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Efficacy of a combined food-response inhibition and attention training for weight loss[☆]

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This Current Opinion in Behavioral Sciences article reviews trials that evaluated an obesity treatment that combines response-inhibition training with high-calorie foods and training designed to reduce attention for high-calorie foods. Two randomized controlled trials suggest that food-response inhibition and attention training produced significant body-fat loss, along with a reduction in valuation of, and reward-region response to, high-calorie foods. However, these effects did not emerge in a third trial, potentially because this trial used more heterogeneous food images, which reduced inhibition learning and attentional learning. Collectively, results suggest that food-response inhibition and attention training can devalue high-calorie foods and result in weight loss, but only if a homogeneous set of high-calorie and low-calorie food images is used.

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Efficacy of a combined food-response inhibition and attention training for weight loss

The increasing worldwide prevalence of obesity continues to be a primary health concern as it accounts for over 2.8 million deaths annually and is the second leading cause of mortality [1]. Unfortunately, the most common treatments, behavioral weight-loss programs, almost never produce lasting weight loss [2], suggesting that it would be useful to evaluate treatments that use a different approach. Herein, we discuss the conceptual rational for dually improving inhibitory control in response to high-calorie foods and reducing attentional bias for high-calorie foods to produce weight loss and trials that have evaluated this combined training. (See Veling et al.; Lawrence et al.; Chambers et al.; etc., this issue, for reviews of studies focusing on only response-inhibition training).

Neural vulnerability factors that predict future weight gain

It is posited that obesity results from elevated reward and attention brain-region response to high-calorie foods and their cues coupled with lower recruitment of inhibitory regions that modulate food reward and attention responsivity [3–5]. In support, prospective research has found that individuals who exhibit greater activity in brain regions implicated in reward processing (striatum, orbitofrontal cortex) in response to high-calorie food images show elevated future ad lib caloric intake [6] and future weight gain [7–10]. Attentional bias for highcalorie food also predicts elevated future ad lib intake [11,12] and future weight gain [13]. Moreover, lower dorsolateral prefrontal cortex activity, an inhibitory region, in response to high-calorie food images, predicted elevated future ad lib intake [14]. Further, lower recruitment of inhibitory-control regions (inferior, middle, and superior frontal gyri) during a delay-discounting task

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predicted elevated future weight gain [15], converging with evidence that lower inhibitory control in response to high-calorie foods predicted elevated future weight gain [16-19]. However, another study did not find a relation between N2 event-related potential data, a neural indicator of inhibitory control, and future weight gain over a 12-week follow-up [20].

Cognitive science tools for reducing neural vulnerability factors for excess weight gain

Data suggest that an intervention that reduces responsivity of reward and attention regions to high-calorie foods and increases responsivity of inhibitory regions may reduce overeating prompted by high-calorie food images and cues, and promote weight loss. There is evidence that manipulations of motor responses to food images reduce reward valuation of high-calorie foods and produce weight loss. Randomized experiments show that relative to control training, computerized go/no-go and stop-signal training in which participants are signaled to repeatedly respond with a button press to lowcalorie food or nonfood images, and repeatedly refrain from pressing a button to high-calorie food images (which we refer to as food-response inhibition training) produced decreased palatability ratings for the high-calorie foods paired with response-inhibition signals and less ad lib intake of those foods [21-24] and reduced reward-region (mid-insula) response to the high-calorie foods [24]. Food- response inhibition training has also been found to produce weight loss among overweight individuals in some trials [22,25,26], but not others ([20,24,25,27,28]; see for a review [29]).

There is also evidence that attention training can reduce attentional bias toward high-calorie food cues, which should decrease the potential for these cues to induce eating in the absence of hunger. Randomized experiments found that dot-probe training in which attentional bias for high-calorie foods is reduced and attentional bias for low-calorie foods is increased reduces attentional bias for and intake of high-calorie foods relative to control trainings that increase attention for high-calorie foods and reduce attention for low-calorie foods [30-32]. However, an attention-modification task lacking a behavioral response element [12] did not reduce attentional bias that emerged in the dot-probe training that included behavioral responses, implying that the motorresponse element of attention training may be essential. Additionally, a recent meta-analysis found no differences between obese and lean individuals in attention bias to palatable foods, suggesting that targeting attention-bias training alone might be insufficient to change eating behavior [33].

