Archival Report

Daily Social Isolation Maps Onto Distinctive Features of Anhedonic Behavior: A Combined Ecological and Computational Investigation

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ABSTRACT

BACKGROUND: Loneliness and social isolation have detrimental consequences for mental health and act as vulnerability factors for the development of depressive symptoms, such as anhedonia. The mitigation strategies used to contain COVID-19, such as social distancing and lockdowns, allowed us to investigate putative associations between daily objective and perceived social isolation and anhedonic-like behavior.

METHODS: Reward-related functioning was objectively assessed using the Probabilistic Reward Task. A total of 114 unselected healthy individuals (71% female) underwent both a laboratory and an ecological momentary assessment. Computational modeling was applied to performance on the Probabilistic Reward Task to disentangle reward sensitivity and learning rate.

RESULTS: Findings revealed that objective, but not subjective, daily social interactions were associated with motivational behavior. Specifically, higher social isolation (less time spent with others) was associated with higher responsivity to rewarding stimuli and a reduced influence of a given reward on successive behavioral choices.

CONCLUSIONS: Overall, the current results broaden our knowledge of the potential pathways that link (COVID-19-related) social isolation to altered motivational functioning.

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The consequences of social isolation for symptoms and maladaptive behaviors have been widely investigated in clinical and preclinical studies. For example, it is well established that chronic social isolation elicits anhedonia (operationalized as a reduction in sucrose intake and sucrose preference) in rodents (1-3). Converging evidence from human investigations points to the bidimensionality of the construct: on the one hand, social isolation (or social disconnectedness) has been defined in terms of a restricted network of relationships and objective lack of interaction with others or participation in social activities (4,5). On the other hand, emotional isolation (5), also called perceived isolation (6), has been conceptualized as the subjective perception of lack of companionship, intimacy, and support and seems to be more strictly associated with feelings of loneliness, a sense of dissatisfaction with one's connections, and qualitative, rather than quantitative, social deficiencies (7). Importantly, studies of humans suggest that both dimensions of social isolation and perceived isolation predict anhedonic symptoms, depression, anxiety (8,9), poorer mental health (10), and morbidity more generally (11). Conversely, a higher level of social support is a protective factor for anhedonia (12), suicide, and suicide risk (13) and is associated with a better quality of life and reduced functional decline in individuals with depression (14).

Most longitudinal studies that have attempted to shed light on causal relationships between loneliness and depressive

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symptoms have pointed to the perception of social isolation, loss of significant others, and a poor interpersonal network as risk factors for the development of depression (15–17), while others have described reciprocal influences between the 2 constructs (8,18).

The reviewed findings are consistent with the assumption of the social baseline theory, according to which the social network is the primary source of safety (19) and of the social safety theory, which further hypothesizes that social isolation and rejection upregulate inflammatory activity mainly via bidirectional neural-immune communication, ultimately increasing all-cause mortality (20,21).

The current study was conducted during a major period of social deprivation amid strict COVID-19 measures. Even following the conclusion of national lockdowns, Italy (including the area in which the current study was performed) continued to face prolonged restrictions at the regional level until early 2022, namely constraints on the number of individuals permitted in public or private spaces, implementation of curfews, and the transition to online learning and remote work. This situation afforded us the opportunity to investigate the effects of varying degrees of loneliness and isolation in a naturalistic setting.

The aim of the current study was to investigate whether (objective or perceived) social isolation would be associated

with anhedonic symptoms in an unselected healthy sample. Importantly, an ecological momentary assessment (EMA) was implemented to provide a daily estimation of the number and duration of (i.e., objective social isolation) as well as the desire and need for (i.e., perceived isolation) social interactions. Moreover, assessment of anhedonic symptoms did not rely on self-reports but instead was evaluated based on participants' performance on the Probabilistic Reward Task (PRT) (22), in which anhedonic behavior is operationalized as the ability to modify behavior as a function of rewards. Overall, we hypothesized positive associations between objective and subjective measures of social isolation and anhedonic behavior. Although it has been historically conceptualized as loss of pleasure (23), anhedonia is a complex and multifaceted symptom that may arise from impairments in several aspects of reward processing, including blunted reward learning or reward sensitivity (24,25). To parse the specific associations of each of these components with social isolation, computational modeling of trial-level performance on the PRT was implemented (26). Given the absence of previous investigations of this issue, we did not have a specific hypothesis about which of the 2 components would be more affected by social isolation

Numerous studies have explored the relationship between objective or perceived social isolation and their associations with the multifaceted construct of anhedonia. However, the current study stands out by aiming to investigate this relationship not only through ecological assessment but also through a behavioral task. By dissecting behavioral performance into 2 components, we aimed at a closer understanding of which aspects of motivational functioning are more involved in this relationship. This approach holds promise to provide a more nuanced understanding of the mechanisms that underlie the interplay between social isolation and anhedonia.

METHODS AND MATERIALS

Participants

This study is part of a larger research project (27), although the sample is not overlapping. Participants were recruited from among university students and the general population from May 2021 to November 2021. The protocol was approved by the local institutional review board (Protocol No. 1170/2021). The final sample included 114 participants (81 female, mean age 22.24 \pm 2.93 years, range 20–30 years) (see Supplemental Section S1 for exclusionary criteria).

Procedure

After signing the informed consent form, participants were asked to complete a set of online questions assessing socio-demographic, lifestyle, and medical information (e.g., age, weight, nicotine and alcohol consumption, physical activity) and a validated questionnaire to evaluate self-reported loneliness. Next, a laboratory visit was scheduled during which participants were asked to perform the PRT. Participants then received a full explanation of the EMA procedure and were asked to keep electronic diaries on their smartphones for 4 consecutive days, after which they were debriefed and compensated with the money that they won during the PRT.

Assessment of Perceived Social Isolation

The Italian version of the Revised UCLA Loneliness Scale (28,29) was administered; it is a 20-item questionnaire that assesses the subjective experience of loneliness and subjective feelings of social isolation (detailed in Supplemental Section S2). Internal consistency in the current study was $\alpha = 0.867$. Questions regarding the specific COVID-19 situation were also administered (see Supplemental Section S3 for the full list).

Probabilistic Reward Task

The PRT (22) is a well-validated signal detection task recommended by the Research Domain Criteria (30) as a probe of participants' ability to modify their behavior as a function of reward, thus providing an objective measure of reward responsiveness and reward learning (detailed in Supplemental Section S4 and Figure S1) (31).

Performance was analyzed with respect to response bias (log b), an empirically derived measure of systematic preference to choose the more frequently rewarded stimulus (rich). ΔResponse bias was computed as the systematic preference to identify the more frequently rewarded stimulus (response bias) during block 3 minus response bias during block 1 (22). To evaluate overall task performance, we also examined discriminability, which is participants' ability to distinguish between the 2 stimuli reflecting task difficulty, as well as accuracy and reaction times (see Supplemental Section S4 for formulas and secondary analyses).

