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A framework for periodized nutrition for athletics

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Abstract

Over the last decade, in support of training periodization there has been an emergence around the concept of nutritional periodization. Within athletics (track and field), the science and art of periodization is a cornerstone concept with recent commentaries emphasizing the underappreciated complexity associated with predictable performance on demand. Nevertheless, there are numerous training, with varying levels of evidence, that feature sport and event specific purposeful sequencing of different training units [long (macro-cycle; months), medium (meso-cycle; weeks) and short (micro-cycle; days and within-day duration)] training cycles and sessions. Indeed, implementation of strategic temporal nutrition interventions (macro, meso and micro) can support and enhance training prescription and adaptation, as well as acute event specific performance. However, a general framework on how, why and when nutritional periodization could be implemented has not yet been established. It is beyond the scope of this review to highlight every potential nutritional periodization application. Instead, this review will focus on a generalized framework, with specific examples of macro-, meso- and micro-periodization for the macronutrients of carbohydrate (CHO), and by extension fat. More specifically, we establish the evidence and rationale for situations of acute high CHO availability, as well as the evidence for more chronic manipulation of CHO coupled with training. The topic of periodized nutrition has made considerable gains over the last decade, but is ripe for further scientific progress and field application.

Key words: performance, track and field, nutrient timing, macro, meso, micro

Introduction

The concept and underpinnings of periodization are deeply rooted in the history of athletics (track and field). Indeed, the seminal scientists and coaches who developed the principles of periodization include Dr. Hans Selye with his General Adaptation Syndrome model (Selye, 1950), followed by Matveyev, Bonderchuck and Bompa (contributions reviewed by (Issurin, 2010)). Numerous approaches to periodization have emerged from this work, including classical, block, polarized and complex models (Issurin, 2010; Kiely, 2012). Within this diversity is a central theme: the purposeful sequencing of different training units [long (macro-cycle; months), medium (meso-cycle; weeks) and short (micro-cycle; days and within-day duration)] so that athletes can attain the desired readiness to perform optimally for targeted events on demand (Stone et al., 1981). However, it is now appreciated that the chaos and complexities of the individual, exposed to various stimuli (physical, emotional, genetic), are probably much more complicated than most periodization purists would want to admit (Kiely, 2018). Furthermore, the impact of nutrition on training adaptation and performance needs to be recognized.

The 2007 International Association of Athletics Federations (IAAF) Nutrition Consensus presented the first formal opportunity to provide theoretical guidelines for nutrition periodization, with suggestions of the approximate energetic and macronutrient demands of different training phases within a yearly periodized training plan (Stellingwerff et al., 2007). Recently, periodized nutrition has been defined as: “the planned, purposeful, and strategic use of specific nutritional interventions to enhance the adaptations targeted by individual exercise sessions or periodic training plans, or to obtain other effects that will enhance performance in the longer term (Jeukendrup, 2017a).” Athletics might be considered the sport most suited to the application of periodized nutrition, given the diversity of the bioenergetic and biomechanical demands of the numerous different events. By extension, given the unique demands of all athletic disciplines and events, it is beyond the scope of this review to highlight every potential nutrition periodization approach. Instead, we will focus on a general framework

highlighting the various considerations in the implementation of periodized nutrition. This review will then highlight an emerging nutritional periodization concept around the various approaches to carbohydrate (CHO) and fat periodization; from a macro-cycle [weeks to months; e.g. chronic low-CHO high-fat (LCHF)] to meso-cycle (several days to weekly; e.g. high dietary to upregulate CHO oxidation) and micro-cycle CHO periodization approaches (within day and between training session; e.g. acute CHO-manipulation).

A framework for periodization of nutrition

The godfather of the American fitness industry Jack Lalanne has often remarked, “Exercise is King while nutrition is Queen; put them together and you have a Kingdom!” Of course, nutrition strategies play a supportive role in enhancing acute training stimuli into optimal training adaptation. For example, optimizing protein (PRO) intake, PRO quality and timing achieves only a minor adaptive hypertrophic response without the potent stimulus of resistance exercise. However, nutrition serves an important function given that elite athletes have 1400-1800 eating occasions per year, while training 300-800 times. Accordingly, figure 1 highlights a theoretical framework that is fundamental for the prioritization and optimization of nutrition periodization that practitioners can apply to a myriad of nutrition interventions. First, the coach and the entire support staff should have a full and comprehensive understanding of the event-specific physiological, neuromuscular, structural and psychological determinants for success. Second, the performance gaps of the individual athlete should be quantified as best as possible against these performance determinants. From this construct, the coach will strategically develop the various macro- (months to years), meso- (weeks to months) and micro-cycles (days to within days) aspects of training periodization and its specific sessions, using these as to bridge between the goals and the gap. Indeed, the integration of peer-reviewed evidence from training studies with the tacit knowledge of elite coaches (Nash et al., 2006) produces the “Science” and “Art” of elite training prescription and periodization (Kiely, 2012). The periodized training schedule provides a framework for the sports nutrition professional to match nutritional strategies to support training outcomes. Table

1 highlights the rationale involved in implementing macro-, meso- and micro-periodized nutrition recommendations. Conversely, figure 1 outlines the process, and highlights several periodized nutrition examples, such as the macronutrient (CHO and PRO), micronutrient (iron) and ergogenic aid (creatine) examples of macro-, meso- and micro-periodization, respectively. However, the narrative of this review will exclusively focus on energy and CHO periodization.