Combined food-response inhibition and attention training

We hypothesized that combining food-response inhibition training and food-attention training might make the training more engaging and effective in reducing highcalorie food intake [34]. We therefore conducted a randomized pilot trial that tested the hypothesis that a multifaceted training protocol, including both food-response inhibition training and attention training, would produce greater reductions in body fat compared with a parallel placebo- response inhibition and attention training with nonfood images [35]. Participants in the intervention were trained to respond to and direct attention toward low-calorie foods and to inhibit responses to and direct attention away from high-calorie foods. Training consisted of 4 weekly 50 min visits in the lab during which participants completed stop-signal training, go/no-go training, respond-signal training, dotprobe training, and visual search training. We used highcalorie and low-calorie foods that were tailored to the preferences of participants, making this a personalized precision-medicine intervention. Overweight or obese adults who completed the intervention showed greater body-fat loss from pretest to posttest than controls who completed placebo training. Intervention versus control participants also showed greater reductions in brain reward (putamen, mid-insula) and attention- (inferior parietal lobe) region response to, and palatability ratings and monetary valuation of, high-calorie foods, consistent with the thesis that training reduces valuation of highcalorie foods.

A follow-up trial found that adding this food-response inhibition and attention training to a dissonance-based obesity-prevention program, which produced significant reductions in body weight and reduced future onset of overweight and obesity [36], resulted in significantly greater body-fat loss from pretest to posttest versus completing the obesity-prevention program and placebo response and attention training with nonfood images [37]. Training consisted of 6 weekly 20 min training visits in the lab during which participants completed stop-signal training, go/no-go training, dot-probe training, and visual search training. High-calorie and low-calorie food images were again tailored to participants' preferences. However, we did not observe significant reductions in palatability or monetary valuation of the high-calorie food images or attentional bias for those images, providing mixed support for the thesis that response and attention training reduces valuation of high-calorie foods.

Given the growing evidence that this multifaceted foodresponse inhibition and attention training produced

body-fat loss in two trials, we conducted a large randomized trial to examine whether food response and attention training would produce significantly greater reductions in body fat compared with placebo response and attention training involving nonfood images over a longer follow-up in a larger sample. We also tested the hypothesis that the food-response inhibition and attention training would reduce functional magnetic resonance imaging fMRI-assessed reward and attention-region response to high-calorie food images and that this would mediate the effects of the intervention on body-fat loss effects. Participants were 179 community-recruited adults with overweight/obesity who completed assessments at pretest, posttest, 3-, 6-, and 12-month follow-ups. The training consisted of 4 weekly visits in the lab during which participants completed stop-signal training, go/nogo training, dot-probe training, and visual search training [38]. Similar to our randomized pilot trial, we used highcalorie and low-calorie foods that were tailored to the preferences of participants. Participants randomized to the intervention showed significantly greater increases in palatability ratings of low-calorie foods compared with placebo controls. However, participants who completed the intervention did not show body-fat loss, reductions in palatability ratings and monetary valuation, or brain reward and attention-region response to high-calorie foods compared with placebo controls.

The lack of effects in this third trial appears to be related to weaker learning in the training evaluated in this trial versus the earlier trials. Specifically, there was no difference in commission errors in the go/no-go task between foods (100% predictive of a response) and filler stimuli (50% predictive) in the intervention group, suggesting that participants did not form an association between high-calorie foods and inhibition. Participants in the current trial also made three-times more no-go commission errors to high-calorie foods (2.9%) than in our pilot trial (1%). Given the important role of associative inhibition learning and accuracy in mediating the effects of food go/no-go training [39,40], the weaker learning may have contributed to the null findings. Further, the increase in attentional bias from pre- to posttest was less pronounced relative to our earlier pilot, both in the intervention and control groups.