Computation Modeling of the PRT. To parse specific components of reward processing, the participants' trial-level PRT data were fitted with 4 reinforcement learning models based on previously established methods (detailed in Supplemental Section S5) (26). Group priors in these models were computed via expectation maximization (32), and subject-specific parameters were estimated through Laplace approximation of the posterior distributions. Integrated grouplevel Bayesian information criterion factors were used to compare the model fits. The most parsimonious account of the data was provided by the action model, with a group-level log Bayes factor compared with the second-best belief model of 68 (which represents very strong evidence). This approach allowed for the derivation of 2 main parameters: 1) reward sensitivity, with higher scores denoting that a participant is more sensitive to rewards (mean = 2.02, SD = 0.35), and 2) learning rate (mean = -5.35, SD = 1.9), which measured the ability to learn from reward feedback, with higher scores indicating a stronger effect of reward on successive behavior.

In accordance with Huys et al., parameters were computed in the transformed space to prevent issues with non-Gaussianity (see Supplemental Section S6 for details) (26).

Ecological Momentary Assessment

The EMA was the last step of the research protocol and involved repeated sampling of individuals' current experiences in real time and in natural environments with the purpose of minimizing recall bias and maximizing ecological validity (33).

Participants received preprogrammed emails containing a link to their electronic diary on Qualtrics.com. These emails

were sent randomly within a 2-hour time window from the first Wednesday after the laboratory session until the following Saturday. On each of the 4 days, participants received 6 surveys, which were randomly distributed throughout their selfreported waking period and began at least 30 minutes after they woke up. The EMA questions were only available for 25 minutes after the initial notification. Answers were provided on a visual analog scale, and each diary took 1 to 2 minutes to be filled out. Based on previous studies (34), ad hoc questions were used to assess the quantity ("Since the last electronic diary, did you have any social interaction?"), duration ("How long did this interaction last?"), subjective perception ("How do you rate the valence of such interaction?"), and desire or need for social interactions ("Since the last electronic diary, have you missed interacting with people?" or "How hard have you desired to interact with someone?"). Participants had to choose "yes" or "no" for the question about the presence of interaction while the other answers were provided on a visual analog scale ranging from 1) 0 = "completely negative" to 10 = "completely positive" for valence; 2) 1 minute to 120 minutes or more for duration; and 3) 0 = "not at all" to 10 = "very much" for desire or need.

Data Analysis

First, assumptions of normality, linearity, homogeneity of variances, and sphericity were checked. Then, the influence of potential confounders, such as age, sex, alcohol, and smoking, on the main variables of interest was assessed using correlational and *t* test analyses. The variables that yielded significant results in this preliminary check were included as covariates in all subsequent analyses.

Regarding PRT performance, 4 separate general linear models were performed. First, the general linear models were computed for response bias and discriminability as outcomes and block (block 1, block 2, block 3) as the within-participant variable. For accuracy and reaction time, stimulus (lean vs. rich) was included as an additional factor. Behavioral results regarding performance on the PRT have been reported elsewhere (27) and will not be detailed here (see Figure S1 for a graphical representation).

For the EMA component, a multilevel model was carried out with Mplus version 8.7 software (35) using robust maximum likelihood estimators with a full information maximum likelihood approach under the missing-at-random assumptions to handle missing data. Participants with <30% of valid assessments on the EMA measures were not retained for this analysis as they were in previous EMA studies in the context of anhedonia and motivational behavior (36,37).

The models were hypothesis driven and informed by the previously reviewed rodent and human findings on the detrimental effects of social isolation on anhedonic symptoms. A mediational chain was specified at the between-participants level: trait loneliness predicting valence of social interaction, which in turn predicted the duration of daily social interaction, which in turn predicted Δ response bias (model 1) and the 2 components of the PRT (model 2). Duration of daily social interaction was also partialled out for trait loneliness, and trait loneliness was specified as being associated with the 2 components of the PRT. In model 2, the computational-based

components of the PRT (i.e., reward sensitivity and learning rate) were treated as separate dependent between-participants variables within the overall nomological network (see Supplemental Section S8 for details on the models).

RESULTS

No correlations or differences emerged in the main variables of the study with respect to age, sex, or nicotine and alcohol consumption, with the exception of Δ response bias, which was significantly higher in nonsmokers than smokers ($t_{112} = 2.06$, p = .02). Thus, general linear models were performed controlling only for smoking status also given previous findings of significant effects of nicotine craving on response bias (38).

The sample included 5 participants who lived alone, 7 who lived with a partner, 23 who lived with roommate(s), and 79 who lived with their family. Regarding consequences of COVID-19, 40 participants reported that they had undergone quarantine measures during the 6 months prior to the start of the study. Most participants (n = 81) highlighted a negative influence of COVID-19 restrictions, and 67 participants reported perceived social isolation. The average Revised UCLA Loneliness Scale score was 38.34 ± 8.37 (normative scores are 37.06 ± 10.91 for men and 36.06 ± 10.11 for women) (28).

In general, after excluding 1.76 of outlier trials, an increase in response bias from the first block to the following blocks emerged together with a concomitant increase in accuracy and a decrease in reaction time for rich (vs. lean) trials from block 1 to block 2 and 3. As expected, no effects of discriminability emerged. Collectively, these patterns confirm that the PRT elicited the intended behavioral effects. Consistent with previous accounts (26), reward sensitivity and learning rate were inversely associated (r = -0.59, p < .001) (see Table 1, Figure 1, and Supplemental Section S7 for outliers).

EMA Models

Eight participants reported a very large proportion of missing data points on EMA variables (>70% of the total), and consequently, they were excluded from the analyses. Thus, the final sample for EMA models comprised 106 individuals, who did not differ from those excluded from the analyses on any sociodemographic or between-person variables. The final average proportion of valid data points for the EMA measures was 59.34% (SD = 15.04), with an average number of data entries (16.14, SD = 3.93, range: 9-24) that was comparable to that obtained in previous studies that have examined social interactions (39,40).

Descriptive statistics and scatterplots related to the EMA models are summarized in Table 1 and Figure 1. Model 1 failed to show significant prediction of Δresponse bias by valence or duration of daily social interactions (detailed in Figure S2).

