when AN patients do not show learning rate disadvantages compared to HCs in decision-making involving food-unrelated information. This study provides valuable insights into the 31 reinforcement learning processes of individuals with AN and emphasizes the need to consider the influence of food-related information on cognitive functioning in this patient population. 33 The findings have potential implications for the development of interventions targeting decision- making processes in individuals with AN

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Introduction

Anorexia Nervosa (AN) is one of the most common eating disorders characterized by distorted body perception and pathological weight loss, particularly in its restricting type (R-AN) (American Psychiatric Association, 2022). Lifetime prevalence for AN has been reported at 1.4% for women and 0.2% for men (Galmiche, Déchelotte, Lambert, & Tavolacci, 2019; Smink, Hoeken, & Hoek, 2013), with a mortality rate that can be as high as 5-20% (Qian et al., 2022). Treating AN is extremely challenging (Atwood & Friedman, 2020; Linardon, Fairburn, Fitzsimmons-Craft, Wilfley, & Brennan, 2017), highlighting the importance of gaining a deeper understanding of its underlying mechanisms (Chang, Delgadillo, & Waller, 2021).

Executive functions have gained significant attention in the research on understanding 49 the mechanisms underlying Anorexia Nervosa (AN). Impairments in executive processes, such as cognitive inflexibility, decision-making difficulties, and inhibitory control problems, have been identified as potential risk and perpetuating factors in AN (Bartholdy, Dalton, O'Daly, Campbell, & Schmidt, 2016; Guillaume et al., 2015; Wu et al., 2014). Within this domain, Reinforcement Learning (RL) in the context of associative learning has received considerable interest. In fact, the presence of persistent maladaptive eating behaviors, despite experiencing negative consequences, along with evidence suggesting changes in responsiveness to rewards and punishments, has led researchers to propose that there may be 57 abnormal reward processing and learning in AN (Schaefer & Steinglass, 2021). Although there is substantial evidence supporting the existence of anomalies in reward sensitivity, our understanding of potential abnormalities specifically related to the learning processes in AN is currently limited. 61

In relation to dysfunctions reward responsiveness among individuals with AN, research has revealed that the intense levels of dietary restriction and physical activity typically

associated with AN can trigger the activation of reward pathways in the brain (Keating, 2010; Keating, Tilbrook, Rossell, Enticott, & Fitzgerald, 2012; Selby & Coniglio, 2020). Additionally, individuals with AN may exhibit diminished reward responses specifically towards food-related stimuli (Wierenga et al., 2014). In a broader sense, research has shown 67 that AN is associated with reduced subjective reward sensitivity and decreased neural response to rewarding stimuli. Moreover, individuals with AN may experience disruptions in processing aversive stimuli, leading to heightened harm avoidance, reduced tolerance for uncertainty, increased anxiety, and heightened sensitivity to punishment (Fladung, Schulze, 71 Schöll, Bauer, & Groen, 2013; Jappe et al., 2011; Keating et al., 2012; O'Hara, Campbell, & Schmidt, 2015). These factors collectively contribute to an altered response to negative feedback and a propensity to actively avoid aversive outcomes (Jonker, Glashouwer, & Jong, 2022; Matton, Goossens, Braet, & Vervaet, 2013). Neuroimaging studies have further supported these findings by revealing neural dysfunctions in individuals with AN regarding their response to loss and aversive taste stimuli (Bischoff-Grethe et al., 2013; Monteleone et al., 2017; Wagner et al., 2007).

Given the crucial importance of RL in acquiring knowledge from past experiences,
extensive research has been conducted to examine potential deficits in RL among individuals
diagnosed with AN (Bischoff-Grethe et al., 2013; Glashouwer, Bloot, Veenstra, Franken, &
Jong, 2014; Harrison, Genders, Davies, Treasure, & Tchanturia, 2011; Jappe et al., 2011;
Matton et al., 2013). However, the findings of these studies have been varied. For instance,
Ritschel et al. (2017) reported impaired RL performance in individuals who had recovered
from AN compared to Healthy Controls (HCs) using a Probabilistic Reversal Learning (PRL)
task, particularly in response to negative feedback. In contrast, Bernardoni et al. (2018)
observed a higher learning rate from punishment among AN patients compared to HCs.
Similarly, Sarrar et al. (2015) found no differences in task performance between individuals
with acute AN and HCs using the Probabilistic Object Reversal Task with neutral stimuli.
Geisler et al. (2018) also found no group differences in a PRL task with neutral stimuli and

monetary feedback.

