

# How does diurnal intermittent fasting impact sleep, daytime sleepiness, and markers of the biological clock? Current insights

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**Abstract:** Mealtimes and feeding schedules may interfere with the circadian system and impact sleep. The practice of intermittent fasting (IF) in its different formats is increasing worldwide. However, most studies addressing the effect of IF on circadian rhythms, daytime sleepiness, and sleep architecture have been conducted during diurnal IF for Ramadan. In this article, we analyze the effect of diurnal IF on the circadian clock, sleep, and daytime sleepiness. In free-living, unconstrained environments that do not control for lifestyle changes such as sleep/wake schedules, sleep duration, and light exposure, studies have demonstrated sudden and significant delays in bedtime and wake time during diurnal intermittent fasting for Ramadan. However, subsequent studies that accounted for lifestyle factors and sleep/wake patterns have reported no changes in markers of the biological clock, daytime sleepiness, or sleep parameters. Nevertheless, several researchers have demonstrated a reduction in the proportion of rapid eye movement stage sleep as the significant alteration in sleep architecture during fasting.

**Keywords:** mealtime, Ramadan, chronotype, sleep architecture, alertness, food, light, caloric restriction

## Introduction

Fasting, mealtime, and sleep may interact and impact the circadian rhythm of various body organs and cells when food is not consumed at a suitable time relative to the timing of the circadian clock of the body.<sup>1,2</sup> Intermittent fasting (IF), in which one voluntarily refrains from food intake for specific times, is an old tradition that is implemented in different practices by various societies around the world.<sup>3,4</sup> Several religions practice periods of fasting in their rituals, including Islam, Christianity, Judaism, Hinduism, and Buddhism.<sup>5</sup> Recently, IF practice has become popular, and there has been great attention to the physiological and metabolic consequences of IF among researchers.<sup>1</sup>

IF is different from caloric restriction (CR), in which caloric intake is lowered for long periods by 20%–40% but meal frequency is preserved.<sup>6</sup> Several types of IF have been described in the literature, including complete fasting every other day,<sup>7</sup> significant CR every other day,<sup>8</sup> eating only a small amount of calories (500–700) 2 consecutive days/week,<sup>9</sup> “time-restricted feeding,” where food intake is restricted to a specified time daily,<sup>10</sup> fasting 1 or 2 days per week and allowing 5–6 days ad libitum ingestion of food, and the spiritual diurnal IF practiced during Ramadan, in which there is total abstinence from food and drink during daytime (dawn to dusk).<sup>11,12</sup>

Most of the previous studies on experimental IF have assessed its effect on metabolic and cardiovascular risks.<sup>1</sup> Most studies addressing the impact of IF on sleep and circadian rhythm have been conducted during diurnal IF for Ramadan. Therefore, in this article,

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we address the impact of diurnal IF on some markers of the circadian clock, sleep parameters, and daytime sleepiness.

## IF for Ramadan

Diurnal IF for Ramadan is distinct from other forms of experimentally studied fasting for several reasons; therefore, it needs to be treated as a separate entity. Ramadan fasting comprises diurnal IF from dawn to sunset, where participants refrain from eating and drinking during the specified period. Ramadan is a month of the Arabic (*Hijri*) year and hence follows the lunar system. This implies that Ramadan comes during a different season every 9 years, which could affect the duration of daytime and hence fasting duration, where daytime is longer during the summer than in the winter.<sup>13</sup>

Moreover, diurnal IF is practiced for the full month of Ramadan; this long duration may permit greater adjustment to the fasting protocol than that typically occurs in other forms of experimental IF. Furthermore, several studies have shown that Ramadan is accompanied by lifestyle changes that may influence sleep and circadian rhythms (Figure 1).