Estimates from the empirical EMA model 2 are reported in Figure 2. The model showed substantial fit to the data: Satorra-Bentler $\chi^2_2 = 0.88$, p = .88. The unconditional intraclass correlation coefficients associated with the valence of social interactions and duration of daily social interactions largely varied across EMA occasions (approximately 90%), but a nonignorable component of their variability was associated with stable individual differences. In other words, scores on these variables varied to a consistent degree between

Table 1. Descriptive Statistics and Zero-Order Correlations for the Ecological Momentary Assessment Model Variables (n = 106)

		Descriptive Statistics			Correlations					
		Mean (SD)	Skewness	Kurtosis	1	2	3	4	5	6
1	ΔResponse Bias	0.09 (0.28)	0.33	0.13	_	0.20ª	0.30ª	0.00	-0.02	-0.05
2	Learning Rate	-5.35 (1.9)	1.06	0.06	_	_	-0.59 ^b	0.12 ^c	0.22ª	0.02
3	Reward Sensitivity	2.02 (0.35)	-1.57	4.73	_	_	_	-0.15 ^c	-0.27ª	-0.04
4	Valence	5.81 (3.59)	-0.53	-1.08	_	_	_	_	0.47 ^b	-0.36 ^b
5	Duration	4.31 (3.77)	0.35	-1.40	_	_	_	0.64 ^b	_	-0.15
6	Loneliness	34.47 (0.81)	0.80	0.66	_	_	_	_	_	_

ΔResponse bias indicates response bias during block 3 minus response bias during block 1; learning rate (in transformed space) indicates the ability to learn from reward feedback; reward sensitivity (in transformed space) indicates the internal worth of an external reward; valence indicates the valence of social interaction; duration indicates the duration of social interaction (objective measure of social isolation); loneliness indicates self-reported levels of loneliness assessed by the Revised UCLA Loneliness Scale. Mean, SD, skewness, and kurtosis refer to the between-participants level (for the ecologic momentary assessment variables, these are calculated across all available measurement occasions). Correlations below the diagonal pertain to the within-participants level, and those above the diagonal pertain to the between-participants level.

assessments, although approximately 10% of their overall variability was explained by enduring individual characteristics. Trait loneliness significantly explained the stable component of the valence of social interactions stemming from EMA assessment (standardized $\beta=-0.36,\,p<.001)$, which in turn significantly explained the stable component of the duration of daily social interactions (standardized $\beta=0.48,\,p<.001)$. Most importantly, the stable component of the duration of daily social interactions significantly explained both computational-based PRT parameters, exerting a negative effect on reward sensitivity (standardized $\beta=-0.28,\,p<.01)$ and a positive effect on learning rate (standardized $\beta=0.22,\,p<.05)$. Overall, the nomological network explained 8% of the total variance in reward sensitivity and 5% of the total variance in the learning rate scores.

DISCUSSION

Combining a laboratory session and an EMA, the current study evaluated whether daily perceived and objective social isolation were associated with anhedonic behavior assessed by performance on the PRT, which provides an objective measure of the ability to adapt behavior as a function of reinforcement history.

The hypothesis of an association between social isolation and dysregulation in motivated behavior was driven by existing evidence of positive cross-sectional and longitudinal associations between social isolation and anhedonic symptoms in human and animal studies (8,38) and of perceived social support as a protective factor for anhedonia and depressive symptoms in general (12,41).

Unexpectedly, momentary levels of perceived and objective social isolation, assessed over 4 consecutive days, were not significantly related to changes in response bias, which is the systematic preference in identifying the more frequently rewarded stimulus across blocks. However, the application of a trial-by-trial computational modeling analysis to PRT choice data allowed us to distinguish 2 critical parameters that underlie such systematic preference, namely reward sensitivity

and learning rate. Previous research that attempted to disentangle single critical parameters in the construct of anhedonia described reward sensitivity as the immediate hedonic impact of rewards (or the behavioral equivalent of consummatory pleasure) and learning rate as the participant's ability to learn from reward feedback, which is shaped by prediction error mechanisms (26).

In the current study, the daily duration of social interactions was negatively associated with reward sensitivity: thus, lesser time spent interacting with others (i.e., higher social isolation) was unexpectedly associated with higher responsivity to rewarding stimuli. Conversely, the duration of daily social interactions was a positive predictor of learning rate, meaning that higher social isolation was associated with an impaired ability to learn from reward feedback, specifically a reduced influence of a given reward on successive behavioral choice.

This conflicting pattern of results is not new in the scientific literature on the effects of social isolation. Behaviorally, the increased reward responsiveness that we found in association with high social isolation resembles the effects of the construct of social craving, which was recently developed by Tomova et al. to describe the urge and desire for social interactions that seems to develop following forced isolation in humans and animals (42,43). Conversely, the association of less frequent social interactions with lower reward learning may suggest an impaired ability to learn from reward feedback that may reflect a cognitive adaptation to reduced social stimuli. This could indicate a diminished reinforcement learning capability, possibly due to a generalized decrease in motivation or engagement stemming from prolonged social isolation. This closely resembles what could be termed social despair. Although not explicitly defined as such, this pattern has been observed in both clinical (44) and preclinical studies, particularly during extended or chronic periods of isolation (45). Chronic isolates appear to exhibit a series of profound interpersonal deficits, involving cognitive, affective, and behavioral aspects, linked to reduced motivation and impaired reward learning. In rodents, this is reflected in increased immobility and reduced swimming and climbing behaviors, diminished

 $^{^{}a}p < .01.$

 $^{^{}b}p < .001.$

 $^{^{}c}p < .05.$

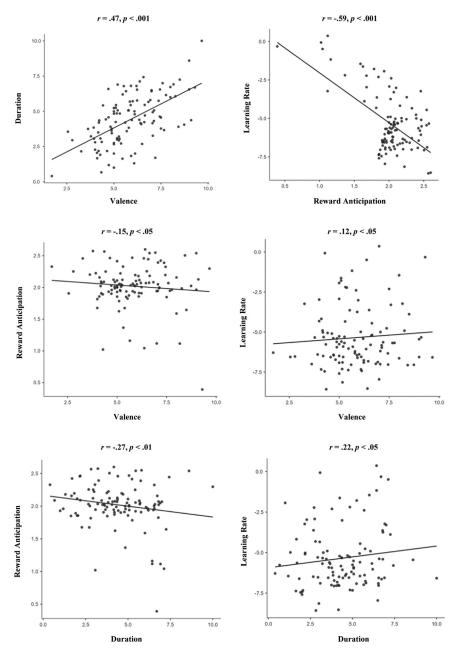


Figure 1. Scatterplots of the between-level associations between learning rate (in transformed space) and reward sensitivity (in transformed space) and momentary ecological assessment of valence and duration of daily social interaction.

sucrose preference, and impaired spatial memory and learning (46–48). Compared with transiently lonely humans, chronically lonely individuals make more self-derogatory internal and stable attributions, seem to prefer passive coping strategies, expect social failure, and fail to seek solutions (49,50).

The opposite associations of social isolation with rewarding sensitivity and learning rate is not surprising if the different neurobiological mechanisms that underpin each parameter are considered. Previous investigations primarily related reward sensitivity and consummatory pleasure to the opioidergic system; specifically, the hedonic impact and liking facet of

motivation have been linked to μ opioid signaling in regions such as the shell of the nucleus accumbens (51). On the other hand, preclinical and clinical studies have shown that reward prediction error is primarily tracked by phasic firing of dopaminergic neurons in the midbrain (52–54). Coherently, the application of the computational model on PRT choice data by Pizzagalli *et al.* showed that administration of a low dose of pramipexole (a dopaminergic agonist acting as an antagonist at low doses) reduced learning rate but not sensitivity (27,55). Thus, it is plausible that the obtained opposite associations between social isolation and the 2 computational factors map

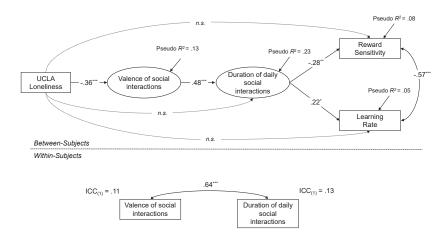


Figure 2. Completely standardized estimates from ecological momentary assessment (EMA) model 2. Dotted lines represent statistically nonsignificant paths. One-headed arrows represent direct effects, while double-headed arrows represent covariances. Squares indicated observed variables, while circles represent latent (between participants-level) components of EMA measures. *p < .05, **p < .01, ***p < .001. CFI, comparative fit index; ICC, intraclass correlation coefficient; RMSEA, root mean square error of approximation; SB, Satorra-Bentler; SRMR, standardized root mean square residual; TLI, Tucker-Lewis index.

onto different and partially dissociable neurobiological

 $SB\chi^{2}_{(df=2)} = .88 \;, p = .88 \;, RMSEA = 0; CFI = 1.00; TLI = 1.00; SRMR_{within} = .000; SRMR_{hatronom} = .014 \;, SRMR_{hatronom} = .014 \;,$

mechanisms.