Periodization of energy intake

Energy intake (EI) is a primary nutritional characteristic since it: 1) establishes the baseline from which intakes of the macronutrients (including muscle substrates) are derived; 2) influences the capacity of the diet to achieve micronutrient targets within nutrient-density constraints; and 3) allows the manipulation of physique via the interaction of training and energy balance. Given that the energy cost of training is often the major determinant of an athlete's energy expenditure, and that the training load varies markedly across micro-, meso- and macro-cycles of the training plan, energy intake should also vary between days, weeks and training phases. In addition, some athletes may desire to manipulate their EI in attempts to alter physical and structural characteristics (e.g. reduce body fat or body mass and/or gain muscle mass), but must appreciate the risks versus rewards of such manipulations. Accordingly, energy manipulations should be strategically integrated into the annual plan to minimize the effects on training quality or competition performance, and strategies should aim to maintain sufficient energy availability (EA) to reduce the acute and chronic issues associated with the development of Relative Energy Deficiency in Sport [RED-S; (Mountjoy et al., 2018)]. For a specific review of low energy availability and RED-S in track and field athletes, the reader is referred to review by Melin et al. in the current series (Melin et al., 2018).

To undertake periodization of energy intake, the nutrition professional and coach need to undertake a full audit of step #3 (within Figure 1) of our framework through a nutrition lens. This assessment should include estimations of total caloric and substrate utilization for each unique type of workout, which then influences the fluctuations or manipulations of energy and

macronutrient intake that needs to be integrated into daily eating patterns, as well as an appreciation of these elements over a given training meso- and macro-cycle. Many indirect protocols can be implemented to provide information on EEE and fuel use; these include indirect calorimetry to estimate fuel utilization linked to internal load (e.g. heart rate) and/or external load (e.g. running speed) and various technologies that can estimate caloric expenditures. It should be stressed, however, that accurate measurements of EI (Larson-Meyer et al., 2018) and the energy expenditure of exercise (EEE) and activities of daily living (Murakami et al., 2016) are extremely difficult to make; indeed, daily mismatches of as little as 300 kcal [1200 kJ; (Torstveit et al., 2018)], which are within the typical errors of measurement of both EI and energy expenditure, can have profound effects on EA over the long term. In light of current caution around the measurement or prescription of “optimal EA”, it is both unwise and impractical to suggest that athletes should chronically “micromanage” the monitoring and manipulation of EA or EI within training or competition environments. However, having an understanding of general needs and how they fluctuate can allow the athlete to develop behavioral practices that allow EI to track with EEE (e.g. to consume recovery snacks, or within session energy intake, on days or during periods of higher EEE), and perhaps, to focus on energy goals during targeted periods of body composition manipulation. The emerging concept of periodization of body composition allows characteristics to be manipulated within an individualized range across different phases of the annual plan according to short- and long-term issues of training adaptation, health and performance (Heydenreich et al., 2017; Stellingwerff, 2018). A case study of a 9-year history of body composition management of an elite female middle-distance runner (Stellingwerff, 2018) illustrates the gradual progression to optimal physique over the athlete’s career as well as the manipulation within a season.

Carbohydrate and fat periodization

A key goal of training is to enhance the various metabolic pathways to improve the capacity and/or rate of ATP production, particularly to reduce or delay the onset of factors that may make these pathways limiting for competition performance. Some of the principles by which these metabolic pathways can be enhanced include increasing the size of the storage and availability of endogenous fuels (capacity) and enhancing the body's ability to make use of exogenous fuels consumed just prior to and during exercise (power). Further metabolic enhancement can also occur by improving the delivery of nutrients and oxygen to the working muscle, reducing the accumulation of by-products that might disturb cellular homeostasis or metabolic regulation, or improving the efficiency (economy) of these pathways to produce ATP across a range of absolute and relative exercise intensities. Carefully organized dietary interventions to enhance the exercise-nutrient interactions can augment the role that training alone already provides. The availability of modern laboratory technologies to investigate cellular signaling events over the past decade has expanded insights into the role of nutritional support in promoting adaptations to exercise. It is now known that many substrates, and in particular muscle glycogen and plasma free fatty acids, act not only as fuels for the exercise bout being undertaken, but also as regulators of the cellular and whole body adaptation to exercise, and specifically endurance exercise (Hansen et al., 2005; Hulston et al., 2010; Morton et al., 2009; Yeo et al., 2008). In relation to this, the reader is directed to a recent review in which a range of acute and chronically applied strategies to manipulate fat and CHO availability are defined and explained, at least in relation to endurance events, to address some confusion over terminology, application and theoretical basis (Burke et al., 2018a). This body of work was initially termed the “train-low (smart): compete high” paradigm and involved the integration of reduced CHO availability for carefully chosen training sessions to enhance markers of training adaptation, while approaching competition with high CHO availability to maximize performance and recovery (Burke, 2010). More recently, the concept of periodization of CHO availability has been explained using the theoretical model of “fueling for

the work required” whereby CHO availability is adjusted in accordance with the demands and goals of the specific training session to be completed (Impey et al., 2018). With this approach, total daily CHO intake and its distribution over the day can be modified day-by-day and meal-by-meal (i.e. within the micro-cycle) to manipulate CHO availability for each exercise occasion.