## Review method

The literature search started on the May 1, 2018, with the following keywords: “sleep,” “sleep pattern,” “daytime sleepiness,” “polysomnography,” “sleep architecture,” “circadian rhythm,” “intermittent fasting,” and “Ramadan,” using PubMed (80 results), Clarivate (94 results), and Google Scholar (105 results). In addition, the reference lists of the

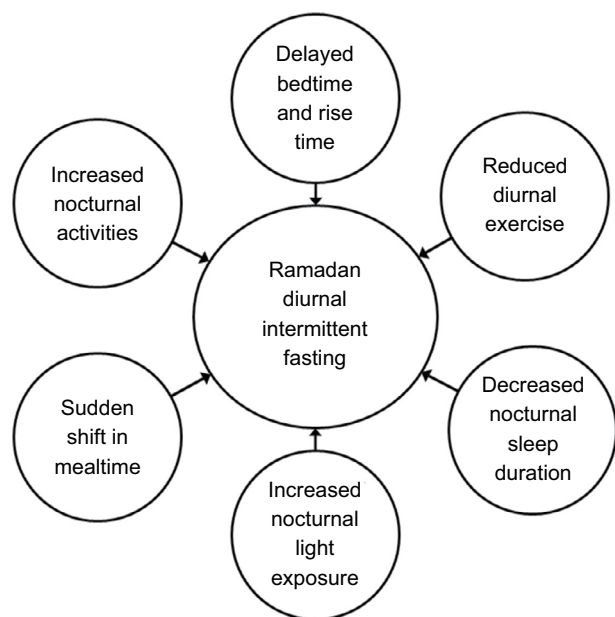
identified articles were also searched for further references. The following inclusion criteria for human studies were used: children, men, and women were accepted as participants, and both randomized controlled trials and nonrandomized trials were accepted.

## Impact of diurnal IF on the circadian rhythm

All cells of the human body have circadian clocks, which are generally classified into peripheral and central clocks based on their anatomical place. The central master clock is located in the suprachiasmatic nucleus (SCN) of the hypothalamus. On the other hand, the peripheral clocks are located in different organs and tissues.<sup>14</sup> Both central and peripheral clocks act to preserve the circadian rhythm of different tissue physiology via controlling “tissue-specific gene expression.”<sup>14</sup>

With the aim of ensuring external synchrony between the body and the surrounding environment, besides internal synchrony and correct chronological alignment between the central and peripheral clocks, the timing of the body circadian system must be calibrated repeatedly.<sup>14</sup> The main entrainment factor (or “Zeitgeber”) for the SCN is light, while peripheral circadian clocks are affected by “neurohumoral modulation.”<sup>15</sup> Therefore, certain behaviors such as the timing of feeding and the timing of abstinence of food may affect the circadian rhythm.<sup>1,12,16</sup> Data that address the impact of IF on the circadian rhythm are limited, and most of those data address the impact of Ramadan diurnal IF on the circadian rhythm. However, current evidence shows that high caloric consumption can lead to alterations in “clock gene expression” in the SCN and other central nuclei.<sup>17,18</sup> In addition, animal studies have shown that temporal food restriction and enforced meal times are powerful synchronizers for body clocks in peripheral tissues, while maintaining a fixed light–dark cycle exposure.<sup>19</sup> Moreover, it has been shown that mealtime schedules can trigger the activation of neural circadian clock genes in several species, which is assumed to be controlled by a “food-entrainable clock.”<sup>22,20</sup>

Almost all physiological systems in the body follow a circadian rhythm, which is reflected by measuring certain hormones and core body temperature.<sup>21</sup> During Ramadan diurnal IF, 2–3 meals are taken after sunset: breakfast (a light meal) is taken at sunset, dinner is taken following night prayer in some countries (~1–3 hour after sunset), and a “predawn meal” (Suhur), which means that food consumption, is moved to the hours of darkness. This food intake shift partly inverts the normal circadian pattern of food intake. In theory, this