A large body of clinical and preclinical literature points to a role of dopaminergic and opioidergic neuromodulation in regulating social interactions and bonding formation, as well as in social deprivation or social distress (45,56,57). According to preclinical evidence, the endogenous opioid system not only regulates physical pain but also social distress and plays a role in the attribution of value to social interactions (58,59). Coherently, a positron emission tomography study conducted with healthy humans found increased activation in the μ opioid receptor system in the ventral striatum, midline thalamus, amygdala, and periaqueductal gray in individuals exposed to social rejections (60). When comparing individuals with a diagnosis of major depressive disorder to healthy control individuals, however, the same authors found reduced endogenous opioid release during social rejection in brain regions implicated in stress and reward processing (61).

Social isolation, on the other hand, appears to be more specifically related to dopaminergic transmission; in rodents, brief periods of social isolation enhance motivation to seek contact and social interactions with conspecifics, a behavior underpinned by an increase in excitatory inputs to midbrain dopaminergic neurons of the dorsal raphe (42). Again, when chronic isolation is considered, opposite patterns of depressive-like behaviors (increased immobility and despair and reduced sucrose preference indicating impaired reward sensitivity and anhedonia) have been described (48). Consistently, encouraging results linking loneliness (perceived isolation) to weaker activation in reward-related brain regions during exposure to socially rewarding stimuli (vs. objects) has emerged in humans (62). However, these results were not replicated by D'Agostino et al. in a similar functional magnetic resonance imaging paradigm (63). To summarize, social isolation and social distress may be modulated by the dopaminergic and opioidergic systems, which in turn may have a preferential impact on reward learning and sensitivity, respectively.

Importantly, most of the reviewed clinical literature concerns loneliness and thus the subjective feeling of isolation involving

concepts of the quality of and satisfaction with interactions rather than objective social disconnectedness and lack of relationships per se. According to a large body of research, loneliness represents a greater threat to mental health than physical isolation and seems to be implicated in depression, suicide, alcohol and substance use, poor sleep habits, and dementia (8,9). This is notable considering the current lack of association between self-reported levels of loneliness as assessed by the UCLA Loneliness Scale and reward learning or sensitivity. However, it should be noted that scores on the UCLA Loneliness Scale were negatively associated with the valence of social interaction but not with its daily duration, which confirms the potential of this scale to primarily assess perceived rather than objective social deprivation. Although the literature linking loneliness assessed by the UCLA Loneliness Scale and momentary assessment of social isolation is not extensive, an increasing interest in the issue has evolved, especially during periods of lockdowns and restrictions of public life. As expected, such studies have positively linked daily loneliness to negative subjective states (64) but have failed to find statistically significant associations between quantity and duration of social conversations and scores on the UCLA Loneliness Scale (65).

The unexpected current lack of association between perceived isolation (scores on the UCLA Loneliness Scale) and the components of reward sensitivity and learning may be well explained by the fact that the study was conducted during times when pandemic-related social distancing protocols were in effect. When considering both objective and subjective social isolation, the determinants of objective isolation (living alone, having a small social network, etc.) are considered to be among the most powerful predictors of loneliness (66,67). It is plausible that when objective social disconnectedness reaches an extreme, as in the situation of the severe social restrictions imposed worldwide in the past few years, its effects outweigh the effects of subjective perception of isolation.

The main limitation of the current research is its correlational design, which does not allow for conclusions about causal relationships between the investigated variables (full limitations

are detailed in Supplemental Section S9). Although far from unraveling the complex relationships between social isolation and the multifaceted construct of anhedonia, the current study provides preliminary insights into impaired motivational functioning in circumstances of higher daily objective social isolation. Importantly, the use of computational modeling allowed us to add a level of complexity to these well-known associations. Given the involvement of different neurobiological pathways in learning rate and reward sensitivity, replication of the current results may provide meaningful insights for pharmacological and psychotherapeutic early interventions. Because anhedonia is one of the main detrimental mental health effects of the COVID-19 outbreak and its related lifestyle changes and social restrictions (68-71), understanding the pathways through which forced social isolation influences the specific components of reward learning and sensitivity is important. It is possible that different durations of social isolation distinctively influence each component of reward processing. For example, we speculate that while reward learning deteriorates progressively in response to longer periods of deprivation, sensitivity to reward is enhanced as a coping strategy or compensatory mechanism. Future studies may investigate this hypothesis in humans by administering the PRT during acute mandatory isolation (72) and/or in chronically isolated individuals. In the future, researchers could build on current findings by including a larger and more diverse group of participants in terms of age, sex, and ethnicity to examine whether these variables moderate the observed outcomes.