Strategies to promote CHO utilization during exercise

Carbohydrate, from muscle glycogen stores and plasma glucose, provides an effective and dominant fuel source for performance across a wide variety of events in athletics (Hawley et al., 2015). Indeed, it has been known for nearly a century that 100% CHO produces ~5.5% more ATP per liter of oxygen consumed than compared to 100% fat oxidation (Krogh et al., 1920). Therefore, the achievement of high CHO availability (defined as CHO stores available to provide the substrate needs of an event or training session) is a key goal of competition nutrition and the reader is directed to reviews of the individualized competition needs across events in track and field (Burke et al., 2018b; R. J. Costa et al., 2018; Slater et al., 2018; Stellingwerff et al., 2018; Sygo et al., 2018). In addition to promoting training quality, undertaking sessions with high CHO availability can enhance the pathways of oxygen-independent glycolysis and CHO oxidation (Cox et al., 2010).

Promoting training adaptation via low CHO availability

The rationale for deliberately promoting low CHO availability in relation to a training session is that commencing and/or recovering from a proportion of training sessions with reduced CHO availability activates acute cell signaling pathways that increase many of the hallmark muscle adaptations to endurance training such as mitochondrial biogenesis, angiogenesis and increased lipid oxidation (Figure 2). The landmark studies in the field have typically manipulated pre-exercise muscle glycogen availability (Hansen et al., 2005; Hulston et al., 2010; Morton et al., 2009; Yeo et al., 2008), given the rationale that glycogen concentration is a potent regulator of key cell signalling kinases (e.g. AMPK, p38), transcription factors (e.g. p53, PPAR) and transcriptional co-activators e.g. PGC-1 α , (Bartlett et al., 2013; Hearn et al., 2018; Psilander et al., 2013; Yeo et al., 2010). However, it is now recognized

that practical models of CHO periodization must extend beyond just manipulating pre-exercise muscle glycogen availability. Accordingly, CHO availability is defined as the sum of the current individual endogenous (i.e. muscle and liver glycogen) and exogenous CHO (i.e. CHO consumed before and/or during exercise) that is available to sustain the *required* training or racing intensity and duration (Impey et al., 2018). According to this definition, it is possible to have insufficient CHO availability (even if exercise is commenced with high pre-exercise muscle glycogen stores) if an inadequate dose of exogenous CHO is consumed during prolonged exercise to sustain the desired intensity (Coyle et al., 1986). Alternatively, it is possible to commence exercise with reduced muscle glycogen, but can still be considered to have sufficient CHO availability if the exogenous CHO consumed during exercise permits the completion of the desired training intensity and duration (Widrick et al., 1993).

The various main approaches to manipulating CHO availability in relation to training are presented in Table 2. Essentially, it is now recognized that manipulation both endogenous and exogenous CHO availability before (Hansen et al., 2005; Yeo et al., 2008), during (Akerstrom et al., 2006; Morton et al., 2009) and after training sessions (Marquet et al., 2016; Pilegaard et al., 2005) may lead to an augmented training response; at least in terms of acute (e.g. cell signalling) and chronic (e.g. changes in enzyme protein content and activity) skeletal muscle metabolic adaptations (Figure 2). Despite the theoretical rationale for incorporating a periodization of CHO availability into an elite athlete’s training program, balancing the risk versus reward can be challenging for a number of reasons. First, the majority of studies to date have been conducted using cycling-based or knee extensor exercise models [78% of 27 studies, (Impey et al., 2018)]. These modalities may more naturally lend themselves to train-low sessions given the non-weight bearing activity and lower eccentric loading compared with running. Second, the CHO requirements of the typical training sessions undertaken by elite track and field athletes are not well known and practitioners must use theoretical knowledge of glycogen utilization from relevant laboratory and field based studies to organize appropriate manipulation of training and diet. A recent meta-analysis which provides more information in

