Breakfast and exercise contingently affect postprandial metabolism and energy balance in physically active males

Javier T. Gonzalez1*, Rachel C. Veasey1, Penny L. S. Rumbold2 and Emma J. Stevenson1

1Brain, Performance and Nutrition Research Centre, School of Life Sciences, Northumbria University, Northumberland Building, Newcastle upon Tyne NE1 8ST, UK
2Department of Sport and Exercise Sciences, School of Life Sciences, Northumbria University, Northumberland Building, Newcastle upon Tyne NE1 8ST, UK

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Abstract
The present study examined the impact of breakfast and exercise on postprandial metabolism, appetite and macronutrient balance. A sample of twelve (blood variables n 11) physically active males completed four trials in a randomised, crossover design comprising a continued overnight fast followed by: (1) rest without breakfast (FR); (2) exercise without breakfast (FE); (3) breakfast consumption (1859 kJ) followed by rest (BR); (4) breakfast consumption followed by exercise (BE). Exercise was continuous, moderate-intensity running (expending approximately 2·9 MJ of energy). The equivalent time was spent sitting during resting trials. A test drink (1500 kJ) was ingested on all trials followed 90 min later by an ad libitum lunch. The difference between the BR and FR trials in blood glucose time-averaged AUC following test drink consumption approached significance (BR: 4·33 (SEM 0·14) v. FR: 4·75 (SEM 0·16) mmol/l; P=0·08); but it was not different between FR and FE (FE: 4·77 (SEM 0·14) mmol/l; P=0·65); and was greater in BE (BE: 4·97 (SEM 0·13) mmol/l) v. BR (P=0·012). Appetite following the test drink was reduced in BR v. FR (P=0·006) and in BE v. FE (P=0·029). Following lunch, the most positive energy balance was observed in BR and least positive in FE. Regardless of breakfast, acute exercise produced a less positive energy balance following ad libitum lunch consumption. Energy and fat balance is further reduced with breakfast omission. Breakfast improved the overall appetite responses to foods consumed later in the day, but abrogated the appetite-suppressive effect of exercise.

Key words: Appetite; Fasted state; Glycaemia; Fat oxidation

Regular breakfast consumption has been inversely associated with BMI1, yet it is not clear whether this association is due to differences in energy expenditure, metabolism or energy intake. Although the ostensible benefits of regular breakfast consumption could be due to improved diet composition with breakfast cereals1, rather than meal pattern per se, acute consumption of breakfast can enhance glucose tolerance, insulin sensitivity and subjective and physiological satiety responses to a test drink2.

A recent position statement concluded that further research is required in regular exercisers with regards to meal pattern, metabolism and appetite regulation5, as research in exercising individuals in this area is sparse. However, this population do use diet/exercise strategies, such as training in the fasted state, to control body fat/mass and improve metabolic adaptations to training6. Exercise attenuates adverse dietary outcomes such as fat-induced glucose intolerance5, and the nutritional state in which exercise is performed can modulate the magnitude of these improvements5. Exercise in the fasted state results in a greater reliance on fat as a substrate6 and has led to its use as a tool to reduce body fat by athletes4. Training in the fasted state also leads to enhanced fat transporter protein mRNA content5, mitochondrial enzyme activity and maximal aerobic capacity7, making exercise in the fasted state an attractive proposition for both recreational and elite athletes. On the other hand, high carbohydrate availability during exercise training may result in improved body composition, as gains in fat-free mass are amplified, whilst fat loss is similar8. Hence, although there is a suggestion that exercise in the fasted state can maximise some benefits already associated with exercise, ensuing effects on appetite and metabolism are not entirely clear.

The regulation of acute energy balance involves (not exclusively) the exposure and sensitivity to the circulating

Abbreviations: AUCINS/GLU, serum insulin AUC to blood glucose AUC ratio; BE, overnight fast followed by breakfast and exercise; BR, overnight fast followed by breakfast and rest; FE, overnight fast followed by exercise without breakfast; FR, overnight fast followed by rest without breakfast; GLP-1, glucagon-like peptide 1; ISIMatsuda, Matsuda insulin sensitivity index; VAS, visual analogue scale.