To shed light on the potential reinforcement RL deficits in AN, researchers have incorporated food-related information into the PRL paradigm. For instance, Zang et al. (2014) demonstrated that individuals with binge-eating disorder exhibited poorer performance when exposed to food-related feedback, indicating a vulnerability to food-related cues. However, attempts to replicate these findings in AN have yielded conflicting results. Hildebrandt et al. (2015) reported increased inflexibility in AN individuals using a PRL task with food-related feedback. In contrast, Hildebrandt et al. (2018) found no differences in PRL performance between AN patients and healthy controls (HCs) when employing the same paradigm.

Given the inconsistent findings in behavioral experiments regarding RL in AN, we propose that these discrepancies may be partly attributed to the predominant use of general stimuli instead of stimuli specifically relevant to the disorder (Schaefer & Steinglass, 2021). Furthermore, when disorder-related information has been incorporated, it has typically been limited to the feedback provided after the participant's choice, with the stimuli presented during the decision-making process unrelated to the disorder. This approach primarily emphasizes the consequences of the choices, neglecting the contextual factors surrounding the decision-making process.

From a theoretical perspective, the methodological choices made in previous studies overlook the critical role of context in learning processes. Contextual learning (Heald, Lengyel, & Wolpert, 2023) draws upon the notion, grounded in the human memory literature, that memory retrieval depends on the match between the conditions during learning and testing. When a mismatch occurs, retrieval is impaired. Applying this concept to RL in AN (Rosas, Todd, & Bouton, 2013), it can be hypothesized that contextual factors, such as individual characteristics, long-term goals, and situational influences, can contribute to impaired RL specifically in decision-making related to the disorder. This impairment can

occur even when RL performance remains intact for decisions unrelated to the disorder (Haynos, Widge, Anderson, & Redish, 2022). In other words, we propose that intermittent impaired RL performance in AN may arise from contextual factors that activate specific learning modes, rather than indicating a fundamental alteration in the underlying learning processes in the brain (for a discussion, see Bernardoni et al., 2021).

To examine the proposed hypothesis, we conducted a study utilizing a modified version of the standard PRL task. In contrast to previous studies that utilized general stimuli (Schaefer & Steinglass, 2021), we incorporated two distinct contexts in our task. One context involved choices between a stimulus related to the disorder (e.g., a caloric food) and a stimulus unrelated to the disorder (e.g., a lamp), while the other context involved choices between two stimuli unrelated to the disorder (i.e., flower vs objects).

Based on the responses to food stimuli often seen in individuals with AN, which
typically exhibit reduced reward, increased aversion, and inhibition (Haynos, Lavender,
Nelson, Crow, & Peterson, 2020), we hypothesized that there would be a more conservative
learning rate for disorder-related choices compared to disorder-unrelated choices within a
PRL task. Additionally, we anticipated a lower learning rate for disorder-related choices in
individuals with AN compared to HCs. Conversely, we expected no learning abnormalities in
individuals with AN for choices unrelated to the disorder.

135 Methods

The study, which adhered to the Declaration of Helsinki, was approved by the
University of Florence's Ethical Committee (Prot. n. 0178082). All eligible participants
provided informed consent and willingly agreed to participate in the study.

## 139 Participants

We tested the hypothesized learning asymmetry in PRL performance across three groups: individuals diagnosed with DSM-5 restricting-type Anorexia Nervosa (R-AN), a

control group comprising of healthy individuals (HCs), and individuals who are at risk of
developing eating disorders (RIs). Demographic and clinical characteristics of the sample are
shown in Table 1. Details about participant selection, inclusion criteria, and sample
characteristics are given in the Supplementary Information (SI).

# 146 Procedure

During the initial session, participants underwent a clinical interview to determine 147 their eligibility for the study. Those who met the criteria and were selected proceeded to 148 anthropometric measurements and completed a battery of psychometric scales. In a 149 subsequent session, participants completed the PRL task. Participants were told they were 150 going to play a simple computer game with the objective of accumulating as many "virtual 151 euro" as possible. During the PRL task, participants were presented with two stimuli 152 simultaneously on a screen and were instructed to select one within a 2.5-second time limit 153 by pressing a key. Trials were presented in an interleaved manner, with a randomly drawn 154 inter-trial interval ranging from 0.5 to 1.5 seconds. Following each trial, a euro coin image 155 was displayed as a reward for correct responses, while a strike-through image of a euro coin 156 served as a punishment for incorrect responses. Feedback was provided for 2 seconds after each trial.