this regard, clearly highlights the impact of intensity and duration on glycogen use (Areta et al., 2018), but practitioners need to consider the use of the internal and external training load metrics outlined in Figure 3, to make best estimates. Of particular note to track and field athletes are the negative effects of reduced CHO (and energy) availability on markers of bone turnover (Sale et al., 2015), which contrast with the benefits of immediate post-exercise feeding of CHO and PRO (Townsend et al., 2017). Although this needs to be confirmed in a longitudinal model, prolonged periods of training with reduced CHO availability may potentially increase the risk for stress fractures, in runners who are already at risk for RED-S and stress fractures (Heikura et al., 2017). Indeed, low CHO availability may mediate disturbed reproductive function in its own right (Loucks, 2014), creating another pathway to explain the increase in musculoskeletal injuries seen with low energy availability (Rauh et al., 2014). Similarly, reduced CHO availability during training may lead to increased susceptibility to illness owing to the role of CHO in modulating post-exercise immune responses (R. J. S. Costa et al., 2005). Finally, evidence that the improved metabolic adaptations associated with periodization of low CHO availability within an athlete’s training program actually translates to improved performance outcomes is equivocal. Indeed, Impey et al. (2018) reported that, although periodically completing endurance-training sessions (e.g. 30-50% of training sessions) with reduced CHO availability modulates the activation of acute cell signalling pathways (73% of n=11 studies) and oxidative skeletal muscle adaptations (78% of n=9), only 37% of studies demonstrate improvements in performance.

The development of practical approaches to sport-specific CHO periodization requires consideration of the individual athlete’s training (e.g. intensity, duration, body composition, altitude, ambient temperature etc) and competitive goals (e.g. time required to attain and taper for peak performance) within the specific micro-, meso- and macro-cycle. An illustration of the implementation of periodized CHO availability in the real world was provided by a 16-week case study of three elite marathon runners (Stellingwerff, 2012). These athletes undertook a weekly average of 2.5 sessions involving low CHO availability during the first 12 weeks.

However, during the subsequent 4 week period which focused on competition preparation, nutritional strategies shifted towards an increased frequency of practicing CHO fueling during training sessions (2.5 sessions per week), coupled with a reduction in low CHO training sessions (Stellingwerff, 2012). In this study, with regard to training with low CHO availability, fasted training was reported to be physiologically and psychologically “easier” to integrate than the more strenuous low muscle glycogen training sessions. Regardless of the specific approach, it is intuitive to avoid chronic periods (>3-5 days), or consecutive performance, of the various “train-low” modalities due to the potential negative effects of both low CHO and inadequate total energy availability on training quality, competitive performance and health outcomes (Burke et al., 2017; Havemann et al., 2006; Mujika, 2018). Rather, careful day-to-day periodization is likely to maintain metabolic flexibility and still allow the completion of high-intensity and prolonged duration workloads on heavy training days. Intuitively, train-low may be best left to those training sessions that are not as CHO dependent and where the intensity and duration is not likely to be compromised by reduced CHO availability (e.g. steady-state type training sessions performed at intensities below the lactate threshold); however significant work remains on how to best periodize CHO with the numerous training permutations (Table 2). As with all nutritional strategies, application of CHO periodization should only be done in conjunction with appropriately qualified nutrition professionals and be continually refined and optimized.

Fat: a relatively unlimited fuel source and driver of adaptation

Although endurance-trained athletes have an enhanced capacity for fat oxidation compared to untrained, this adaptation is clearly not maximized from training alone since it can be doubled, or even tripled, by chronic adaptation to a LCHF diet (Burke et al., 2002; Burke et al., 2017; Carey et al., 2001). Various models have been used to achieve these high fat oxidation rates including non-ketogenic CHO-restricted diet (NK-LCHF; typically 65% energy as fat and <20% energy from CHO to prevent ketosis while reducing CHO intake to levels below the fuel costs of daily training) and the more restrictive ketogenic LCHF diet [K-

LCHF: typically <50 g/d CHO and 75-80% fat; (Burke et al., 2018a)]. However, studies have shown that as little as 5 days of exposure to HFLC diets, while continuing to undertake both high volume and intensity of training, achieves a robust retooling of the muscle to increase intramuscular triglycerides stores, and enhance the mobilization, transport, uptake and oxidation of fats (for review see Burke, 2015).

By itself, however, chronic adaptation to a NK-LCHF does not translate into clear improvements in endurance performance, except in specific scenarios or individuals (Burke, 2015). Therefore, a micro-periodization protocol which (theoretically) could sequentially enhance the capacity of both fat- and CHO-based fuel oxidation prior to an endurance event has been proposed; this involves a 5-6 d fat adaptation phase followed by acute restoration of CHO availability just prior to and during the endurance task (Burke et al., 2002; Carey et al., 2001; Havemann et al., 2006). Investigation of this protocol showed that the muscle retooling achieved by the fat-adaptation phase is robust in continuing to promote higher rates of fat oxidation in the face of aggressive strategies to restore high CHO availability during the endurance task (Burke et al., 2002; Carey et al., 2001). Despite substantially reduced rates of muscle glycogen utilization during the early part of the exercise task there was no clear enhancement of a subsequent time trial in any of these studies. One apparent explanation for this outcome is that, rather than sparing glycogen utilization, chronic exposure to a high fat diet causes an impairment of CHO oxidation during exercise due to a reduction on glycogenolysis and a downregulation of the activity of the pyruvate dehydrogenase complex [PDH; (Stellingwerff et al., 2006)]. Data from rodent models suggests that such inhibition of PDH activity may actually be due to the effects of reduced muscle glycogen availability. Indeed, commencing exercise with reduced muscle glycogen augments the activation of peroxisome proliferator activated receptor- δ (PPAR- δ) (Philp et al., 2013), a transcription factor that up-regulates pyruvate dehydrogenase kinase 4 (PDK4) that, in turn, phosphorylates and inactivates PDH. In human muscle, however, we have recently shown that the down-regulation of CHO metabolism appears to be a consequence of the high-fat diet and increased