*Corresponding author: J. T. Gonzalez, fax +44 191 243 7012, email javier.gonzalez@northumbria.ac.uk
hormonal and metabolic milieu\(^\text{(9)}\), which underscores the importance of determining these changes concomitant with measuring energy balance. Exercise training improves glucose tolerance\(^\text{(9)}\), yet acute exercise effects are less lucid\(^\text{(10–13)}\). Muscle glucose uptake is increased after exercise\(^\text{(14)}\), as assessed in rat hindlimb muscle. However, both this method and the most commonly used technique for assessing insulin sensitivity in human subjects (the euglycaemic–hyperinsulinaemic clamp) possess some caveats. First, they ignore the gastrointestinal response to food ingestion. Direct contact of nutrients with L-cells in the intestine stimulates secretion of glucagon-like peptide 1 (GLP-1), which potentiates insulin secretion and sensitivity and reduces food intake\(^\text{(9)}\). GLP-1 exists in two active forms; in human subjects, the primary circulating form is GLP-1\(_{7–36}\). Acute exercise has been shown to increase GLP-1 concentrations in the fed state\(^\text{(15)}\). Therefore, GLP-1 may be an important mediator in the acute regulation of energy homeostasis regarding breakfast consumption and exercise.

Second, provision of nutrients other than glucose can influence glucose tolerance and insulin sensitivity. Protein, for example, stimulates insulin and/or incretin hormone influence glucose tolerance and insulin sensitivity. Protein, consumption and exercise. Acute regulation of energy homeostasis regarding breakfast Therefore, GLP-1 may be an important mediator in the acute regulation of energy homeostasis regarding breakfast consumption and exercise.

The protocol was approved by the School of Life Sciences Ethics Committee at Northumbria University.

**Preliminary measurements**

Participants undertook preliminary tests to establish: (1) the relationship between \(O_2\) uptake and running speed on a flat treadmill (Woodway ELG, Woodway) using a four-stage, 16 min test; (2) their \(V_{\text{O2peak}}\) using an incremental treadmill test, whereby the gradient was increased by 1\% min to exhaustion, as described previously in detail\(^\text{(29)}\). The duration of the exercise period in the main trials was calculated from submaximal \(O_2\) uptake and \(CO_2\) values in order to expend 2.9 MJ (693 kcal) of energy whilst running at a speed estimated to elicit 60\% \(V_{\text{O2peak}}\). This value was chosen to equate to approximately 1 h on average, whilst maintaining similar energy expenditure across participants. On the same day, participants were familiarised with the visual analogue scales (VAS) to assess subjective appetite sensations in main trials, and it was verbally confirmed that participants did not have any particular disliking of foods contained in the test meals.

**Experimental design**

All participants completed four trials in a randomised (performed by J. T. G with Research Randomizer version 3.0; http://www.randomizer.org/), crossover design separated by \(\geq 7\) d comprising a continued overnight fast followed by: (1) rest without breakfast (FR); (2) exercise without breakfast (FE); (3) breakfast consumption (1859 kJ) followed by rest (BR); and (4) breakfast consumption followed by exercise (BE). By necessity of the design (food intake and exercise), the intervention was not blinded. All trials were performed under similar laboratory conditions (ambient temperature, humidity and pressure; all \(P>0.05\); data not shown). Food and fluid diaries were kept for the day preceding the first trial and participants were instructed to replicate this for all subsequent trials. Alcohol, caffeine and vigorous activity were prohibited for 24 h prior to trials.

On trial days, participants arrived in the laboratory at 07.30 hours after a 10–14 h fast and a cannula was inserted into an antecubital vein for blood sampling. After baseline samples of expired gas and VAS were taken, in breakfast trials (BE and BR), participants consumed a porridge breakfast. In fasting trials (FE and FR), participants were permitted to consume water only, which was consumed \(ad\ li\ bit\ um\) on the first exercise and non-exercise trials, and water consumption was replicated for the following exercise and non-exercise trials (Fig. 1). Following 120 min of rest, during exercise trials (BE and FE), participants ran on a treadmill at 61.1 (SEM 0.6)\% \(V_{\text{O2peak}}\) for 59 (SEM 2) min based on the \(a\ priori\) estimated energy expenditure. Treadmill speed was adjusted accordingly on the first trial to obtain the appropriate \(V_{\text{O2}}\). Changes in speed were noted for duplication in subsequent exercise trials. In resting trials (BR and FR), participants rested for the equivalent amount of time as the exercise trials.

Within 20 min of exercise termination, participants ingested a chocolate milk test drink. Following a 90 min postprandial
period, a homogeneous *ad libitum* test lunch was provided. Participants were provided with an initial 430 g (3694 kJ; 882 kcal) portion of the test meal, which was replaced upon completion. The test meal was terminated when the participant instructed that they felt ‘comfortably full’. Participants were constantly reminded to follow this instruction and were always presented with fresh, warmed portions before participant-induced termination to ensure that the end of a portion was not the reason for meal termination. Remaining food was then removed and weighed out of the sight of the participants to determine energy intake.

**Anthropometric measurements**

Body mass was determined to the nearest 0·1 kg using balance scales (Seca) upon arrival at the laboratory, immediately prior to and following exercise, where participants wore only light clothing. Height was measured to the nearest 0·1 cm using a stadiometer (Seca).