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REFERENCES

- Grippo AJ, Lamb DG, Carter CS, Porges SW (2007): Social isolation disrupts autonomic regulation of the heart and influences negative affective behaviors. Biol Psychiatry 62:1162–1170.
- Wallace DL, Han M-H, Graham DL, Green TA, Vialou V, Iñiguez SD, et al. (2009): CREB regulation of nucleus accumbens excitability mediates social isolation-induced behavioral deficits. Nat Neurosci 12:200–209.
- Acero-Castillo MC, Ardila-Figueroa MC, Botelho De Oliveira S (2021): Anhedonic type behavior and anxiety profile of Wistar-UIS rats subjected to chronic social isolation. Front Behav Neurosci 15:663761.
- De Jong Gierveld J, Hagestad GO (2006): Perspectives on the integration of older men and women. Res Aging 28:627–637.
- Weiss RS (1974): Loneliness: The Experience of Emotional and Social Isolation. Cambridge, MA: MIT Press.
- Cornwell EY, Waite LJ (2009): Social disconnectedness, perceived isolation, and health among older adults. J Health Soc Behav 50:31–48.
- Pinquart M, Sorensen S (2001): Influences on loneliness in older adults: A meta-analysis. Basic Appl Soc Psych 23:245–266.
- Cacioppo JT, Hughes ME, Waite LJ, Hawkley LC, Thisted RA (2006): Loneliness as a specific risk factor for depressive symptoms: Crosssectional and longitudinal analyses. Psychol Aging 21:140–151.
- Hagerty BM, Williams RA (1999): The effects of sense of belonging, social support, conflict, and loneliness on depression. Nurs Res 48:215–219
- Slavich GM, O'Donovan A, Epel ES, Kemeny ME (2010): Black sheep get the blues: A psychobiological model of social rejection and depression. Neurosci Biobehav Rev 35:39–45.
- Cacioppo JT, Hawkley LC (2009): Perceived social isolation and cognition. Trends Cogn Sci 13:447–454.
- Grey I, Arora T, Thomas J, Saneh A, Tohme P, Abi-Habib R (2020): The role of perceived social support on depression and sleep during the COVID-19 pandemic. Psychiatry Res 293:113452.
- Motillon-Toudic C, Walter M, Séguin M, Carrier J-D, Berrouiguet S, Lemey C (2022): Social isolation and suicide risk: Literature review and perspectives. Eur Psychiatry 65:e65.
- Rytsälä HJ, Melartin TK, Leskelä US, Sokero TP, Lestelä-Mielonen PS, Isometsä ET (2007): Predictors of long-term work disability in Major Depressive Disorder: A prospective study. Acta Psychiatr Scand 115:206–213.
- Heikkinen R-L, Kauppinen M (2004): Depressive symptoms in late life: A 10-year follow-up. Arch Gerontol Geriatr 38:239–250.
- Loades ME, Chatburn E, Higson-Sweeney N, Reynolds S, Shafran R, Brigden A, et al. (2020): Rapid systematic review: The impact of social isolation and loneliness on the mental health of children and adolescents in the context of COVID-19. J Am Acad Child Adolesc Psychiatry 59:1218–1239.e3.
- Simbi CMC, Zhang Y, Wang Z (2020): Early parental loss in childhood and depression in adults: A systematic review and meta-analysis of case-controlled studies. J Affect Disord 260:272–280.
- Luo Y, Hawkley LC, Waite LJ, Cacioppo JT (2012): Loneliness, health, and mortality in old age: A national longitudinal study. Soc Sci Med 74:907–914.
- Coan JA (2010): Adult attachment and the brain. J Soc Pers Relat 27:210–217.
- Slavich GM (2020): Social safety theory: A biologically based evolutionary perspective on life stress, health, and behavior. Annu Rev Clin Psychol 16:265–295.
- Slavich GM (2022): Social Safety Theory: Understanding social stress, disease risk, resilience, and behavior during the COVID-19 pandemic and beyond. Curr Opin Psychol 45:101299.

- Pizzagalli DA, Jahn AL, O'Shea JP (2005): Toward an objective characterization of an anhedonic phenotype: A signal-detection approach. Biol Psychiatry 57:319–327.
- 23. Ribot T (1896): La Psychologie des Sentiment. Paris: Felix Alcan.
- Der-Avakian A, Pizzagalli DA (2018): Translational assessments of reward and anhedonia: A tribute to Athina Markou. Biol Psychiatry 83:932–939.
- Pizzagalli DA (2022): Toward a better understanding of the mechanisms and pathophysiology of anhedonia: Are we ready for translation? Am J Psychiatry 179:458–469.
- Huys QJ, Pizzagalli DA, Bogdan R, Dayan P (2013): Mapping anhedonia onto reinforcement learning: A behavioural meta-analysis. Biol Mood Anxiety Disord 3:12.
- Castellano P, Gigli V, Ghezzi V, Ang Y-S, Schettino M, Pizzagalli DA, Ottaviani C (2023): Momentary gustative-olfactory sensitivity and tonic heart rate variability are independently associated with motivational behavior. Int J Psychophysiol 186:1–9.
- Russell D, Peplau LA, Cutrona CE (1980): The revised UCLA Loneliness Scale: Concurrent and discriminant validity evidence. J Pers Soc Psychol 39:472–480.
- Solano L, Coda R (1994): Relazioni, Emozioni, Salute: Introduzione Alla Psicoimmunologia. Padova: Piccin.
- Insel T, Cuthbert B, Garvey M, Heinssen R, Pine DS, Quinn K, et al. (2010): Research Domain Criteria (RDoC): Toward a new classification framework for research on mental disorders. Am J Psychiatry 167:748–751.
- Johnstone V, Alsop B (2000): Reinforcer control and human signaldetection performance. J Exp Anal Behav 73:275–290.
- Do CB, Batzoglou S (2008): What is the expectation maximization algorithm? Nat Biotechnol 26:897–899.
- Shiffman S, Stone AA, Hufford MR (2008): Ecological momentary assessment. Annu Rev Clin Psychol 4:1–32.
- Bailey T, Shahabi L, Tarvainen M, Shapiro D, Ottaviani C (2019): Moderating effects of the valence of social interaction on the dysfunctional consequences of perseverative cognition: An ecological study in major depression and social anxiety disorder. Anxiety Stress Coping 32:179–195.
- Muthén LK, Muthén BO (1998–2018): Mplus User's Guide, 8th ed. Los Angeles, CA: Muthén & Muthén.
- Bakker JM, Goossens L, Lange I, Michielse S, Schruers K, Lieverse R, et al. (2017): Real-life validation of reduced reward processing in emerging adults with depressive symptoms. J Abnorm Psychol 126:713–725.
- Grippo AJ, Carter CS, McNeal N, Chandler DL, LaRocca MA, Bates SL, Porges SW (2011): 24-hour autonomic dysfunction and depressive behaviors in an animal model of social isolation: Implications for the study of depression and cardiovascular disease. Psychosom Med 73:59–66.
- Peechatka AL, Whitton AE, Farmer SL, Pizzagalli DA, Janes AC (2015): Cigarette craving is associated with blunted reward processing in nicotine-dependent smokers. Drug Alcohol Depend 155:202–207.
- Bernstein MJ, Zawadzki MJ, Juth V, Benfield JA, Smyth JM (2018): Social interactions in daily life: Within-person associations between momentary social experiences and psychological and physical health indicators. Journal of Social and Personal Relationships 35:372–394.
- Vella EJ, Kamarck TW, Flory JD, Manuck S (2012): Hostile mood and social strain during daily life: A test of the transactional model. Ann Behav Med 44:341–352.
- Peirce RS, Frone MR, Russell M, Cooper ML, Mudar P (2000): A longitudinal model of social contact, social support, depression, and alcohol use. Health Psychol 19:28–38.
- Matthews GA, Nieh EH, Vander Weele CM, Halbert SA, Pradhan RV, Yosafat AS, et al. (2016): Dorsal raphe dopamine neurons represent the experience of social isolation. Cell 164:617–631.
- Tomova L, Tye K, Saxe R (2021): The neuroscience of unmet social needs. Soc Neurosci 16:221–231.
- Heinrich LM, Gullone E (2006): The clinical significance of loneliness: A literature review. Clin Psychol Rev 26:695–718.