fat availability rather than CHO restriction alone (Leckey et al., 2018). The consequences of reduced CHO utilization within the tricarboxylic acid cycle are likely to manifest in a reduced capacity for ATP production at high intensities. For example, in a study in which the fat adaptation-CHO restoration model was undertaken prior to a 100 km cycling time trial, there was a trend towards impairment of overall performance, but a significant reduction in ability to complete 1km and 4km sprints undertaken at ~90% peak power output interspersed within the protocol (Havemann et al., 2006). This reduction in power output matches the ~30% reduction in PDH and estimated glycogenolysis during a 1min sprint at 150% of peak power output found by Stellingwerff et al. (2006).

The proposed benefits of chronic adaptation to ketogenic LCHF diets has also received recent attention in both lay (Brukner, 2013) and scientific press (Noakes et al., 2014; Volek et al., 2014) due to the proposed benefits of increased fat oxidation and exposure to high levels of circulating ketone bodies. Indeed, an early study featuring 4 weeks of LCHF adaptation to such CHO restriction showed the muscle's plasticity in being able to increase fat utilization to maintain exercise capacity at modest power outputs (~60% VO₂max; (Phinney et al., 1983)). However, Phinney et al. already noted that this feat was achieved in the face of impairments of capacity to undertake exercise at higher intensities. A more recent investigation of 3.5 weeks of exposure to a K-LCHF diet in elite race walkers demonstrated an apparent rationale for Phinney et al.'s findings (Burke et al., 2017). In this study, some of the highest rates for fat oxidation ever reported in the literature were achieved following adaptation to the K-LCHF diet, under conditions of either fasting or intake of fat during a prolonged exercise protocol. However, this was associated with an increase in the oxygen cost (i.e. reduced economy) of walking at a range of speeds relevant to the competitive events in this discipline (20 and 50 km), and the K-LCHF diet did not improve 10,000m performance compared to the CHO-supported dietary groups (Burke et al., 2017).

Taken together, chronic LCHF interventions have been shown to 1) decrease CHO oxidation (Burke et al., 2002; Carey et al., 2001; Havemann et al., 2006) via decreased

glycogen utilization and PDH activation (Stellingwerff et al., 2006); 2) result in decreased exercise economy due to lower ATP production associated with increased fat oxidation (Burke et al., 2017); 3) and probably cause a decrease of the intestinal glucose transport proteins (SGLT-1) reducing the capacity for intestinal absorption. This would decrease the effectiveness of CHO feeding strategies, thus increasing the risk of gut disturbances (Jeukendrup, 2017b). It is important that coaches and athletes understand the metabolic demands and limiting factors in their events (Figure 1), and trial specific interventions on an individual level (Figure 3), and realize that all sustained majority of track and field events are exceptionally CHO dependent. Indeed, there is continued interest in *increasing* rather than *decreasing* CHO dependence to improve performance of events of sustained high-intensity exercise at intensities around the “lactate threshold” (e.g. the marathon) to take advantage of the greater economy of ATP production from the oxidation of this fuel (Burke et al., 2018b).

Conclusions & Future Directions

The numerous various events in athletics, all with unique bioenergetics, biomechanical and structural performance determinants, lend themselves to endless permutations of potential periodized nutrition interventions. Therefore, this review took a holistic approach to develop a nutrition periodization framework to guide practitioners in the field across these nutrition periodization options, while grounding them in targeted intervention first principles (Figure 1 & Table 1). From this framework the literature has been examined regarding macro, meso and micro-periodization of CHO availability, and consequently, fat periodization (Table 2 & Figure 2), with considerations and recommendations to individualize and test in the field, as required (Figure 3). Obviously, this paper just focused on an example of nutrition periodization by examining the research in the macro- meso- and/or micro-manipulating of CHO, and by extension. However, Figure 1 highlights various other nutrition examples of periodization that are beyond the scope of this review, but we would highlight a few other papers in this series that also feature periodized nutrition examples for protein (Phillips et al., 2018), supplements (Peeling et al., 2018), CHO fueling (Burke et al., 2018b) and middle-

distance athletes (Stellingwerff et al., 2018). With an appreciation of the complexities of training periodization, future directions in this field should include better quantification of knowledge and application of existing periodized approaches in elite athletes, as well as systematically controlled CHO periodization approaches over prolonged training blocks in larger cohorts of athletes. Other scientific areas for future investigation also include the meso and/or macro-periodization of ergogenic aids in attempts to enhance training adaptations and performance. In summary, the field of periodized nutrition has made considerable gains over the last decade, but is ripe for further progress.