**Test meals**

The breakfast consisted of 72 g oats (Oatso Simple Golden Syrup, Quaker Oats) and 360 ml semi-skimmed milk (Tesco) and provided 1500 kJ of energy (358 kcal; 17 % protein, 63 % carbohydrate and 23 % fat). The test drink was 500 ml of chocolate milk (Yazoo, Campina Limited) and contained concentrations, 4 ml was dispensed into an EDTA vacutainer containing 100 µl aprotinin and immediately centrifuged at 3000 rpm at 4°C for 10 min. Plasma was stored for later determination of GLP-17–36 using an immunoassay (Phoenix Pharmaceuticals, Inc.). Remaining whole blood from 10 ml samples was allowed to stand for 30 min in a non-anticoagulant tube before being centrifuged at 3000 rpm at 4°C for 10 min. Aliquots of serum were then stored for later determination of NEFA (WAKO Diagnostics) and insulin (DIAsource Immuno-Assays S.A.) concentrations in duplicate. All plasma/serum samples were stored at −80°C. The intra-assay CV were 5·6 and 7·2 % for NEFA and insulin, respectively. Inter-assay CV were 8·1, 3·6 and 18·5 % for NEFA, insulin and GLP-17–36, respectively. In order to reduce the inter-assay variation, samples from each participant were analysed during the same run where possible. It was decided that it was unnecessary to adjust analyte concentrations to account for plasma volume changes, as exercise of a similar and greater intensity and duration does not result in changes in plasma volume[15,25].

**Energy expenditure and substrate oxidation**

Expired gas samples were collected using an online gas analysis system (Metalyzer 3B, Cortex) calibrated using gases of known concentrations and a 3 l syringe. Participants wore a facemask and after a 2 min stabilisation phase, 5 min samples were obtained and averaged at baseline, at every 30 min after breakfast consumption (or equivalent time in breakfast omission trials) and at 5, 15, 30, 50, 70 and 90 min following consumption of the test drink. Expired gas was continuously sampled throughout the exercise and averaged over each 5 min period, ignoring the first 5 min to allow for steady-state values.

Substrate metabolism was calculated, assuming negligible protein oxidation, with \( V_{O2} \) and \( V_{CO2} \) production values using stoichiometric equations and was adjusted during exercise to account for the contribution of glycogen to metabolism[20]:

\[
\text{Rate of fat oxidation at rest and during exercise (g/min)} = (1.695 \times V_{O2}) - (1.701 \times V_{CO2}).
\]

\[
\text{Rate of carbohydrate oxidation at rest (g/min)} = (4.585 \times V_{CO2}) - (3.226 \times V_{O2}).
\]

\[
\text{Rate of carbohydrate oxidation during exercise (g/min)} = (4.210 \times V_{CO2}) - (2.962 \times V_{O2}).
\]

\( V_{O2} \) and \( V_{CO2} \) are measured in litres/min.

Energy expenditure was calculated based on fat, glucose and glycogen concentrations providing 40·81, 15·64 and 17·36 kJ/g of energy, respectively. At rest, calculations were based on glucose providing all of the carbohydrate for metabolism, whereas during moderate-intensity exercise,
carbohydrate oxidation is met by both glucose and glycogen providing a 20 and 80 % contribution, respectively.(26)

**Subjective ratings**

Paper-based 100 mm VAS were completed at baseline, prior to and immediately following breakfast and at every 30 min thereafter until exercise (or equivalent time points in breakfast omission trials); further, VAS were completed immediately following exercise and after test drink consumption and at 30 min intervals thereafter. Final VAS were completed following termination of the test meal. Questions asked were used to determine hunger, fullness, satisfaction and prospective food consumption. An overall appetite score was calculated using the following formula, as previously used(27):

\[
\text{Overall appetite} = (\text{hunger} + \text{prospective food consumption}) + (100 - \text{fullness}) + (100 - \text{satisfaction}) / 4.0.
\]

**Statistical analysis**

Due to difficulties associated with blood collection, data for GLP-1 7–36 and NEFA concentrations and appetite sensations were converted into AUC using the trapezoidal rule. Indices of insulin secretion and sensitivity, post-test drink serum insulin AUC to blood glucose AUC ratio (AUCINS/GLU) and Matsuda insulin sensitivity index (ISIMatsuda) were calculated as described previously(28,29). Unless otherwise stated, all data are presented as mean values with their standard errors represented by vertical bars. * Mean value for the FE trial was significantly different from that of BE trial (P<0.05). † Mean value for the FR trial was significantly different from that of BE trial (P<0.05). § Mean value for the FR trial was significantly different from that of BE trial (P<0.05). ¶ Mean value for the BR trial was significantly different from that of BE trial (P<0.05). * Mean value for the BR trial was significantly different from that of FE trial (P<0.05). || Mean value for the BR trial was significantly different from that of BE trial (P<0.05). ‡ Mean value for the FE trial was significantly different from that of BE trial (P<0.05).