One explanation for the weaker learning in the present trial versus the past two trials that produced body-fat loss effects [35,37] is that we used more diverse food images in both the low- and high-calorie food categories than in the trainings for those earlier trials. In the earlier trials, we included only images of fruits and vegetables in the low-calorie (go/attend) category and fewer categories of unhealthy food images. In the current trial, we included other types of low-calorie foods taken from ten different categories, including whole-grain foods, sushi, eggs, fish, and lean meats. The high-calorie foods also

encompassed a more diverse range of 10 subcategories, including sweet foods, pizza, meats, fast food, and drinks. It is possible that the boundaries may have been 'blurred' by the diversity of food images included in both the low- and high-calorie food categories, resulting in weaker associative learning at the category level. Although some studies have demonstrated associative learning and devaluation of food at the item-specific level, these have generally included fewer food items (e.g. 20) compared with the number of items (80) in the current trial [21]. We therefore recommend that future studies use images from more narrow, distinct, and 'meaningful' categories of low-calorie and high-calorie foods to promote stimulus-response learning at the category level [41]. Such learning could also be encouraged by giving participants more explicit instructions about the categories in the task and what to attend to, and by including fewer different food images. Alternatively, a recent meta-analysis found that interleaved practice (i.e. category items presented in an interleaved fashion with items of other categories) with labeled category items can improve learning of heterogeneous categories [42]. Refining the training tasks so that participants focus more on inhibition than on go-responding (i.e. switching their attention away from go stimuli) might also contribute to stronger inhibitory learning. Finally, future research should include sensitive measures of stimulusresponse learning within the training tasks, such as inhibition accuracy to foods versus fillers [22] or memory for stimulus-response contingencies [43] to check that the target mechanisms have been successfully modified [44].

The COVID-19 pandemic may also have made it more difficult to detect body-fat loss effects in the third trial because the lockdown contributed to much higher attrition at the 3-, 6-, and 12-month follow-ups than we observed in past trials of obesity interventions, making it impossible to evaluate the longer-term effects of the intervention. Moreover, lockdowns have been related to increased unhealthy food consumption and reduced physical activity, particularly among overweight individuals [45,46], which may further reduce the ability to find long-term effects. Interestingly, the d effect size for body-fat loss effects from pretest to 12-month follow-up based on the observed means was .32, implying that we might have been able to detect longer-term effects on body fat if the pandemic did not result in such high attrition.

Conclusions

Elevated brain reward and attention-region response and weaker inhibitory-region response to high-calorie foods has predicted future weight gain, suggesting that an intervention that reduces reward and attention-region response and increases inhibitory-region response to such foods might reduce overeating. Findings from cognitive psychology suggest that combining food-response inhibition and attention training might prove useful in reducing neural vulnerability factors that predict future weight gain. Two randomized controlled trials suggest that food-response inhibition and attention training produced significant body-fat loss, along with a reduction in valuation of the high-calorie foods. However, these effects did not emerge in a third trial, most likely because this trial used a more heterogeneous set of the food images, which attenuated learning. It will be important for future research to test whether using more homogeneous sets of images of high-calorie and lowcalorie foods results in larger reductions in valuation of the high-calorie foods and body fat. It would also be useful for future research to evaluate adaptations to food response and attention training that contribute to larger and more reliable intervention effects (e.g. such as gradually fading the response and inhibition cues [28]). Finally, it would be valuable for additional prospective brain-imaging studies to identify individual differences in neural responsivity that predict future weight gain, which may isolate additional neural vulnerability factors to target in future experimental therapeutics trials.

Conflict of interest statement

Nothing declared.

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References and recommended reading

Papers of particular interest, published within the period of review, have been highlighted as:

- · of special interest
- Word Health Organization: Obesity and overweight. 2021.
 Available: (https://www.who.int/nes-room/facts-in-pictures/detail/6-facts-on-obesity) (June 9, 2021).
- Gokee-LaRose J, et al.: Are standard behavioral weight loss programs effective for young adults? Int J Obes 2009, 33:1374-1380
- Boswell RG, Kober H: Food cue reactivity and craving predict eating and weight gain: a meta-analytic review. Obes Rev 2016, 17:159-177.
- Nederkoorn C, et al.: Control yourself or just eat what you like? Weight gain over a year is predicted by an interactive effect of response inhibition and implicit preference for snack foods. Health Psychol 2010, 29:389-393.
- Stice E, Yokum S: Neural vulnerability factors that increase risk for future weight gain. Psychol Bull 2016, 142:447-471.
- Gearhardt AN, et al.: Neural response to fast food commercials in adolescents predicts intake. Am J Clin Nutr 2020, 111:493-502.
- Demos KE, Heatherton TF, Kelley WM: Individual differences in nucleus accumbens activity to food and sexual images predict weight gain and sexual behavior. J Neurosci 2012, 32:5549-5552.