- Mumtaz F, Khan MI, Zubair M, Dehpour AR (2018): Neurobiology and consequences of social isolation stress in animal model-A comprehensive review. Biomed Pharmacother 105:1205–1222.
- Ibi D, Takuma K, Koike H, Mizoguchi H, Tsuritani K, Kuwahara Y, et al. (2008): Social isolation rearing-induced impairment of the hippocampal neurogenesis is associated with deficits in spatial memory and emotionrelated behaviors in juvenile mice. J Neurochem 105:921–932.
- Koike H, Ibi D, Mizoguchi H, Nagai T, Nitta A, Takuma K, et al. (2009): Behavioral abnormality and pharmacologic response in social isolation-reared mice. Behav Brain Res 202:114–121.
- Zlatković J, Todorović N, Bošković M, Pajović SB, Demajo M, Filipović D (2014): Different susceptibility of prefrontal cortex and hippocampus to oxidative stress following chronic social isolation stress. Mol Cell Biochem 393:43–57.
- 49. Shaver PR, Furman WD, Buhrmester D (1985): Transition to college: Network changes, social skills, and loneliness. In: Duck S, Perlman D, editors. Understanding personal relationships: An interdisciplinary approach. Thousand Oaks: Sage Publications, 193–219.
- Snodgrass MA (1987): The relationships of differential loneliness, intimacy, and characterological attributional style to duration of loneliness. Soc Behav Pers 2:173–186.
- Peciña S, Berridge KC (2000): Opioid site in nucleus accumbens shell mediates eating and hedonic 'liking' for food: Map based on microinjection Fos plumes. Brain Res 863:71–86.
- Bayer HM, Glimcher PW (2005): Midbrain dopamine neurons encode a quantitative reward prediction error signal. Neuron 47:129–141.
- Steinberg EE, Keiflin R, Boivin JR, Witten IB, Deisseroth K, Janak PH (2013): A causal link between prediction errors, dopamine neurons and learning. Nat Neurosci 16:966–973.
- Santesso DL, Evins AE, Frank MJ, Schetter EC, Bogdan R, Pizzagalli DA (2009): Single dose of a dopamine agonist impairs reinforcement learning in humans: Evidence from event-related potentials and computational modeling of striatal-cortical function. Hum Brain Mapp 30:1963–1976.
- Pizzagalli DA, Evins AE, Schetter EC, Frank MJ, Pajtas PE, Santesso DL, Culhane M (2008): Single dose of a dopamine agonist impairs reinforcement learning in humans: Behavioral evidence from a laboratory-based measure of reward responsiveness. Psychopharmacology 196:221–232.
- Cacioppo S, Capitanio JP, Cacioppo JT (2014): Toward a neurology of loneliness. Psychol Bull 140:1464–1504.
- Colonnello V, Petrocchi N, Farinelli M, Ottaviani C (2017): Positive social interactions in a lifespan perspective with a focus on opioidergic and oxytocinergic systems: Implications for neuroprotection. Curr Neuropharmacol 15:543–561.
- Kalin NH, Shelton SE, Barksdale CM (1988): Opiate modulation of separation-induced distress in non-human primates. Brain Res 440:285–292.
- Trezza V, Damsteegt R, Achterberg EJM, Vanderschuren LJMJ (2011): Nucleus accumbens μ-opioid receptors mediate social reward. J Neurosci 31:6362–6370.
- Hsu DT, Sanford BJ, Meyers KK, Love TM, Hazlett KE, Wang H, et al. (2013): Response of the μ-opioid system to social rejection and acceptance. Mol Psychiatry 18:1211–1217.
- Hsu DT, Sanford BJ, Meyers KK, Love TM, Hazlett KE, Walker SJ, et al. (2015): It still hurts: Altered endogenous opioid activity in the brain during social rejection and acceptance in major depressive disorder. Mol Psychiatry 20:193–200.
- Cacioppo JT, Norris CJ, Decety J, Monteleone G, Nusbaum H (2009): In the eye of the beholder: Individual differences in perceived social isolation predict regional brain activation to social stimuli. J Cogn Neurosci 21:83–92.
- D'Agostino AE, Kattan D, Canli T (2019): An fMRI study of loneliness in younger and older adults. Soc Neurosci 14:136–148.
- 64. Badal VD, Lee EE, Daly R, Parrish EM, Kim H-C, Jeste DV, Depp CA (2022): Dynamics of loneliness among older adults during the COVID-19 pandemic: Pilot study of ecological momentary assessment with network analysis. Front Digit Health 4:814179.
- Wang R, Chen F, Chen Z, Li T, Harari G, Tignor S, et al. (2014): StudentLife: Assessing mental health, academic performance and

- behavioral trends of college students using smartphones. Proc 2014 ACM International Joint Conference on Pervasive and Ubiquitous Computing, ACM 3-14.
- Hawkley LC, Hughes ME, Waite LJ, Masi CM, Thisted RA, Cacioppo JT (2008): From social structural factors to perceptions of relationship quality and loneliness: The Chicago health, aging, and social relations study. J Gerontol B Psychol Sci Soc Sci 63:S375– S384
- Routasalo PE, Savikko N, Tilvis RS, Strandberg TE, Pitkälä KH (2006): Social contacts and their relationship to loneliness among aged people – A population-based study. Gerontology 52:181–187.
- 68. Brooks SK, Webster RK, Smith LE, Woodland L, Wessely S, Greenberg N, Rubin GJ (2020): The psychological impact of quarantine and how to reduce it: Rapid review of the evidence. Lancet 395:912–920.
- Hagerty SL, Williams LM (2020): The impact of COVID-19 on mental health: The interactive roles of brain biotypes and human connection. Brain Behav Immun Health 5:100078.
- Medda E, Toccaceli V, Gigantesco A, Picardi A, Fagnani C, Stazi MA (2022): The COVID-19 pandemic in Italy: Depressive symptoms immediately before and after the first lockdown. J Affect Disord 298:202–208.
- Wieman ST, Fields JS, Arditte Hall KA, MacDonald HZ, Liverant GI (2024): Effects of the COVID-19 pandemic on anhedonia, reward exposure and responsiveness, and sleep in college students. Journal of American College Health 72:351–355.
- Tomova L, Wang KL, Thompson T, Matthews GA, Takahashi A, Tye KM, Saxe R (2020): Acute social isolation evokes midbrain craving responses similar to hunger. Nat Neurosci 23:1597–1605.

SUPPLEMENTARY INFORMATION

Daily Social Isolation Maps Onto Distinctive Features of Anhedonic Behavior: A Combined Ecological and Computational Investigation

Gigli et al.

S1. Exclusionary criteria

Exclusionary criteria were (a) history or presence of serious medical conditions; (b) self-reported formal diagnosis of psychiatric disorder or problematic substance use; (c) neurological disorders; (d) use of drugs/medications; and (e) pregnancy or breast-feeding. Participants were informed that they could win up to 20 euros for their participation.

S2. Revised UCLA Loneliness scale

Examples of items are "How often do you feel that you lack companionship?" or "How often do you feel outgoing and friendly?", and answers are provided on a Likert scale from 0 = Never to 4 = Often. The total score ranges from 20 to 80.