**Results**

The participants’ age, height, body mass, BMI and peak O2 uptake (VO2peak) were 23.2 (sd 4.5) years, 178.0 (sd 7.0) cm, 77.2 (sd 5.3) kg, 24.5 (sd 2.0) kg/m2 and 53.1 (sd 5.5) ml/kg per min, respectively.

**Blood glucose**

Blood glucose concentration displayed a trial × time interaction effect (Fig. 2(A); P<0.001). Breakfast consumption reduced time to reach peak blood glucose concentration following test drink ingestion by 10 and 4 min during rest and exercise trials, respectively (P=0.012 and P=0.02, respectively). Peak blood glucose concentration was unaffected by breakfast consumption during resting trials (FR: 5.95 (SEM 0.20) mmol/l; BR: 5.75 (SEM 0.14) mmol/l; P=0.20). No difference was observed in peak or in time to peak blood glucose concentrations in FR v. FE trials (P=0.73 and P=0.28, respectively). However, in BE, blood glucose concentration reached 6.66 (SEM 0.24) mmol/l, significantly greater than FE (5.89 (SEM 0.17) mmol/l; P=0.06) and BR (P=0.036). The difference between the BR and FR trials in AUC for blood glucose approached statistical significance (Fig. 2(B); P=0.09); but it was not significantly different between the FR and FE trials (P=0.65); and was greater in BE v. BR trials (P=0.012).
Integrated effects of breakfast and exercise

Serum insulin

A trial × time interaction effect was observed for serum insulin concentrations \((P<0.001)\), where peak concentrations occurred at 37 (SEM 3) min in the FR trial, and the delay compared with BR (29 (SEM 1) min; \(P=0.09\)) and FE (30 (SEM 4) min; \(P=0.10\)) approached statistical significance. Serum insulin concentrations rose after test drink consumption (Fig. 3(A)) to a similar peak between trials (FR: 682 (SEM 71), BR: 607 (SEM 46), FE: 570 (SEM 72) and BE: 586 (SEM 64) pmol/l; \(P=0.21\)). The greater AUC for serum insulin in FR v. all other trials approached statistical significance (Fig. 3(B); \(P=0.07, P=0.12\) and \(P=0.09\) for BR, FE and BE, respectively).

Indices of insulin secretion and sensitivity

The AUCINS/GLU was similar between FR and BR trials (82 (SEM 7) and 80 (SEM 6) pmol/mmol; \(P=0.45\)), but was reduced by exercise compared with the FR trial (FE: 70 (SEM 7) and BE: 67 (SEM 6) pmol/mmol; \(P=0.03\) and \(P=0.04\) for FE and BE trials, respectively). ISI Matsuda was similar between the trials (12 (SEM 4), 12 (SEM 4), 12 (SEM 4) and 13 (SEM 5) arbitrary units for FR, BR, FE and BE respectively; all \(P=0.05\)).

Serum NEFA

Test drink consumption transiently suppressed NEFA concentrations and a significant trial × time interaction effect was observed (Fig. 4(A); \(P<0.001\)). The time at which the nadir of NEFA concentrations was reached was delayed in the FR trial (81 (SEM 3) min) compared with all other trials (BR: 65 (SEM 3) min, \(P=0.019\); FE: 57 (SEM 3) min, \(P<0.001\); and BE: 55 (SEM 6) min, \(P=0.007\)). The AUC for BR was lower than that for FR and BE trials (Fig. 4(B); \(P=0.01\) and \(P=0.004\), respectively).

Plasma glucagon-like peptide 1\textsubscript{7–36}

There was no trial × time interaction effect or main effects of trial on GLP-1\textsubscript{7–36} concentrations (Fig. 5(A); both \(P>0.05\)).
in the exercise period, an effect which was independent of exercise/rest (Table 1). This resulted in similar carbohydrate balance (intake minus oxidation) post-exercise between FE and BE, in spite of a large difference in carbohydrate balance prior to exercise (pre-exercise: −17 (SEM 2) g and 43 (SEM 2) g, P<0·001; post-exercise: −108 (SEM 7) and −102 (SEM 8) g, P=0·38 for FE and BE trials, respectively). Following consumption of the test drink, energy expenditure and fat oxidation were greater in both exercise trials compared with rest trials, yet carbohydrate oxidation was similar (Table 1).