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Figure 1. A methodological framework required for successful nutrition periodization, including examples of macro- (months), meso- (weeks) and micro-cycle (days / within day) nutrition periodization interventions for carbohydrate (CHO), protein (PRO), iron and creatine.

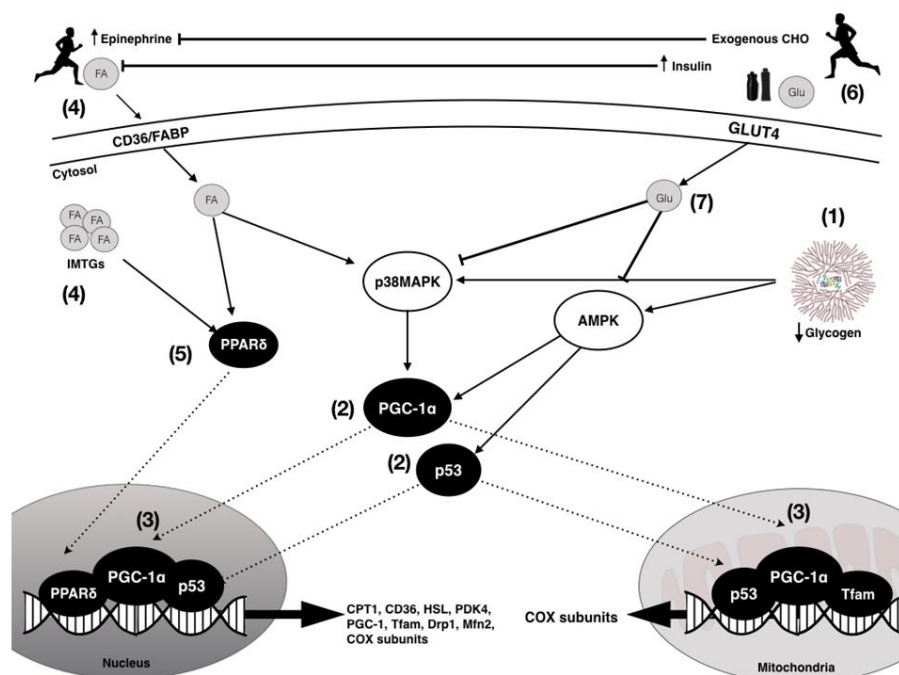


Figure 2. Schematic overview of the potential cell signalling pathways regulating the enhanced mitochondrial adaptations associated with training with low CHO availability. **1)** Reduced muscle glycogen enhances both AMPK and p38MAPK phosphorylation that results in **2)** activation and translocation of PGC-1 α and p53 to the mitochondria and nucleus. **3)** Upon entry into the nucleus, PGC-1 α co-activates additional transcription factors, (i.e. NRF1/2) to increase the expression of COX subunits and Tfam as well as auto-regulating its own expression. In the mitochondria, PGC-1 α co-activates Tfam to coordinate regulation of mtDNA and induces expression of key mitochondrial proteins of the electron transport chain, e.g. COX subunits. Similar to PGC-1 α , p53 also translocates to the mitochondria to modulate Tfam activity and mtDNA expression and to the nucleus where it functions to increase expression of proteins involved in mitochondrial fission and fusion (DRP-1 and MFN-2) and electron transport chain protein proteins. **4)** Exercising in conditions of reduced CHO availability increases adipose tissue and intramuscular lipolysis via increased circulating adrenaline concentrations. **5)** The resulting elevation in FFA activates the nuclear transcription factor, PPAR δ to increase expression of proteins involved in lipid metabolism such as CPT-1, PDK4, CD36 and HSL. **6)** However, consuming pre-exercise meals rich in CHO and/or CHO during exercise increases insulin, which can down-regulate lipolysis thereby negating FFA mediated signalling. **7)** Additionally, CHO intake before or during exercise can also reduce both AMPK and p38MAPK activity thus having negative implications for downstream regulators.

Abbreviations: AMPK; AMP-activated protein kinase, CHO; carbohydrate, COX; cytochrome c oxidase, CPT1; carnitine palmitoyltransferase 1, Drp1; dynamin-related protein 1, FA; fatty acid, FABP; fatty acid binding protein, GLUT4; glucose, HSL; hormone sensitive lipase, IMTG; intramuscular triglycerides, Mfn2; mitofusion-2, p38; mitogen-activated protein kinase, PDK4; pyruvate dehydrogenase kinase 4, PGC-1 α ; peroxisome proliferator-activated receptor gamma coactivator 1-alpha, PPAR δ ; peroxisome proliferator-activated receptor, Tfam; mitochondrial transcription factor A.