- Stice E, Burger KS, Yokum S: Reward region responsivity predicts future weight gain and moderating effects of the TaqlA allele. J Neurosci 2015. 35:10316-10324.
- Yokum S, et al.: Individual differences in striatum activity to food commercials predict weight gain in adolescents. Obesity 2014, 22:2544-2551.
- Yokum S, Gearhardt AN, Stice E: In search of the most reproducible neural vulnerability factors that predict future weight gain: analyses of data from six prospective studies. Soc Cogn Affect Neurosci 2021,nsab013.
- Nijs IM, et al.: Differences in attention to food and food intake between overweight/obese and normal-weight females under conditions of hunger and satiety. Appetite 2010, 54:243-254.
- Werthmann J, et al.: Attention bias for chocolate increases chocolate consumption—an attention bias modification study. J Behav Ther Exp Psychiatry 2014, 45:136-143.
- Calitri R, et al.: Cognitive biases to healthy and unhealthy food words predict change in BMI. Obesity 2010, 18:2282-2287.
- Cornier MA, et al.: Sex-based differences in the behavioral and neuronal responses to food. Physiol Behav 2010, 99:538-543.
- Kishinevsky FI, et al.: fMRI reactivity on a delay discounting task predicts weight gain in obese women. Appetite 2012, 58:582-592
- Evans GW, Fuller-Rowell TE, Doan SN: Childhood cumulative risk and obesity: the mediating role of self-regulatory ability. Pediatrics 2012, 129:e68-e73.
- Francis LA, Susman EJ: Self-regulation and rapid weight gain in children from age 3 to 12 years. Arch Pediatr Adolesc Med 2009, 163:297-302.
- Schlam TR, et al.: Preschoolers' delay of gratification predicts their body mass 30 years later. J Pediatr 2013, 162:90-93.
- Seeyave DM, et al.: Ability to delay gratification at age 4 years and risk of overweight at age 11 years. Arch Pediatr Adolesc Med 2009. 163:303-308.
- Carbine KA, et al.: Does inhibitory control training reduce weight and caloric intake in adults with overweight and obesity? A preregistered, randomized controlled event-related potential (ERP) study. Behav Res Ther 2021, 136:103784.
- Chen Z, et al.: How does not responding to appetitive stimuli cause devaluation: evaluative conditioning or response inhibition? J Exp Psychol Gen 2016, 145:1687-1701.
- 22. Lawrence NS, et al.: Training response inhibition to food is associated with weight loss and reduced energy intake.

 Appetite 2015, 95:17-28.
- Veling H, Aarts H, Stroebe W: Using stop signals to reduce impulsive choices for palatable unhealthy foods. Br J Health Psychol 2013, 18:354-368.
- 24. Yang Y, et al.: Pilot study of food-specific Go/No-Go training for overweight individuals: brain imaging data suggest inhibition shapes food evaluation. Soc Cogn Affect Neurosci 2021,nsab137.
- Allom V, Mullan B: Two inhibitory control training interventions designed to improve eating behaviour and determine mechanisms of change. Appetite 2015, 89:282-290.
- Veling H, et al.: Targeting impulsive processes of eating behavior via the internet. Effects on body weight. Appetite 2014, 78:102-109.
- Forman EM, et al.: Computerized neurocognitive training for improving dietary health and facilitating weight loss. J Behav Med 2019, 42:1029-1040.

Randomized controlled trial that found that the effect of sweet foodspecific inhibitory control training on weight loss was moderated by implicit preference for sweet foods.

Memarian S, et al.: Can sweet food-specific inhibitory control training via a mobile application improve eating behavior in children with obesity? Br J Health Psychol 2021.