There was no detectable difference in ad libitum energy intake at lunch (Fig. 6; P=0·78). Hence, when energy intakes from the breakfast and the test drink are taken into consideration, breakfast trials produced a greater total energy intake (Fig. 6; P<0·001). The variation in the compensation of energy intake to account for the increase in energy expenditure (energy intake on exercise trials minus energy intake on resting trials) ranged from −1916 to 3749 kJ (−458 to 895 kcal) in the fasting trials and from −1447 to 3683 kJ (−346 to 880 kcal) in the breakfast trials. A total of seven individuals consumed less in the FE v. FR trial, four individuals partially compensated for exercise, consuming more in the FE v. FR trial, but not enough to overcome the exercise-induced energy expenditure. Only one participant over-compensated for exercise, consuming more than the exercise-induced energy expenditure in the FE v. FR trial. In breakfast trials, six individuals consumed less in the BE v. BR trial, five partially compensated and only one over-compensated for the exercise-induced energy expenditure. No significant relationship was present between the compensation on fast days and the compensation on breakfast days (r = −0·07, P>0·05).

Energy balance post-lunch was most positive with BR and least positive with FE trials (Fig. 7). There was no detectable difference in carbohydrate balance when breakfast was omitted v. consumed, although the difference at rest approached significance (FR v. BR, P=0·06; FE v. BE, P=0·05; Fig. 7). Yet, fat balance was significantly different between all trials, apart from the FR v. BE trial, albeit in BE, a reduction which approached statistical significance was observed (P=0·06).

Subjective ratings

Feelings of hunger during the exercise period were suppressed in FE v. FR (P=0·015) and BE v. BR trials (P=0·016). This was still the case immediately post-exercise in the FE v. FR trial (P=0·002), yet, in the BE v. BR trial, there was no detectable difference (P=0·45). FE also reduced ratings of prospective consumption during and after exercise v. FR (P=0·028 and P=0·032, respectively), whereas BE did not significantly affect prospective consumption ratings compared with BR (P=0·67 and P=0·15, respectively). Overall appetite rating showed similar findings (Fig. 8(A)), where the change from pre- to during the exercise period was significantly different between the FR and the FE trials (2 (SEM 1) v. −11 (SEM 4); P=0·048), but not between the BR and BE trials (6 (SEM 2) v. 0 (SEM 4); P=0·21).

Breakfast did not influence hunger immediately pre-lunch during exercise trials (P=0·11), but did reduce hunger

Energy intake, metabolism and balance

Energy expenditure, fat oxidation and carbohydrate oxidation did not differ at baseline (P=0·43, P=0·13 and P=0·57, respectively).

In the breakfast postprandial period, energy expenditure was not significantly different between the trials (Table 1). Less fat and more carbohydrate were utilised during the breakfast postprandial period in the breakfast trials (i.e. BE and BR v. fasting trials (i.e. FE and FR) (Table 1; P=0·005 and P<0·001, respectively).

The exercise bout lasted for 59 (SEM 2) min and mean O₂ uptake was similar between the FE and BE trials during this period (2·52 (SEM 0·11) and 2·50 (SEM 0·11) litres/min; P=0·54). In spite of the equivalent amount of external work performed, exercise increased energy expenditure during the breakfast trials (3279 (SEM 50) kJ) compared with that during the fasting trials (2627 (SEM 43) kJ; P<0·001).

Breakfast consumption reduced the reliance on fat as a substrate and subsequently raised carbohydrate metabolism

There was also no difference in AUC (Fig. 5(B)), peak or time to peak GLP-1 7–36 concentrations (P=0·17, P=0·27 and P=0·45, respectively).

Fig. 5. (A) Plasma glucagon-like peptide-1 7–36 (GLP-1 7–36) concentration in response to test drink consumption in the overnight fast followed by rest without breakfast (FR, ○), overnight fast followed by exercise (EX) without breakfast (FE, ▲) and overnight fast followed by breakfast and EX (BE, ▽) trials. BL, baseline; PE, pre-EX. (B) Time-averaged GLP-1 7–36 AUC following test drink consumption. Values are means, with standard errors represented by vertical bars.

Energy intake, metabolism and balance

Energy expenditure, fat oxidation and carbohydrate oxidation did not differ at baseline (P=0·43, P=0·13 and P=0·57, respectively).
in resting trials (P=0.006). The same pattern was observed with prospective consumption (FR v. BR; P=0.005; BR v. FE; P=0.005; FE v. BE; P=0.10). However, immediately prior to lunch, overall appetite was suppressed in the BR trial compared with that in both fasting trials (i.e. FE and FR) (P=0.001 and P=0.005, for rest and exercise, respectively; Fig. 8(B)).