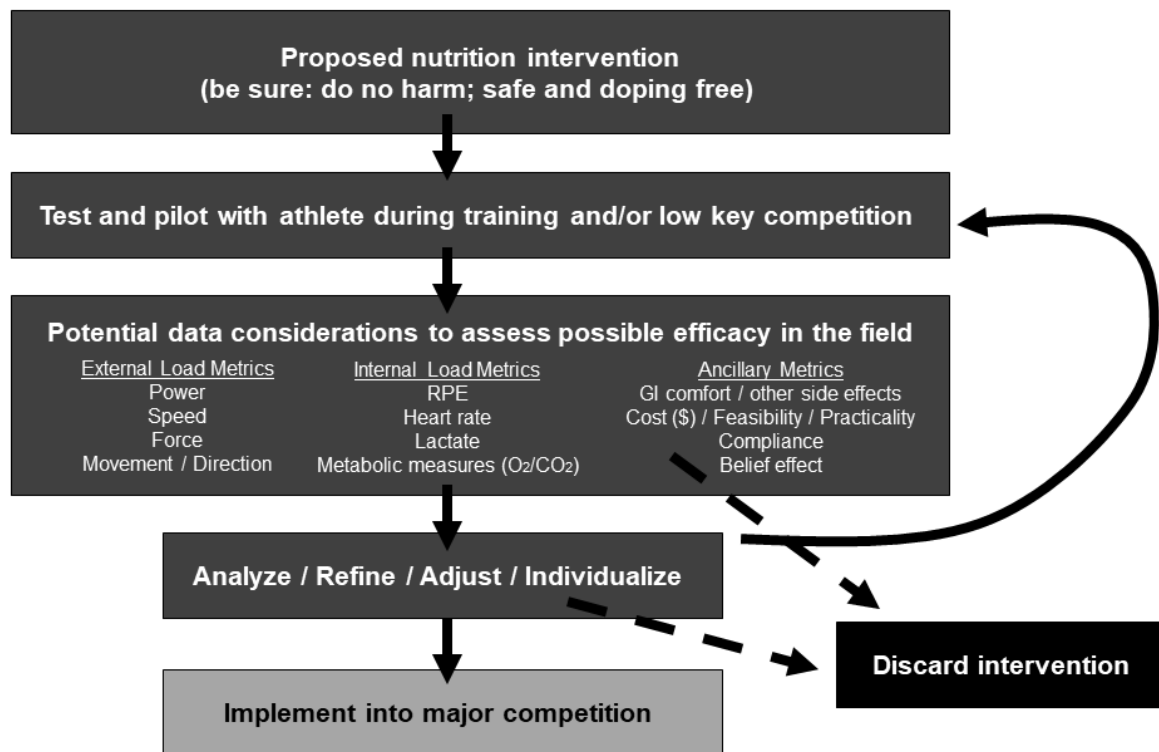


Figure 3. Supplement individualization and piloting framework, including potential data collection variables to help inform protocol and efficacy. GI, gastro-intestinal; RPE, ratings of perceived exertion.

Table 1. Nutritional considerations in relation to macro-, meso- and micro-cycle training periodization.

Performance and/or Periodization Considerations	Nutritional Context
Macro-Periodization (months to weeks)	
What is the specific load and density of this training phase/block for this macro-cycle (stimulus)?	Estimation of required endogenous / exogenous substrates required during this block should guide nutrition advice. Assessment of any nutrition ergogenic aids that synergistically match the macro-periodization.
What are the EA requirements of this macro-phase?	Ensure adequate EI for optimal EA. If required, assess RED-S status indicators as outlined by Mountjoy et al., 2018.
In relation to the training phase / block what are the current and long-term body composition goals? Are changes even necessary?	Strategic team discussions around risk & reward to optimize body composition targets, and develop an individual profile.
Are there any macro health considerations? (Injury / illness profile)	Target potential nutrition interventions to individual athlete health issues.
Meso-Periodization (weeks to days)	
What is the specific load and density of this training phase / block for this meso-cycle (stimulus)?	Estimation of required endogenous / exogenous substrates required due to the specific training stimuli during this block should guide nutrition advice. Consideration of any nutrition ergogenic aids that synergistically match the meso-periodization.
What are EA requirements of this meso-phase?	Ensure adequate EI for optimal EA. If required, assess RED-S status indicators as outlined by Mountjoy et al., 2018.
If a competition block (many competitions over several days to weeks), what are the chronic to acute recovery requirements?	During heavy competition phases, extensive logistical planning and practice is required for general, competition and recovery nutrition interventions.
What environmental training interventions are being implemented in this phase?	Environments (heat, cold, altitude) dictate implementation of various periodized nutrition interventions (e.g. hydration, iron etc.).
Micro-Periodization (days to within day)	
What is the specific load and density of this training phase/block for this micro-cycle (stimulus)?	Assessment of required endogenous / exogenous substrates required due to the specific training stimuli during this block should guide nutrition advice. Consideration of any nutrition ergogenic aids that synergistically match the micro-periodization.
What are the EA requirements of various different types of training days?	Ensure adequate EI for optimal EA, appreciating that there may be day to day EEE and EI variability.
What is the typical training day schedule? (within this specific training block/phase)	Understanding the individual athletes daily and weekly schedule informs micro (within hour) nutrition and supplement recommendations.
What are the acute recovery requirements from a unique / specific single training session?	Estimation of exogenous / endogenous substrates required during individual training sessions, coupled with the training goals and phase, will inform the pre, during and post-training nutrition interventions.
What are the acute recovery requirements from a single competition?	Generally, all recovery interventions are optimized during rounds of a competition, or throughout a competition block to maximize subsequent performance.