**Figure 1** Lifestyle changes that accompany the month of Ramadan and that may affect sleep and circadian rhythms.

shift in mealtime may disrupt the circadian rhythm and biological clock of fasting participants.<sup>12</sup>

## Subjective assessment of chronotype

One study assessed chronotypes using an abridged version of the Horne and Ostberg questionnaire in fasting and nonfasting individuals during Ramadan diurnal IF in a “free-living environment” to define three behavioral groups: morning-type, evening-type, and neither-type.<sup>22,23</sup> Interestingly, the study reported an increase in the evening chronotype among the performers of fasting at both the beginning and the end of Ramadan month compared with baseline. Similarly, an increase in evening chronotype was also reported in nonfasting people during Ramadan, which advocates that lifestyle changes other than fasting may influence the circadian rhythm during Ramadan.<sup>23</sup> This result strengthens the hypothesis that lifestyle changes accompanying Ramadan may affect circadian rhythms.

## Effects of diurnal IF on body temperature

Core body temperature increases during the day and decreases at night. Sleep onset is accompanied by a decrease in core body temperature, whereas an increase in core body temperature initiates the wakefulness process.<sup>21</sup> Studies that assessed changes in core body temperature during Ramadan diurnal IF have reported conflicting results (Table 1). In general, studies that assessed the circadian rhythm during Ramadan diurnal intermittent fasting can be divided into two categories: category 1 encompasses studies that examined the changes in some markers of the circadian clock in an unconstrained “free-living environment” without controlling for confounders and lifestyle changes such as meal composition, caloric intake, total sleep time, energy expenditure, or light exposure,<sup>24–27</sup> and category 2 encompasses studies that controlled for the above confounders.<sup>28–30</sup>

Twenty-four-hour continuous monitoring of rectal temperature in six healthy volunteers during Ramadan revealed delays in both the “bathypase” and the “acrophase” of body temperature (ie, the times at which the calculated minimum and maximum values of body temperature occur, respectively).<sup>25</sup> One more study reported that the circadian rhythm of body temperature was reversed during Ramadan diurnal IF, with significant decreases in oral temperature at the following times: 09:00, 11:00, 13:00, and 16:00, and considerable increases in temperature at 23:00 and 00:00 hours.<sup>26</sup> Conversely, a third paper that assessed the oral temperature of volunteers during the first and third weeks of Ramadan at 08:00, 16:00, and 00:00 using a “high-precision” thermometer while controlling for the composition of meals and sleep duration

reported no changes in body temperature.<sup>30</sup> The above studies did not control for most of the lifestyle changes discussed earlier that may impact the circadian rhythm regardless of fasting. To assess the potential effects of lifestyle changes that accompany Ramadan, a well-designed study recruited young adults with “delayed sleep phase disorder” (evening chronotype) who used to sleep during the daytime and wake up and eat at night.<sup>24</sup> In an unconstrained, free-living environment, the investigators use a portable armband physiological and activity sensor to measure circadian alterations in proximal skin temperature and energy expenditure before Ramadan and during the first 2 weeks of Ramadan.<sup>24</sup> Interestingly, the study revealed a further delay in the “acrophase” of proximal skin temperature and energy expenditure denoting a shift in the circadian pattern. The findings of this study support the hypothesis that factors other than the sudden shift in meal-times may influence sleep patterns and circadian rhythms during Ramadan.<sup>24</sup> Table 1 summarizes the studies that have examined the circadian rhythm of body temperature during diurnal IF for Ramadan.