The PRL task consisted of two blocks, each containing 160 trials (see Figure 1). One block included pairs of food-related and food-unrelated images, while the other block exclusively used food-unrelated images. The images were selected randomly from sets of food-related and food-unrelated categories.

All images used in the study were obtained from the International Affective Picture

System (IAPS) database (Lang et al., 2005). The food-related category consisted of images

of french fries, cake, pancake, cheeseburger, and cupcake (IAPS #7461, 7260, 7470, 7451,

7405), while the food-unrelated category included images of a lamp, book, umbrella, basket,

and clothespin (IAPS #7175, 7090, 7150, 7041, 7052). For the control task, five images were used for each of the two food-unrelated categories, i.e., five images of flowers (IAPS #5000, 5001, 5020, 5030, 5202) and five images of objects (IAPS #7010, 7020, 7034, 7056, 7170). (For further details, see the SI).

# 171 Data analysis

To analyze the temporal dynamics of the two-choice decision-making in the PRL task, 172 we employed a hierarchical reinforcement learning drift diffusion model (RLDDM), as 173 described in Pedersen, Frank, and Biele (2017) and Pedersen and Frank (2020). Cognitive 174 modeling analysis allows us to deconstruct decision-making task performance into its 175 component processes. This approach enables the identification of deviations in the 176 underlying mechanisms that may not be evident in the overall task outcome. The RLDDM 177 consists of two key components: one describes how reward feedback is employed to update 178 value expectations [delta learning rule; Rescorla and Wagner (1972)] and the other describes 179 how an agent uses these expectations to arrive at a decision. In the RLDDM the traditional 180 softmax function is replaced by Drift-Diffusion Model [DDM; Ratcliff and McKoon (2008)] 181 which assumes a stochastic accumulation of evidence on each trial (see SI).

The RLDDM has six basic parameters:  $\alpha^+$ ,  $\alpha^-$ , v, a, t, and z. The  $\alpha$  parameters 183 quantify the learning rate in the Rescorla-Wagner delta learning rule (Rescorla & Wagner, 184 1972); a higher learning rate results in rapid adaptation to reward expectations, while a 185 lower learning rate results in slow adaptation. The parameter  $\alpha^+$  is computed from 186 reinforcements, whereas  $\alpha^+$  is computed from punishments. The drift rate v is the average 187 speed of evidence accumulation toward one decision. The decision boundary is the distance 188 between two decision thresholds; an increase of a increases the evidence needed to make a decision. The increase of a leads to a slower but more accurate decision; a decrease in aresults in a faster but error-prone decision. The non-decision time t is the time spent for stimuli encoding or motor execution (i.e., time not used for evidence accumulation). The

starting point parameter z captures a potential initial bias toward one or the other boundary in absence of any stimulus evidence.

To assess the presence of context-dependent learning, we conditioned the model's parameters on two specific contexts: disorder-related choices and disorder-unrelated choices.

This allowed us to examine how the model's parameters varied in response to these different contextual conditions.

199 Results

## Models selection

To evaluate context-dependent learning, we compared several RLDDM models that 201 varied in their conditioning of the model's parameters on the group (R-AN, HC, RI) and 202 context (disorder-related choices and disorder-unrelated choices). We used the Deviance 203 Information Criterion (DIC) to balance model fit and complexity, selecting the model with 204 the lowest DIC as the best trade-off. The following RLDDM models were examined. Model 205 M1: Standard RLDDM without conditioning. DIC = 39879.444. Model M2: Separate 206 learning rates for positive and negative reinforcements. DIC = 39124.890 Model M3: 207 Group-based  $\alpha^+$  and  $\alpha^-$  parameters. DIC = 39194.763. Model M4: Group and 208 context-based  $\alpha^+$  and  $\alpha^-$  parameters. DIC = 38197.467. Model M5: Group and 209 context-based  $\alpha^+$ ,  $\alpha^-$ , and a parameters. DIC = 36427.448. Model M6: Group and 210 context-based  $\alpha^+$ ,  $\alpha^-$ , a, and drift rate (v) parameters. DIC = 36185.146. Model M7: Group 211 and context-based  $\alpha^+$ ,  $\alpha^-$ , a, v, and non-decision time (t) parameters. DIC = 34904.053. 212 Model M8: Group and context-based  $\alpha^+$ ,  $\alpha^-$ , a, v, t, and starting point (z) parameters. DIC 213 = 34917.762. Among the evaluated models, Model M7 had the lowest DIC, indicating the 214 best trade-off between goodness of fit and model complexity. In Model M7, the parameters 215  $\alpha^+$ ,  $\alpha^-$ , a, v, and t (excluding z) were conditioned on both the group and the context.