S3. Questions regarding the COVID-19 situation

The following questions about individuals' lifestyle, social isolation (objective or perceived) and recent changes in habits due to the COVID-19 outbreak and subsequent restrictions were administered:

- 1. Do you live alone?
- 2. If you do not live alone, with whom do you live?
- 3. Has the spread of the virus and the restrictions forced you to spend a lot of time alone?
- 4. Has the spread of the virus and the restrictions significantly impacted your lifestyle and led to substantial changes?
- 5. Have you ever been in isolation because you contracted COVID-19 or came into contact with someone who tested positive?
- 6. In the past 6 months, have you been in isolation because you contracted COVID-19 or came into contact with someone who tested positive?
- 7. Are you currently in isolation?

S4. Probabilistic Reward Task

The PRT was administered in the laboratory via E-Prime (version 3.0). As depicted in Figure S1, the task involves 300 trials, divided into 3 blocks separated by 30-sec breaks. In each trial, participants saw a cross for 1000-1400 ms in the center of the screen, then a cartoon face without the mouth for

500 ms. After that, a mouth appeared for 100 ms. The mouth could be short (10.00 mm) or long (11.00 mm). Next, the cartoon face without the mouth appeared again for an additional 1500 ms. The difference between the short and long mouths was small (just 1 mm), making it difficult to distinguish them (22). Participants were asked to decide if the mouth was short or long by pressing a button ("v" or "m") on the keyboard, counterbalanced across participants. Importantly, not every accurate answer receives a reward. Moreover, and unbeknownst to participants, correct identification of one of the mouth lengths (defined as the "rich stimulus") was rewarded three times more frequently compared to the other ("lean stimulus"). Within each block, the short and long stimuli were presented equally often in a pseudorandomized sequence with the constraint that the same stimulus was not presented more than three times consecutively. For each block, reward feedback ("Correct!! You won 20 cents") was presented after 40 correct responses for 1500 ms after the participant's choice and was followed by 250ms of blank screen. During each block, correct identifications of the rich stimulus received reward feedback for 30 times while correct identifications of the lean one were followed by the positive feedback only 10 times. A controlled reinforcer procedure was used to provide reward feedback according to a pseudorandom schedule that determined which specific trials were to be rewarded for correct choices. If a participant failed to make a correct identification for a trial in which feedback was scheduled, reward feedback was delayed until the next correct response of the same stimulus type (rich or lean). When the reward was not given because the participant was inaccurate (or accurate, but no feedback was scheduled) a blank screen was displayed for 1750 ms. The total task duration was approximately 24 minutes. Participants were informed that they could potentially win up to €20; however, the actual reward was fixed and not contingent on their performance. Therefore, at the conclusion of the task, all participants received a monetary reward of €20.

Response Bias (*log b*) was calculated as:

$$\log \mathbb{II}b = \frac{1}{2} log \left[\frac{(RICHcorrect + 0.5) \times (LEANincorrect + 0.5)}{(RICHincorrect + 0.5) \times (LEANcorrect + 0.5)} \right],$$

In addition, discriminability (log d) was computed as a control measure of participants' ability to discriminate between the two stimuli. Discriminability reflects task difficulty and was calculated as:

$$\log \mathbb{Z} d = \frac{1}{2} lod \left[\frac{(RICHcorrect + 0.5) \times (LEANcorrect + 0.5)}{(RICHincorrect + 0.5) \times (LEANincorrect + 0.5)} \right],$$

0.5 was added to each variable in order to make the calculation of the Response Bias and discriminability possible in cases in which one of the raw cells was equal to 0.

Following previous studies, secondary analyses evaluating accuracy and reaction times (RT) were performed to assess overall task performance. Overall, trials with RTs less than 150 ms or longer than 1500 ms were excluded to remove outliers; then, remaining trials with RTs (following natural log transformation) falling outside the mean $3 \pm SD$ were considered as additional outliers and excluded.

S5. Computation Modeling of PRT

The 'Stimulus-Action' model adopts the standard Rescorla-Wagner premise. Both stimuli were assumed to be totally distinct, and rewards were associated with separate stimulus-action pairs. The 'Action' model proposed that participants only learned the value of their actions when forming expectations, independently of the stimuli. The third model, 'Belief', assumed that participants are uncertain of which stimulus was actually presented on each trial and, thus, the rewards were associated with a combination of two uncertainty-weighted stimulus-action associations. Finally, the 'Punishment' model examined whether trials in which no reward was delivered were treated as aversive losses.

S6. Computation of Learning Rate and Reward Sensitivity in the transformed space

Learning rate is not constrained to the range of 0-1 and reward sensitivity is not constrained to the range of 0 to +inf. Instead, both parameters are unconstrained from -inf to +inf, but larger values still indicate greater learning rate and reward sensitivity. Assuming ε is the actual learning rate with a constraint from 0 to 1, we have presented learning rate in the transformed space as $\log \frac{\varepsilon}{1-\varepsilon}$, which is unconstrained from -inf to +inf. Similarly, assuming ρ is the actual reward sensitivity with a constraint from 0 to +inf, we have presented reward sensitivity in the log-transformed space as $\log \rho$, which is unconstrained from -inf to +inf.

The role of reward sensitivity and learning rate in achieving optimal performance in the task has been explored. Optimal performance was defined as a 3:1 response ratio of rich to lean choices, reflecting the reward probabilities in the task setup (rich trials rewarded at 60% and lean trials at 20%). This ratio is indicative of optimal decision-making, where choice allocation proportionally matches the reward probabilities – a principle derived from the probabilistic matching law.

For the simulations, we varied RS and LR around the mean values observed in our sample data. Specifically, RS and LR were varied by ± 1 and ± 2 standard deviations from these means. This approach was chosen to reflect a range of behavior that spans from typical to extreme within our observed participant data, thus providing insights into how deviations from the mean affect task performance.

We generated 500 surrogate datasets for each of the 25 RS-LR combinations, with each dataset consisting of 300 trials. We analyzed these datasets using a sliding window of 50 trials to monitor the evolution of the rich:lean response ratio. The results highlighted a nuanced dynamic (Figure S3). For example, if you look at the third row, when learning rate is kept constant at the mean, excessively high RS led participants to disproportionately favour the rich option, deviating from the optimal 3:1 ratio (indicated by the red horizontal line in the plots). Conversely, too low RS resulted in insufficient responsiveness to rewards, failing to leverage the richer option adequately. Regarding LR, if you look at the third column, when RS is kept constant at the mean, high values of LR led to overfitting to recent outcomes, causing fluctuations and instability in choice patterns where the rich:lean response ratio is close to 1. On the other hand, low values of LR hindered timely adaptation to the task's reward contingencies.

These findings illustrate the importance of balancing RS and LR to achieve a response ratio that aligns with the optimal strategy defined by the reward probabilities of the task.

S7. Outlier management

A single outlier at the level of enduring individual differences (i.e., between-persons level) was identified as potentially influential data point using standardized scores >|3| for univariate outliers. Multivariate outliers were identified based on their multivariate Mahalanobis distance (for p < .01). While no multivariate outliers were detected, a potentially influential subject was identified for Valence. Removing this subject from the analyses did not change the substantive results, and differences between the zero-order correlations calculated on the final sample and those excluding the univariate outlier did not exceed |.02|.