There was no detectable difference in AUC for hunger between exercise and rest (P=0.47 and P=0.71 for FR v. FE and BR v. BE trials, respectively). The AUC for overall appetite following consumption of the test drink was greater in the FR trial v. the BR trial (Table 2; P=0.006), and this pattern was still apparent, although it was attenuated when exercise was performed (Table 2; P=0.029). Similar patterns were shown for hunger and prospective consumption AUC and mirrored by fullness and satisfaction AUC (Table 2).

Discussion

The present study attempted to examine the cumulative effects of breakfast consumption and exercise on the metabolic and appetite responses to foods consumed later in the day and on subsequent energy and macronutrient balance. The main findings were that acute breakfast consumption is likely to reduce postprandial glycaemia and insulinaemia at rest. Acute exercise did not affect glucose tolerance when breakfast was omitted, but reduced glucose tolerance when breakfast was consumed; the pertinence of this chronically should be noted with caution, given the benefits of exercise training. Exercise in the fasted state led to a greater transitory reduction in appetite compared with exercise in the fed state. Energy and fat balance were least positive following exercise in the fasted state.

Acute breakfast consumption has been shown to improve glucose tolerance(2). The present findings in physically active males somewhat support the previous data, although the effect may be more trivial in these aerobically fit individuals, with magnitude-based inferences(30) indicating 41 and 59% likelihoods of beneficial and negligible effects, respectively, on glucose tolerance. This could be due to the fact that subjects in the present study are regular exercisers and therefore displaying better basal glucose tolerance(5). Lower fasting blood glucose concentrations (approximately 4·5 v. 4·8 mmol/l) support this proposition. Lower NEFA exposure prior to consumption of the test drink in the BR trial compared with the FR trial is a possible cause of the potential improvement in glucose tolerance, as prolonged NEFA elevations reduce insulin-stimulated glucose disposal by inhibiting insulin signalling(31). The (non-significant) increase in insulinemia and delay in peak insulin concentrations do support this proposition.

Muscle contraction stimulates insulin-independent glucose uptake(14), and thus explains why glucose uptake is augmented following an acute bout of exercise in spite of increased NEFA concentrations, which was observed in the FE and BE trials. Increased glucose uptake is a well-established observation at the muscle(14) and whole-body level(32). Thus, based on insulin clamp studies, it may seem surprising that there was no difference in glucose tolerance between the fasted rest and exercise trials, but this does, in fact,
corroborate with studies using oral glucose tolerance tests. Until now, studies in healthy participants have shown either decreases\textsuperscript{10,11,35–37} or no difference\textsuperscript{12,13,38} in glucose tolerance following acute endurance exercise. In those displaying no difference, the tests were either performed in the fasted state\textsuperscript{13,38} or glucose tolerance was assessed more than 2 h after exercise\textsuperscript{12}. The present study is the first to demonstrate that when nutrients are ingested immediately post-exercise, the effect on acute postprandial glucose kinetics may depend on the nutritional state (fasted or fed) prior to exercise. It may be the accrual of this acute effect that contributes to the attenuated improvements in glucose tolerance seen during exercise training when carbohydrate availability is high\textsuperscript{5}.

Regarding the effects of exercise when fasted, endurance exercise increases the rate of appearance of endogenous glucose\textsuperscript{37}. Therefore, the increase in muscle glucose uptake after exercise\textsuperscript{14} (affecting rate of disappearance) could ostensibly be offset by the increase in splanchnic glucose output (affecting rate of appearance) and, hence, result in an increase in flux, but there was no difference in the systemic concentrations of glucose after exercise compared with that after rest when fasted. Future studies are needed to address whether this is indeed the mechanism at play.

Food consumption prior to exercise also increases splanchnic blood flow during exercise\textsuperscript{60}. As mesenteric blood flow is positively associated with intestinal glucose absorption\textsuperscript{89}, it can be speculated that the increase in blood flow (from breakfast consumption), combined with increased passive absorption (from exercise), results in the greater peak blood glucose concentration in the BE trial compared with the FE trial. However, recent evidence associates the increase in intestinal absorption with reduced gut blood flow occurring during intense exercise and may result in intestinal damage\textsuperscript{80}, indicating faster entry of glucose into the circulation when gut blood flow is reduced (which occurs when exercising after fasting compared with after feeding\textsuperscript{60}). This adds to the confusion in the previous conjecture, as the putative increase in splanchnic blood flow in BE would result in less intestinal cell damage and reduced passive absorption, leading to a lower blood glucose AUC (assuming that endogenous glucose production and glucose disappearance remain constant, which can be presumed due to similar carbohydrate balance post-exercise and thus similar whole-body glycogen concentrations).