# Modelling results

Model M7 was estimated using 15,000 iterations, with a burn-in period of 5,000 iterations. Convergence of the Bayesian estimation was evaluated using the Gelman-Rubin statistic. For all parameters in Model M7, the  $\hat{R}$  values were below 1.1 (maximum = 1.025, mean = 1.002), indicating no significant convergence issues. Collinearity and posterior predictive checks were also used to evaluate model validity (see SI).

To investigate the impact of disorder-related versus disorder-unrelated information on RL learning, we compared the posterior estimates of the RLDDM parameters of M7 between the two conditions (see Table 1).

Table 1
Posterior Parameter Estimates of DDMRL Model M7 by Group (R-AN, HC, RI) and Context of PRL Choice (disorder-related vs. disorder-unrelated information). The learning rates  $(\alpha)$  are shown on a logit scale. The probability (p) describes the Bayesian test that the posterior estimate of the parameter in the disorder-related context is greater than the posterior estimate of the parameter in the disorder-unrelated context. Standard deviations are provided in parentheses.

Group	Par.	Neutral choice	Food choice	p	Cohen's $d$
R-AN	a	1.273 (0.039)	1.442 (0.040)	0.0013	0.802
R-AN	V	1.403 (0.320)	1.776 (0.342)	0.7907	0.190
R-AN	t	0.188 (0.011)	0.174 (0.011)	0.8311	-0.253
R-AN	$\alpha^{-}$	1.815 (1.081)	0.738 (1.096)	0.2349	-0.432
R-AN	$\alpha^+$	1.006 (0.899)	-1.786 (0.756)	0.0098	-1.206
НС	a	1.222 (0.033)	1.314 (0.034)	0.0256	0.474
НС	V	$2.157 \ (0.265)$	1.790 (0.263)	0.1606	-0.358
НС	t	0.183 (0.009)	0.172 (0.009)	0.8228	-0.280
НС	$\alpha^{-}$	2.780 (0.874)	3.442 (0.980)	0.6993	0.298
НС	$\alpha^+$	1.198 (0.680)	1.326 (0.700)	0.5544	0.071
RI	a	$1.245 \ (0.041)$	1.316 (0.039)	0.1026	0.403

Group	Par.	Neutral choice	Food choice	p	Cohen's d
RI	V	2.197 (0.322)	1.849 (0.307)	0.2133	-0.381
RI	t	0.188 (0.011)	0.186 (0.011)	0.5462	0.166
RI	$\alpha^{-}$	2.857 (1.067)	2.904 (1.062)	0.5101	0.015
RI	$\alpha^+$	$1.573 \ (0.847)$	$0.739 \ (0.752)$	0.2247	-0.438

Let consider first the evidence of context-dependent learning from within-group 226 comparisons. We found that, on average, individuals in the R-AN group demonstrate a 227 reduced learning rate in response to positive prediction errors (PEs) for disorder-related 228 choices, as compared to disorder-unrelated choices (Cohen's d = 1.206, p = 0.0098). In 229 contrast, no substantial evidence was found indicating a difference in the learning rate 230 between disorder-related and disorder-unrelated choices in the HC (p = 0.5544) and RI (p =231 0.2247) groups. We found no credible difference in the learning rate from negative prediction 232 errors between disorder-related and disorder-unrelated choices for any of the R-AN (p =233 (0.2349), HC (p = 0.6993), and RI (p = 0.5101) groups. Moreover, we found that both the 234 R-AN (Cohen's d = 0.802, p = 0.0013) and HC (Cohen's d = 0.474, p = 0.0256) groups 235 showed a higher decision threshold for disorder-related choices compared to 236 disorder-unrelated choices. 237

Further evidence of context-dependent learning emerges from between-groups comparisons. When making disorder-related choices, individuals with R-AN displayed a decreased learning rate following positive prediction errors (PEs) compared to both HC and RI. Specifically, the learning rate after positive PEs was lower for R-AN compared to HC, p = 0.0009, Cohen's d = 1.498. Similarly, R-AN exhibited a lower learning rate after positive PEs compared to RI (p = 0.0085, Cohen's d = 1.209). In contrast, no credible difference in the learning rate after positive PEs was found between R-AN and HC (p = 0.4325), as well as between R-AN and RI (p = 0.3232), for choices unrelated to disorder information.