S8. Ecological Momentary Assessment data analysis

EMA observations (Level 1 or within-person level) were nested within individuals (Level 2, or between-person level). While loneliness and Δ Response Bias (as well as its subcomponents, reward

sensitivity and learning rate) represents between-person variables by design (i.e., they were measured once), EMA measures represent within-person variables. In both Model 1 and Model 2, the between-person (stable) components of valence and duration of daily social interaction were modeled at the between-person level (i.e., they were centered at their latent means), and their within-person components were specified as unstructured at Level 1 of analysis (i.e., the within-person level was fully saturated).

Participants with < 30% of valid assessments on the EMA measures were not retained for this analysis; however, findings are the same in terms of effect sizes, statistical, and practical significance when a more stringent criterion (<25%) is adopted.

S9. Limitations of the study

The study relies on self-reported measures using a convenience sample, and data were collected in a single national context. Moreover, it is difficult to completely disentangle objective and perceived social isolation in an unselected sample in which the mismatch between actual and perceived social contacts is expected to be lower compared to what is usually seen in clinical populations (1). Notwithstanding, thanks to the use of an EMA design to capture ongoing daily levels of interactions, distinctive patterns of results for actual and perceived social isolation were found. However, the EMA design does not include inquiries about participants' modes of communication, specifically, whether interactions occurred in person or online. This consideration should have been taken into account, as some research suggests that online interactions may detrimentally affect the sense of social bonding, particularly when employed to alleviate social disconnection (2,3). The existing literature is inconsistent on this topic; while several studies support the compensatory theory of technology use in terms of mental well-being and reduced loneliness during the pandemic (4,5), others demonstrate no correlation between the frequency of virtual social interaction and perceived isolation (6). The last limitation is indeed the fact that the present study was conducted during COVID-19 social restrictions (7,8). It has to be noted, however, that studies investigating the role of social relationships during the pandemic showed how perceived social support and social network size may act as a buffer against COVID-related physical and mental health concerns, fatigue and general psychological distress (9,10).

References

- 1. Wang J, Mann F, Lloyd-Evans B, Ma R, Johnson S (2018): Associations between loneliness and perceived social support and outcomes of mental health problems: a systematic review. *BMC Psychiatry* 18: 156.
- 2. Geirdal AØ, Ruffolo M, Leung J, Thygesen H, Price D, Bonsaksen T, *et al.*, (2021): Mental health, quality of life, wellbeing, loneliness and use of social media in a time of social distancing during the COVID-19 outbreak. A cross-country comparative study. *J Ment Health* 30: 148–155. https://doi.org/10.1080/09638237.2021.1875413.
- Bonsaksen T, Ruffolo M, Price D, Leung J, Thygesen H, Lamph G, et al., (2023):
 Associations between social media use and loneliness in a cross-national population: Do motives for social media use matter? Health Psychol Behav Med 11: 2158089.

 https://doi.org/10.1080/21642850.2022.2158089.
- 4. Sahi RS, Schwyck ME, Parkinson C, Eisenberger NI (2021): Having more virtual interaction partners during COVID-19 physical distancing measures may benefit mental health. *Sci Rep* 11: 18273. https://doi.org/10.1038/s41598-021-97421-1.
- 5. Brown G, Greenfield PM (2021): Staying connected during stay-at-home: Communication with family and friends and its association with well-being. *Hum Behav and Emerg* 3: 147–156. https://doi.org/10.1002/hbe2.246.
- 6. Towner E, Tomova L, Ladensack D, Chu K, Callaghan B (2022): Virtual social interaction and loneliness among emerging adults amid the COVID-19 pandemic. *Curr Res Ecol Soc Psychol* 3: 100058. https://doi.org/10.1016/j.cresp.2022.100058.
- 7. Bzdok D, Dunbar RIM (2022): Social isolation and the brain in the pandemic era. *Nat Hum Behav* 6: 1333–1343.
- 8. Forbes PAG, Pronizius E, Feneberg AC, Nater UM, Piperno G, Silani G, *et al.* (2023): The effects of social interactions on momentary stress and mood during COVID -19 lockdowns. *British J Health Psychol* 28: 306–319.
- 9. Nitschke JP, Forbes PAG, Ali N, Cutler J, Apps MAJ, Lockwood PL, Lamm C (2021): Resilience during uncertainty? Greater social connectedness during COVID-19 lockdown is associated with reduced distress and fatigue. *British J Health Psychol* 26: 553–569.
- 10. Szkody E, Stearns M, Stanhope L, McKinney C (2021): Stress-Buffering Role of Social Support during COVID-19. *Fam Process* 60: 1002–1015.

Figure S1. Illustration of the Probabilistic Reward Task (A) and graphical representation of the current results in terms of Response Bias (B), Accuracy (C), Discriminability (D), and Reaction Times (E).

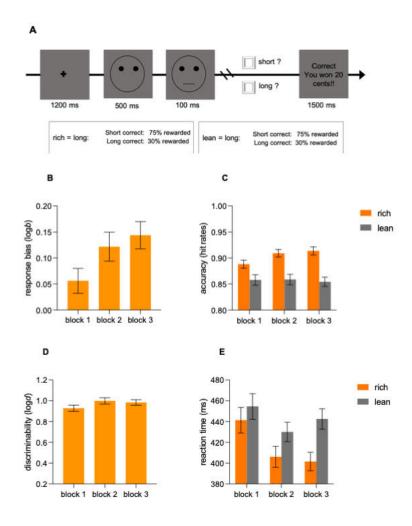
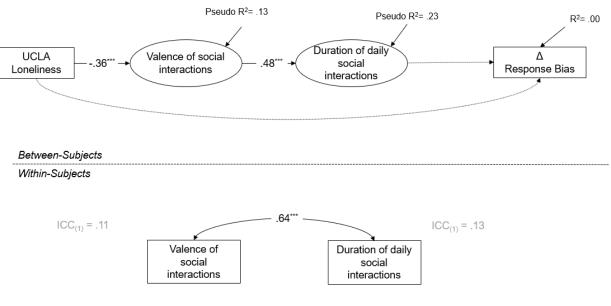


Figure S2. Completely Standardized Estimates from the EMA Model 1.



 ${\rm SB}\chi^{2}_{(d\!f\!=\!1)}=.11\;,p=.73,\,{\rm RMSEA}=0;\,{\rm CFI}=1.00;\,{\rm TLI}=1.00;\,{\rm SRMR}_{within}=.000;\,{\rm SRMR}_{between}=.008$

Note. RMSEA = Root Mean Square Error of Approximation; CFI = Comparative Fit Index; TLI = Tucker-Lewis Index; SRMR = Standardized Root Mean Square Residual. Dotted lines represent non statistically significant paths. One-headed arrows represent direct effects, while double-headed arrows represent covariances. Squares indicated observed variables, while circles represent latent (between-subject level) components of EMA measures. *p < .01, **p < .01, ***p < .001.

Figure S3. Role of reward sensitivity (RS) and learning rate (LR) in achieving optimal performance in the PRT.