## Effects of diurnal IF on melatonin and other hormones

Melatonin is a useful marker of circadian rhythm because it is highly precise and accurate.<sup>26,31,32</sup> Therefore, proper measurement of melatonin level is an excellent method to examine the impact of diurnal IF on the circadian rhythm. One study assessed the blood levels of melatonin every 4 hours (not including the 02:00 time point measurement to avoid sleep disruption) before and at the end of the third week of Ramadan.<sup>27</sup> The study revealed a reduced, delayed night peak and a flattened slope in serum melatonin concentrations during Ramadan.<sup>27</sup> Another study, which measured the salivary melatonin levels of eight healthy volunteers (who performed diurnal IF) over a 24 hours period (08:00, 16:00, and 00:00) at three time points – 1 week before Ramadan and at the end of the first and third weeks of Ramadan – while controlling for total sleep time, revealed a statistically significant fall in melatonin concentrations at 00:00 and 16:00 during Ramadan compared to baseline levels.<sup>30</sup> Although the study reported comparable trends in melatonin circadian pattern during Ramadan and at baseline, the slope was flatter in the former case.<sup>30</sup> However, the above studies failed to measure melatonin late at night, neglecting the possibility of a late-night peak in melatonin concentrations.

None of the above studies accounted for potential confounders that may accompany Ramadan and affect the measured melatonin levels. In a subsequent study,

**Table 1** A summary of the studies that objectively assessed changes in the circadian pattern of body temperature during diurnal intermittent fasting for Ramadan

Study	Study population	Study design	Assessment tool	Study setting	Findings	Study details
Roky et al <sup>25</sup>	8 healthy young adults (age: 20–28 years) Location: Morocco	Case crossover study with repeated measures	Rectal thermistor probe for at least 24 hours	Monitoring at home Controlled for meal composition and time, as well as bedtime and rising time	Delay in acrophase and bathyphase	The participants were of the intermediate chronotype as determined by the Morningness- Eveningness Questionnaire Dinner was served 1 hour before bedtime during Ramadan and 3 hours before bedtime at baseline No objective assessment of prior sleep pattern
Roky et al <sup>26</sup>	10 healthy young adults (age: 20–28 years) Location: Morocco	Case crossover study with repeated measures	High-precision medical oral thermometer at 09:00, 11:00, 13:00, 16:00, 20:00, and 23:00	Monitoring at home Controlled for meal composition and time, and bed and rise time	Reversal of circadian pattern of temperature	Subjects were of the intermediate chronotype as determined by the Morningness-Eveningness Questionnaire Sleep duration was 1 hour shorter during Ramadan than at baseline No objective assessment of prior sleep pattern
BaHammam et al <sup>30</sup>	8 healthy young adults (age: 31.8±2 years) Location: Saudi Arabia	Case crossover study with repeated measures	High-precision medical oral thermometer at 08:00, 16:00, and 00:00	Monitoring in the laboratory Controlled for meal composition	No change	No objective assessment of prior sleep pattern Subjects stayed in the laboratory during monitoring
BaHammam et al <sup>24</sup>	6 healthy young adults with delayed sleep phase disorder (age: 18–24 years) Location: Saudi Arabia	Case crossover study with repeated measures	SenseWear Pro Armband™ that measures proximal skin temperature during the last week of Shaban and the first 2 weeks of Ramadan	Free-living environment	Further delay in temperature acrophase	Subjects belonged to the evening chronotype Sleep patterns were monitored for 2 weeks prior to the study by using sleep diaries (no objective assessment) Sleep/wake schedule and sleep duration during the study were assessed objectively via armband Participants lived in an unconstrained environment during the study

we attempted to overcome those limitations by assessing circadian profiles of serum melatonin during diurnal IF outside Ramadan (to assess the impact of diurnal IF act in the absence of the lifestyle changes associated with Ramadan) and during the second week of Ramadan while controlling for light exposure, sleep pattern, total sleep time, and meal composition.<sup>28</sup> In eight healthy volunteers, serum melatonin was measured at five time points (22:00, 02:00, 04:00, 06:00, and 11:00) three times: 4 weeks before Ramadan where volunteers were asked to perform diurnal IF (fasting outside the month of Ramadan) for 1 week, 1 week before Ramadan during which volunteers were not fasting and living a routine life, and during the second week of Ramadan while performing diurnal IF. The trough in melatonin levels occurred 11:00 during the three studied periods, which shows that diurnal IF has no significant effect on the circadian rhythm of melatonin when other confounders are accounted for. In addition, a cosinor analysis of melatonin levels showed no significant changes in the “acrophase.”<sup>28</sup> Therefore, the previously reported changes in melatonin levels during Ramadan could be related to the lifestyle changes that accompany Ramadan. Nonetheless, more studies with larger sample sizes are required to evaluate the effects of diurnal IF on melatonin levels and circadian patterns. Table 2 summarizes the studies that have objectively addressed changes in the circadian pattern of melatonin during diurnal IF for Ramadan.<sup>27,28,30</sup>