Concerning the learning rate after negative PEs, we found that R-AN showed a lower learning rate compared to HC, but only for disorder-related choices: (p = 0.0274, Cohen's d)247 = 1.144). Individuals with R-AN showed a higher decision threshold for disorder-related 248 choices compared to both HC (Cohen's d = 0.622, p = 0.0068) and RI (Cohen's d = 0.454, p = 0.454249 = 0.0118) participants. No credible group differences were found for disorder-unrelated 250 choices. Additionally, we observed that both HC (Cohen's d = 0.520, p = 0.0344) and RI 251 (Cohen's d = 0.529, p = 0.0392) participants exhibited a faster accumulation of evidence and 252 more confident decision-making, as indicated by a higher average drift rate parameter, 253 compared to individuals with R-AN. This difference was only evident for disorder-unrelated 254 choices. Finally, no credible differences were found, for both within-group and between-group 255 comparisons, regarding the non-decision time parameter (t).

#### 257 Preferential choices

To investigate the presence of a bias against food choices in individuals with R-AN 258 during the PRL task, regardless of their past action-outcome history, we analyzed the 259 frequency of food choices in PRL blocks where a food image was paired with a neutral image. 260 Our results show that the AN-R group did not exhibit a bias against the food image, with a 261 proportion of food choices estimated at 0.49, 95% CI [0.46, 0.51]. Furthermore, there were no 262 credible differences in food choices between the R-AN group and the HC group (contrast 263 R-AN - HC = -0.007, 95% CI [-0.037, 0.024]) or between the R-AN group and the RI group 264 (contrast R-AN - RI = 0.013, 95% CI [-0.019, 0.046]). 265

# 266 Comorbidity

We conducted a further statistical analysis to investigate whether the conservative
learning behavior observed in individuals with R-AN could be explained by comorbid
conditions. Using model M7, we categorized individuals with R-AN based on the presence or
absence of diagnosed comorbidities. Our analysis revealed no credible differences in
parameters between the two groups. Specifically, for the disorder-related context, the

parameter differences were as follows:  $\Delta \alpha^- = 2.614$ , 95% CI [-3.173, 8.364];  $\Delta \alpha^+ = -0.635$ , 95% CI [-4.301, 2.449];  $\Delta a = -0.034$ , 95% CI [-0.188, 0.124];  $\Delta v = 0.230$ , 95% CI [-1.203, 1.586];  $\Delta t = 0.002$ , 95% CI [-0.050, 0.055]. For the disorder-unrelated context, the parameter differences were:  $\Delta \alpha^- = -0.768$ , 95% CI [-6.570, 4.401];  $\Delta \alpha^+ = -1.739$ , 95% CI [-6.184, 1.654];  $\Delta a = -0.126$ , 95% CI [-0.281, 0.025];  $\Delta v = 0.744$ , 95% CI [-0.453, 1.886];  $\Delta t = -0.003$ , 95% CI [-0.057, 0.052].

## 278 Discussion

Our findings reveal a context-dependent learning asymmetry in individuals with R-AN specifically in the positive learning rate. This within-group asymmetry is observed when comparing the performance in the PRL task for disorder-related choices versus disorder-unrelated choices. Importantly, no similar difference is found in the two control groups.

The presence of context-dependent learning asymmetry is also supported by
between-group comparisons. Individuals with R-AN exhibited lower learning rates for both
positive and negative prediction errors compared to the HC group, and specifically for
positive prediction errors compared to the RI group, but these differences were observed only
for disorder-related choices. In contrast, no credible differences in learning rates were found
among the three groups for disorder-unrelated choices.