Randomized controlled trial that found sweet food-specific go/no-go training via a mobile app reduced sweet food choice and intake but not

- 29. de Klerk MT, Smeets PAM, la Fleur SE: Inhibitory control as a potential treatment target for obesity. Nutr Neurosci 2022,1-16.
- Kakoschke N, Kemps E, Tiggemann M: Attentional bias modification encourages healthy eating. Eat Behav 2014, **15**:120-124.
- 31. Kemps E, Tiggemann M, Hollitt S: Biased attentional processing of food cues and modification in obese individuals. Health Psychol 2014. 33:1391-1401.
- Kemps E, et al.: Attentional retraining can reduce chocolate consumption. J Exp Psychol Appl 2014, 20:94-102.
- 33. Hagan KE, et al.: A systematic review and meta-analysis of attentional bias toward food in individuals with overweight and obesity. Appetite 2020, 151:104710.

This meta-analytic review synthesizes the available evidence on attentional bias to palatable food in persons with overweight/obesity. The results showed no significant differences between obese and lean individuals in attentional bias to palatable foods. The review highlights the need for improving psychometric properties of attentional bias measures and the examination of potential moderators.

- Stice E, et al.: Training motor responses to food: a novel treatment for obesity targeting implicit processes. Clin Psychol Rev 2016. 49:16-27.
- 35. Stice E, et al.: Pilot test of a novel food response and attention training treatment for obesity: Brain imaging data suggest actions shape valuation. Behav Res Ther 2017. 94:60-70

Randomized controlled trial that found that food response/attention training reduced valuation of high-calorie foods and reward region response to high-calorie foods, and produced weight loss compared to placebo training.

- 36. Stice E, et al.: An experimental therapeutics test of whether adding dissonance-induction activities improves the effectiveness of a selective obesity and eating disorder prevention program. Int J Obes 2018, 42:462-468.
- 37. Stice E, et al.: Enhancing efficacy of a dissonance-based obesity and eating disorder prevention program: experimental therapeutics. *J Consult Clin Psychol* 2021, **89**:793-804.

Randomized controlled trial that found that adding food response/attention training to a dissonance-based obesity prevention program resulted in significantly greater body fat loss compared to completing the obesity prevention program with a placebo training.

- 38. Stice, E., Yokum, S., Gau, J., Veling, H., Lawrence, N., Kemps, E.: Efficacy of a food response and attention training treatment for obesity: a randomized placebo controlled trial. submitted.
- Jones A, et al.: Inhibitory control training for appetitive behaviour change: a meta-analytic investigation of mechanisms of action and moderators of effectiveness. Appetite 2016, 97:16-28.
- 40. Porter L, et al.: Exploring strategies to optimise the impact of food-specific inhibition training on children's food choices Front Psychol 2021, 12:653610

This paper describes the findings of two randomized control trials that suggest that the effect of energy-dense food-specific go/no-go training on food choices in children is moderated by the accuracy on energydense no-go trials.

- 41. Serfas BG, Florack A, Büttner OB, Voegeding T: What does it take for sour grapes to remain sour? Persistent effects of behavioral inhibition in go/no-go tasks on the evaluation of appetitive stimuli, Motivation Sci 2017, 3:1-18.
- 42. Brunmair M, Richter T: Similarity matters: a meta-analysis of interleaved learning and its moderators. Psychol Bull 2019, 145:1029-1052

This meta-analysis synthesizes the available evidence on the effects of interleaved learning across different materials, settings, and populations. The results showed an effect for inductive learning of complex visual material with high similarity between categories but not within categories.

- 43. Chen Z, et al.: Go/no-go training changes food evaluation in both morbidly obese and normal-weight individuals. J Consult Clin Psychol 2018, 86:980-990.
- 44. Wiers RW, Boffo M, Field M: What's in a trial? On the importance of distinguishing between experimental lab studies and randomized controlled trials: the case of cognitive bias modification and alcohol use disorders. J Stud Alcohol Drugs 2018, 79:333-343.
- 45. Poelman MP, et al.: Eating behavior and food purchases during the COVID-19 lockdown: a cross-sectional study among adults in the Netherlands. *Appetite* 2021, **157**:105002.
- Robinson E, et al.: Obesity, eating behavior and physical activity during COVID-19 lockdown: a study of UK adults. Appetite 2021

This study found that the 2020 COVID-19 social lockdown had a negative influence on weight-related behaviors among adults with higher BMI. Results show that during the lockdown, higher BMI was associated with lower levels of physical activity and diet quality, and greater frequency of overeating