Most studies assessing melatonin levels during diurnal IF have consistently reported a reduction in measured melatonin levels, even during short-term experimental fasting.<sup>30</sup> Several theories have been suggested to understand the decrease in melatonin levels during fasting. The nocturnal increase in cortisol levels during Ramadan is a possible mechanism.<sup>33,34</sup> In addition, it has been proposed that a decrease in the glucose supply during fasting may result in a decrease in melatonin synthesis,<sup>35</sup> as it has been shown in rats that mild hypoglycemia decreases melatonin levels, and administration of glucose improves the decrease in melatonin levels.<sup>36</sup> However, previous studies have shown that hypoglycemia does not happen in healthy individuals during diurnal IF.<sup>37</sup> Reduction in tryptophan had been proposed as a possible mechanism of the reduction in melatonin levels during Ramadan diurnal IF, as tryptophan is essential for melatonin synthesis. Nevertheless, this proposed mechanism is improbable because short duration diurnal IF is not likely to cause a reduction in tryptophan.<sup>36</sup>

Both leptin and ghrelin hormones are influenced by fasting, mealtime, and sleep. Hence, the abrupt shift in mealtime

during diurnal IF for Ramadan may entrain the circadian rhythms of both hormones.

Two previous studies have examined the impact of diurnal IF on the circadian patterns of leptin using a cosinor analysis. The first study was conducted on free-living participants and revealed a major shift (~5 hours) in the peak and nadir serum leptin concentrations in the third week of Ramadan.<sup>38</sup> However, the amplitude and 24 hours mean concentrations of leptin levels revealed no significant changes compared to the levels before Ramadan. Another study reported a significant decrease in plasma leptin concentration at 22:00; however, a cosinor analysis demonstrated no significant changes in the circadian rhythm of leptin.<sup>29</sup> The sudden change in mealtime during Ramadan may explain the nocturnal decrease in plasma leptin levels. The difference between the findings of the two studies could be explained by the fact that the second study controlled for lifestyle changes and environmental conditions, while the first study did not, which might have resulted in a delay in the circadian rhythm of leptin in the first study.

So far, only one study has evaluated circadian rhythm of ghrelin during Ramadan diurnal IF while controlling for eating habits and environmental conditions.<sup>29</sup> A cosinor analysis revealed no significant changes in the “acrophase” of the circadian rhythm of ghrelin.<sup>29</sup>

In summary, the above-reviewed studies suggest that during Ramadan diurnal IF, there is a delay in the circadian clock. However, when eating habits and environmental conditions are accounted for, diurnal IF does not significantly affect the circadian rhythm of body temperature and melatonin.

## Effects of diurnal IF on sleep

### Subjective assessment of sleep

When assessing the influences of diurnal IF during Ramadan on sleep, it is essential to consider the accompanying lifestyle changes during the holy month. For example, in some Muslim countries, the beginning of work is delayed and work duration is shortened during Ramadan. Some of the earlier studies that assessed sleep patterns during Ramadan used sleep diaries and self-reported data. Studies that assessed sleep/wake pattern in three Muslim countries revealed a significant sudden delay in bedtime and wake time during Ramadan in performers of diurnal IF.<sup>21,23,30,39,40</sup>

Interestingly, one study demonstrated bedtime delay during Ramadan even in nonfasting subjects.<sup>23</sup> This documented delay in bedtime and rise time reflects the delay in the start of work during Ramadan.