Support for context-dependent learning in R-AN is also provided by the DDM
parameters of the hDDMrl model. Specifically, we observed that the R-AN group exhibited a
higher decision threshold (parameter "a" in the hDDMrl model) compared to the HC and RI
groups, but this difference was only evident in the context of disorder-related choices. This
suggests that individuals with R-AN displayed a more cautious or conservative
decision-making behavior specifically in relation to disorder-related choices (see also Caudek,
Sica, Cerea, Colpizzi, & Stendardi, 2021; Schiff, Testa, Rusconi, Angeli, & Mapelli, 2021).

Further support of context-related learning in R-AN comes from the result which 297 indicate that both healthy control (HC) and at-risk (RI) participants exhibited a faster 298 accumulation of evidence and displayed more confident decision-making, as reflected by a 299 higher average drift rate parameter, compared to individuals with restrictive anorexia 300 nervosa (R-AN). However, this difference was specifically observed for disorder-unrelated 301 choices. It is noteworthy that individuals with R-AN displayed slower evidence accumulation 302 and less confident decision-making specifically in disorder-unrelated contexts, whereas this 303 group difference was not observed for disorder-related choices. This finding further supports 304 the notion of context-dependent learning in individuals with R-AN, particularly in the 305 context of food-related information.

Further evidence of context-related learning in R-AN comes from the analysis of the 307 drift rate parameter. Individuals with R-AN exhibited slower evidence accumulation and less 308 confident decision-making compared to the control groups, specifically in the context of 309 disorder-unrelated choices. Conversely, no credible group differences were observed for 310 food-related choices. These results suggest that individuals with R-AN may allocate greater 311 cognitive resources to process salient information in the disorder-related context, which leads 312 to similar evidence accumulation rates in decision-making compared to the control groups. 313 In contrast, they exhibit a slower evidence accumulation rate when faced with less salient 314 disorder-unrelated choices. 315

The analysis of preferential choices supports the conclusion that the learning
performance asymmetry observed in individuals with R-AN is not due to a preferential
selection of the disorder-unrelated image during the learning task. Additionally, our analysis
examining the relationship between the model's parameters and the presence of
comorbidities indicates that the learning performance asymmetry in individuals with R-AN
cannot be attributed to comorbid conditions.

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## General discussion

In this study, we investigated reinforcement learning using a behavioral paradigm that included two distinct learning contexts: one involving choices related to food and the other involving choices unrelated to food. We compared the performance of patients with R-AN to age-, gender-, and education-matched healthy controls, as well as healthy controls at-risk of developing eating disorders. Consistent with our hypotheses, our findings revealed a lower learning rate in the disorder-related context for individuals with R-AN, whereas both healthy participants and at-risk individuals learned equally well in both contexts.

In PRL tasks, a participant's performance can be influenced by two potential factors. 330 First, there may be a learning impairment, where participants struggle to accurately update 331 the value of the stimuli. Second, there may be a decision impairment, where participants 332 may still select the wrong stimulus despite having intact learning processes. Our results show 333 that individuals with R-AN may struggle with both accurately updating the value of 334 disorder-related stimuli and making appropriate decisions based on this information. 335 However, we did not observe similar impairments in decision making for disorder-unrelated 336 choices. These findings provide evidence for context-dependent learning in individuals with 337 R-AN, where the inclusion of disorder-related information negatively impacts their RL 338 performance. It is important to note that this effect is specific to the disorder-related context 339 and does not suggest a generalized RL deficit in individuals with R-AN. Thus, our results 340 challenge the notion of a domain-general RL mechanism impairment in this population (see 341 Bernardoni et al., 2021). 342

Previous studies have demonstrated that reward and punishment processing in individuals with AN is influenced by stimulus properties and contextual factors. For instance, predictable and controllable behaviors such as calorie counting or purging are often perceived as rewarding, providing individuals with a sense of control and accomplishment. Conversely, unpredictable and uncontrollable situations, such as social outcomes, can be perceived as punishing, leading to heightened anxiety and distress (Haynos et al., 2020).
While previous studies have predominantly examined the impact of context on the subjective value attributed to experiences in AN, our study expands on this research by demonstrating that context plays a crucial role in the actual learning process itself (Heald et al., 2023).
This goes beyond solely influencing subjective value and provides valuable insights into how reward and punishment processing operates in AN.