The present study used an exercise intensity that was lower (61 % \(\text{V}_{\text{O2peak}}\) v. 70 % of maximum power output) than that of van Wijck et al.\textsuperscript{40}. At lower intensities (55 % \(\text{V}_{\text{O2peak}}\)), the exercise-induced reduction in splanchnic blood flow is abolished\textsuperscript{60}. This makes it tempting to presume that other factors, such as heat or mechanical stresses or changes in hormone concentrations, contribute to the increase in intestinal glucose absorption following exercise\textsuperscript{41}. Another factor at play could be reductions in insulin sensitivity of non-exercised (upper limb) muscle following exercise\textsuperscript{12}. Clearly, this area has great scope for future work, pertinent to the understanding of the impact of food intake and exercise on subsequent whole-body glucose tolerance.

The AUC\textsubscript{INS/GLU} was lower in both exercise trials compared with the FR trial, whereas IS\textsubscript{Matsuda} was similar between trials, suggesting that postprandial insulin secretion is reduced immediately following exercise, but insulin sensitivity is unaffected\textsuperscript{28,29}. This strengthens the assumption that the change in glucose kinetics seen in the present study is due to a difference in the glucose rate of appearance.

The finding that GLP-1\textsubscript{7–36} concentrations were not different between trials is in accordance with the proposition that glucose entered the circulation via passive absorption. Intravenous infusion of glucose mirroring the plasma glucose profile to oral ingestion does not augment GLP-1 concentrations\textsuperscript{43}. Therefore, as GLP-1\textsubscript{7–36} concentrations were not different between trials, this provides support for elevated glucose appearance from passive absorption, as greater GLP-1\textsubscript{7–36} secretion would not occur. GLP-1\textsubscript{7–36} is also a potent incretin
Integrated effects of breakfast and exercise

Fig. 8. Overall appetite. Overall appetite sensations during (A) the breakfast postprandial and exercise (EX) periods and (B) following test drink consumption in the overnight fast followed by rest without breakfast (FR, ◦), overnight fast followed by breakfast and rest (BR, ◌), overnight fast followed by EX without breakfast (FE, □) and overnight fast followed by breakfast and EX (BE, ▲) trials. BL, baseline; DE, during EX; EE, end of EX; PL, post-lunch.

Overall appetite (mm)

0 20 40 60 80 100

Overall appetite (mm)

0 20 40 60 80 100

Time (min)

Time post-drink (min)

hormone, stimulating insulin secretion and also suppressing appetite(39). Thus, as GLP-1<sub>7–36</sub> concentration did not differ between trials, it would seem that other factors are playing a role in enhanced insulin action and appetite suppression with breakfast consumption. It should be noted that GLP-1<sub>7–36</sub> may interact with neurons expressed locally in L-cells, prior to being rapidly degraded on entry into the circulation, where its clearance can exceed cardiac output by two to three times(44). Hence, GLP-1<sub>7–36</sub> can still influence appetite in spite of no detectable rise in its plasma concentrations.

There was evidence of delayed suppression of NEFA following consumption of the test drink in the FR trial compared with the BR trial, suggestive of metabolic inflexibility, again associated with insulin resistance. Exercise uncoupled the link between breakfast, NEFA and insulin concentrations, whereby, in both the FE and BE trials, insulin and NEFA concentrations were similar prior to and following consumption of the test drink. Increased NEFA availability during and following exercise is required to support higher rates of fat oxidation by skeletal muscle as carbohydrate is used to replenish glycogen stores(11). As such, NEFA flux is raised, and, as insulin-resisting effects of NEFA on muscle seem to be time dependent(31), turnover may be more important than NEFA concentrations for insulin sensitivity.

Exercise transiently suppressed hunger and overall appetite. This is a common phenomenon(15,16,45), yet less is known about the effect of nutritional status on the ability of exercise to influence appetite. The present study found that, compared with rest, exercise suppressed hunger and overall appetite to a greater extent when fasted compared with the fed state (approximately 17% v. 9%, respectively). Nevertheless, it should be noted that appetite was higher in the fasting state prior to exercise. To our knowledge, this is the first crossover study to demonstrate the effect of exercise in fasted and fed conditions on appetite sensations compared with resting trials in the equivalent nutritional state.