**Table 2** A summary of the studies that objectively assessed changes in melatonin during diurnal intermittent fasting for Ramadan

Study	Study population	Study design	Assessment tool	Study setting	Findings	Study details
Bogdan et al <sup>27</sup>	10 healthy male volunteers (age: 32–40 years) Location: France	Case crossover study with repeated measures	Blood samples were obtained every 4 hours, omitting the 02:00 time point, before and on the 23rd day of Ramadan	Free-living environment Controlled for meal timing and composition Did not control for light exposure, sleep schedule, or social habits that accompany Ramadan	A decreased and delayed night peak and a flattened slope of serum melatonin concentration in Ramadan	Volunteers slept 1 hour less during Ramadan than before Ramadan Melatonin concentrations were not measured late at night, which fails to address the possibility of a late peak in melatonin concentration
BaHammam et al <sup>30</sup>	8 healthy young adults (age: 31.8±2 years) Location: Saudi Arabia	Case crossover study with repeated measures	Saliva samples were collected at three time points over a 24-hour period (08:00, 16:00, and 00:00) before and on the 7th and 21st days of Ramadan	In-laboratory monitoring Controlled for meal timing and composition Controlled for sleep duration Did not control for light exposure or social habits that accompany Ramadan	A significant decrease in melatonin concentrations at 00:00 and 16:00 during Ramadan Melatonin profiles continued to show the same trend during Ramadan, but with a flatter slope	Melatonin concentrations were not measured late at night, which fails to address the possibility of a late peak in melatonin concentration
Almeneessier et al <sup>28</sup>	8 healthy young adults (age: 26.6±4.9 years) Location: Saudi Arabia	Case crossover study with repeated measures	Blood samples were collected at 22:00, 02:00, 04:00, 06:00, and 11:00 before Ramadan and while performing fasting outside Ramadan month and on the second week of Ramadan	In-laboratory monitoring Controlled for light exposure, sleep schedule, sleep duration, energy expenditure, and meal composition	Intermittent fasting during Ramadan has no significant effect on the circadian pattern of melatonin	Assessed melatonin level when volunteers were fasting outside Ramadan month to control for lifestyle changes that accompany Ramadan



With regard to sleep duration, conflicting results were reported during Ramadan. Whereas some studies reported a reduction of sleep duration during Ramadan,<sup>39–41</sup> others reported no significant alterations in nighttime sleep duration during Ramadan.<sup>21,23</sup> Nevertheless, in a study that assessed sleep duration objectively using actigraphy in eight fasting (diurnal IF) and eight nonfasting volunteers in a free-living environment during Ramadan, bedtime and wake time were delayed in the fasting volunteers, and a significant reduction in total sleep time was observed in the fasting group.<sup>42</sup>

The discrepancies between the above studies may reflect cultural and lifestyle variations among countries or the use of different subjective assessment tools.<sup>12,13</sup>

## Objective assessment of sleep

### Sleep architecture

A few studies have objectively measured sleep parameters during the month of Ramadan among volunteers practicing diurnal IF using in-laboratory polysomnography and home-based ambulatory unattended polysomnography.<sup>25,30,43,44</sup> Table 3 summarizes the studies that have evaluated sleep architecture using polysomnography during Ramadan diurnal IF. In studies that controlled for sleep/wake pattern, diurnal IF had no effect on nonrapid eye movement sleep stages, stage shifts, arousal index, or measured cardiorespiratory parameters.<sup>25,30,44</sup> One study objectively assessed sleep among young athletes and revealed no changes in total sleep time during Ramadan diurnal IF; however, the study reported a significant increase in the number of awakenings.<sup>43</sup> Nevertheless, a major limitation of the last study is the fact that the study athletes suddenly changed their primary sleep time from night to daytime, and the study did not account for naps. Therefore, the abrupt changes in bedtime and rise time may have accounted for the observed changes in sleep architecture.<sup>17,45</sup>