Performance and/or Periodization Considerations	Nutritional Context
What are the training or competition specific interventions to optimize performance? (from tapering, to warm-up to sport psychology)	Competition phase tends to offer unique nutrition periodization challenges, such as body comp optimization during tapering, optimizing recovery protocols, to acute competition specific ergogenic aids (e.g. caffeine, sodium bicarbonate etc.).

CHO, carbohydrate; EA, Energy availability = (Energy Intake (EI) - Exercise Energy Expenditure (EEE)) / Fat Free Mass;

Table 2. Overview of practical approaches to manipulate endogenous and exogenous CHO availability within CHO periodization strategies. For a summary of the terminology and rationale of different strategies of periodized fuel support for training and competition, see Burke et al., 2018a while the principles underpinning skeletal muscle adaptations, training and performance outcomes are detailed in the review of Impey et al., 2018.

Practical Approach	Practical Implementation / Theoretical Rationale	Supporting References
Train low (glycogen) session	<ul style="list-style-type: none"> Both muscle and liver glycogen are reduced during an initial morning training session. CHO intake is then withheld in recovery or sub-optimal intakes occur such that a second session is completed in the afternoon or early evening with reduced pre-exercise CHO availability. Depending on the timing of both sessions, the total time considered in a state of low CHO availability could range from 3-8 hours. Commencing exercise with low muscle glycogen stores and/or sustaining exercise intensity and/or duration to a specific level of absolute glycogen depletion is associated with the activation of key cell signaling proteins (e.g. AMPK, p38, PPAR, PGC-1α,) which achieve a coordinated up-regulation of the nuclear and mitochondrial genomes. Over a chronic training period, this may increase oxidative enzyme protein content/activity, up-regulate whole body and intramuscular lipid metabolism, with potential improvements in exercise performance and capacity. 	(Hulston et al., 2010; Morton et al., 2009; Yeo et al., 2010; Yeo et al., 2008)
Train low (fasted) session	<ul style="list-style-type: none"> Breakfast is consumed after training and no form of CHO is consumed during exercise resulting in significant elevated FFAs. This approach would predominantly target reduced liver glycogen (associated with fasting in the overnight period) though depending on the CHO intake consumed in the recovery period after the last training session, pre-training muscle glycogen may also be considered low. Exercise undertaken in fasted conditions leads to increased metabolic stress for the muscle, central nervous system and/or liver gluconeogenesis and leads to up-regulation of AMPK and signaling pathways that increase expression of proteins regulating substrate transport (e.g. GLUT 4 and CD36, FABPm, respectively) and substrate utilization (e.g. PDK4, HK, CS, β-HAD). 	(Akerstrom et al., 2006; De Bock et al., 2008; Van Proeyen et al., 2011)
Recovery low / sleep low strategy	<ul style="list-style-type: none"> Both muscle and liver glycogen are reduced during an evening training session. CHO intake is then withheld in recovery or sub-optimal intakes occur such that a second session is completed on the subsequent morning with reduced pre-exercise CHO availability. Depending on the timing of both sessions, the total time considered in a state of low CHO availability could range from 8-14 hours. Restricting CHO intake in the post-exercise period therefore maintains post-exercise muscle and liver glycogen at reduced levels as well as prolongs the duration of post-exercise elevations in circulating FFA availability. The interactive effects of changes in substrate availability may sustain the post-exercise upregulation of cell signaling (e.g. AMPK, p53, PGC-1α) pathways thus leading to increases in the adaptive response to the session. The sleep-low train low model has been associated with improved exercise performance in trained triathletes. 	(Bartlett et al., 2013; Marquet et al., 2016; Pilegaard et al., 2005)

Practical Approach	Practical Implementation / Theoretical Rationale	Supporting References
Train high (glycogen + exogenous CHO) session	<ul style="list-style-type: none"> • Training sessions deliberately commenced with high muscle and liver glycogen following optimal pre-training (e.g. CHO “loading” of 6-12 g/kg body mass, pre-training meal of 1-3 g/kg body mass) and in-training fueling (e.g. CHO consumed during exercise at a rate of 30-90 g per hour). • In this approach, promotion of high training intensity, duration, training of the gut and practicing in competition fueling are the goals. 	(R. J. S. Costa et al., 2017; Cox et al., 2010)
Amalgamation of targeted CHO availability training & recovery approaches	<ul style="list-style-type: none"> • The above approaches are deliberately amalgamated over a 24 h-48 h period. In this model, an initial train high session may be completed to promote training intensity and duration followed by a second train low session that is performed 24-48 hours later with reduced CHO availability. The second session may have arisen as a result of a combination of sleep-low, recover low and a low daily absolute CHO intake. • Using this model, CHO availability can be adjusted, before, during and/or after each training session in an attempt to take advantage of the cellular signaling responses that may occur with CHO restriction at each of these time-periods. 	(Impey et al., 2018; Stellingwerff, 2012)

CHO, carbohydrate; FFA, free fatty acids.