Harmonious with preceding research(15,16), the exercise-induced suppression of appetite was abolished within 30 min of exercise termination and appetite was subsequently similar between exercise and rest trials until lunch. Breakfast consumption, however, reduced overall appetite following test drink consumption by approximately 17 and 14% in the rest and exercise trials, respectively. Despite a 10% reduction in appetite ratings with breakfast consumption, no detectable difference in energy intake between trials was observed at lunch. This occurred regardless of the additional 1859 kJ of energy consumed during breakfast and approximately 2423 kJ of energy expended during exercise. Subsequently, energy intake was higher in breakfast trials. Observational data corroborate the present findings with daily energy intake increase in regular breakfast consumers compared with ommitters(41). Yet, when BMI was measured, it was still inversely associated with breakfast consumption(41), suggesting that it may be the increased energy expenditure and the improved metabolic responses to food consumption that result in better weight maintenance.

The outcome that exercise did not influence subsequent energy intake is in accord with most of the prior research in this area, although some have found an increase in immediate energy intake(46). It may be that individual variation exists, whereby some individuals’ drive to eat following exercise is dominated by hedonic processes(47). This leads to a divergence between those who compensate for extra energy expenditure by increasing intake and non-compensators, who fail to increase intake in the face of an increase in expenditure. In the present study, the range of compensation for exercise-induced energy expenditure was large (5665 kJ of energy separated the individual who over-compensated and the individual who under-compensated the greatest). This variation in the compensation of energy expenditure is likely to account for the variation seen in body fat changes in an exercise intervention (reviewed by Caudwell et al.(48)). It is interesting to note that there was no significant relationship between the degree of compensation to exercise on fasted trials and breakfast trials, suggesting that those who over-compensate during exercise in one nutritional state (i.e. the
fasted/fed state) may not over-compensate in the opposing circumstance. Another possibility is that exercise energy expenditure is gradually compensated for by energy intake, which is likely to require a period of several weeks, and even then is not likely to be fully compensated for(49).

The higher total energy intake with breakfast trials and the exercise-induced energy expenditure led to energy balance being most positive in the BR trial and least positive in the FE trial. BE resulted in an approximately 1110 kJ reduction in energy balance compared with FR. When taken in concert with the similar appetite sensations to resting trials, exercise may provide a more attractive option for restricting energy availability compared with omitting breakfast. Interestingly, in spite of differing quantities of carbohydrate and fat oxidised with all trials, carbohydrate balance was remarkably similar between the FE and BE trials, whereas fat balance was 3-fold more positive in the BE trial. This may not be as clear at rest, as the difference between the FR and BR trials in carbohydrate balance did approach a statistically significant difference (Fig. 7), but was higher than that in exercise trials. At least in the short term, the regulation of carbohydrate stores is more tightly regulated than fat stores(50). The findings of the present study add that consumption/omission of breakfast will not alter carbohydrate balance, whereas exercise can reduce carbohydrate balance.

The increased energy expenditure observed during exercise with breakfast consumption was provided by a higher rate of carbohydrate oxidation, this has previously been reported(50–52) and may be magnified during running due to the weight-bearing component(53). The relevance of this with respect to energy balance was, however, trivial, as energy balance was lower in the FE trial compared with the BE trial.

This controlled experimental study involved the provision of a popular breakfast food consumed prior to a bout of exercise or rest in physically active males, with a structure similar to the eating patterns in Western society. It could be viewed that a caveat with the present study is that the participants were physically active and that a sedentary population would benefit more from exercise/diet-induced improvements in metabolism and appetite. However, those who regularly exercise still utilise energy/carbohydrate restriction in order to regulate body composition(40). Therefore, the results are pertinent to these populations; yet, it would undoubtedly be of virtue to investigate these responses in other populations (females, sedentary and obese) to extrapolate findings to a wider population. Moreover, future work should examine whether there is a difference in energy intake in subsequently consumed meals over a longer duration.

It is also of merit to recognise that the environmental conditions were similar between trials, which is important due to the potential effect of temperature on appetite and energy intake(46).

The findings of the present investigation suggest that in an acute setting, energy intake from breakfast and energy expenditure from exercise are not compensated for at lunch. Consequently, energy balance was most positive following breakfast and rest and least positive following breakfast omission and exercise. When exercise is performed, it may be more pertinent to omit breakfast if a negative fat balance is desirable, although the findings of the present study are unable to predict the longer-term outcomes of energy and fat balance due to the single-meal design, and as such this conclusion should be interpreted with caution.

The present study aimed to explore the effect of breakfast and exercise on the metabolic and appetite responses to subsequent food consumption. The findings indicate that breakfast ingestion may improve the metabolic and appetite responses to subsequently consumed foods when sedentary. When breakfast is consumed, subsequent postprandial glycaemia is higher following exercise, yet care should be taken during the interpretation for chronic effects, as exercise training almost always confers a benefit for glucose tolerance and insulin sensitivity. Exercise also resulted in an ephemeral reduction in appetite, which is greater when performed fasted.

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