Other recent studies have focused on investigating context-specific learning in eating 354 disorders. One task specifically designed for this purpose is the two-step Markov decision 355 task, which distinguishes between automatic or habitual (model-free) learning and controlled 356 or goal-directed (model-based) learning. For instance, studies conducted by Foerde et al. 357 (2021) and Onysk and Seriès (2022) employed similar experiments using the two-step task 358 paradigm. Foerde et al. (2021) compared a monetary two-step task and a food-related 359 two-step task, while Onysk and Seriès (2022) utilized stimuli unrelated to food or body 360 images (i.e., pirate ships and treasure chests) with rewards associated with body image 361 dissatisfaction. The results of these studies consistently demonstrated that individuals with 362 AN tend to exhibit a stronger inclination towards habitual control over goal-directed control 363 across different domains compared to healthy controls. However, no significant differences 364 were observed in learning rates as a function of context, nor between AN patients and 365 healthy controls, according to these findings. In contrast, the present study reveals that, in individuals with R-AN, the learning process per se can be influenced by contextual (disorder-related) information, even when such information is not directly relevant to the task outcome.

The hypothesis proposing that reinforcement learning (RL) anomalies in individuals
with anorexia nervosa (AN) may be influenced by contextual factors carries significant
implications for treatment strategies. Currently, Cognitive Remediation Therapy (CRT) is
utilized to address cognitive inflexibility in AN and other eating disorders. CRT involves

cognitive exercises and behavioral interventions aimed at improving central coherence 374 abilities, reducing cognitive and behavioral inflexibility, and enhancing thinking style 375 comprehension (Tchanturia, Davies, Reeder, & Wykes, 2010). A key aspect of CRT is to 376 avoid addressing symptom-related themes and instead utilize neutral stimuli in cognitive and 377 behavioral exercises. This approach aims to establish a therapeutic alliance and reduce 378 drop-out rates, particularly among individuals with AN. However, recent evidence suggests 379 that CRT may not consistently improve central coherence abilities, cognitive flexibility, or 380 symptoms associated with eating disorders (Hagan, Christensen, & Forbush, 2020; 381 Tchanturia, Giombini, Leppanen, & Kinnaird, 2017). In response to these findings, Trapp et 382 al. (2022) propose modifications to address practical challenges encountered in the 383 application of CRT. They question the use of neutral stimuli and draw support from Beck's 384 cognitive theory of depression (Beck & Alford, 2009). This proposition aligns with the hypothesis of our study. If further studies consistently demonstrate that maladaptive RL is context-dependent, it would necessitate a shift in intervention approaches.

There are few important limitations and questions for future research. 1) One aspect 388 to consider is the use of symbolic rewards and punishments in our study, represented by 389 images of a one euro coin and a barred representation of a one euro coin, respectively. These 390 rewards and punishments were merely symbolic, and it is unclear how the use of concrete, 391 non-symbolic rewards and punishments would impact the findings. Additionally, the 392 subjective value of one euro, or the loss of one euro, may vary among participants. Therefore, 393 future studies could aim to determine the equivalence of subjective values for rewards and punishments to enhance the understanding of the underlying processes. 2) Our study only included individuals with R-AN who were not in the most severe stage of the illness, as they were recruited from a center for voluntary medical and psychological support. We did not 397 examine R-AN patients who require hospitalization due to the life-threatening nature of 398 their illness. It is possible that at the later stages of the illness, associative learning abilities, 399 which were preserved in the present sample under neutral conditions, may become impaired.

Therefore, investigating the impact of illness severity on context-dependent learning in R-AN 401 patients is an important avenue for future research. 3) While we observed no difference in 402 the choice behavior of R-AN patients, as measured by the relative frequency of image choices, 403 when selecting between a neutral image and a food image, we did find a slower learning rate 404 and lower decision threshold for R-AN patients compared to healthy controls in the RLDDM 405 model when compared to choosing between two neutral images. It is possible that the higher 406 "salience" of food images compared to neutral images could be better captured by other 407 measures, such as fixation length or the number of fixations, rather than solely relying on the 408 relative frequency of image choices. This warrants further exploration in future studies. 4) It 409 is worth noting that our study excluded women under the age of 18. However, this age range 410 is a critical period as the onset of AN during this stage may have a more profound impact on 411 associative learning, given the ongoing cognitive development and less-developed protective factors. Therefore, future studies should take into consideration the inclusion of participants 413 in this age range to better understand the influence of context-dependent learning in R-AN. References

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