A recent study controlled for the lifestyle changes associated with Ramadan, such as sleep/wake pattern, caloric consumption, meal composition, and light exposure to eliminate variables other than meal timing that might affect sleep architecture; it objectively assessed sleep architecture at baseline, during diurnal IF outside Ramadan, during Ramadan, and during a nonfasting period after Ramadan.<sup>44</sup> Rapid eye movement (REM) sleep proportion decreased while fasting during and outside Ramadan compared with the baseline; however, the proportion of REM sleep reverted to normal during recovery (after Ramadan).<sup>44</sup> Three other studies reported a reduction in REM sleep during Ramadan.<sup>25,30,43</sup> In agreement with the above, studies that assessed the effect

of experimental fasting on animals reported similar changes in sleep architecture.<sup>46</sup> REM sleep disappeared in piglets following 18 hours of fasting, which reverted to normal after feeding.<sup>46</sup>

Several hypotheses have been proposed to account for the decrease in REM sleep during fasting; however, the precise mechanisms are unknown.<sup>12</sup> In addition, the clinical consequences of reduced REM sleep during fasting are unknown. Proposed theories to understand the decrease in REM sleep during fasting include the documented nocturnal rise in cortisol and insulin,<sup>47,48</sup> and the possible increase in nocturnal body temperature as a result of eating exclusively at night during diurnal IF.<sup>25,26</sup> Increased nocturnal temperature could result in a reduction in REM sleep proportion.<sup>49,50</sup> Another proposed mechanism is the interruption of sleep in the early morning (predawn) for the predawn meal (Suhur), which is the period in which the major fraction of REM sleep usually occurs.<sup>30</sup>

### Sleep latency

With regard to sleep latency during Ramadan IF, studies have reported conflicting results. Roky et al showed a significant increase in sleep latency and a significant reduction in nocturnal sleep time.<sup>25</sup> However, in that study, dinner was served 30 minutes before going to bed; the short time between dinner and bedtime could have affected sleep latency. Later studies that provided dinner at an earlier time (3–3.5 hours before bedtime) and controlled daytime naps revealed no change in sleep latency.<sup>13,30,44</sup>

The above-reviewed studies suggest that when dinner is provided early and daytime naps are controlled for, diurnal IF does not have major influence in sleep architecture. The only change in sleep architecture consistently reported is the reduction in REM sleep.<sup>44</sup> However, in a free-living environment that does not control for confounders that usually accompany Ramadan such as mealtimes or the daytime naps, sleep latency and sleep architecture may be disturbed.<sup>43</sup>

### Daytime sleepiness

A discussion of the effect of diurnal IF on sleep would not be complete without reviewing the impact of daytime fasting on daytime sleepiness. Several studies have assessed the interaction between daytime sleepiness and diurnal IF during Ramadan both subjectively<sup>21,23,30,39,40,42,44,51</sup> and objectively.<sup>30,44,51</sup> However, these studies reported conflicting results, which may reflect the differences in the used assessment tools, the lack of objective assessment or the failure to control for potential confounders that could impact daytime

**Table 3 A summary of the studies that assessed sleep architecture using polysomnography during diurnal intermittent fasting for Ramadan**

Study	Study population	Study design	Test used for assessment	Study details	Findings
Roky et al <sup>25</sup>	8 young healthy adults (age: 20–28 years) Location: Morocco	Case crossover study with repeated measures	Ambulatory 8-channel unattended PSG	Unattended PSG Dinner was served at 22:30, and PSG recording started at 23:30 Did not objectively account for the prior sleep/wake pattern before assessing sleep Did not monitor for daytime naps prior to overnight sleep study Meals during and outside Ramadan were according to a fixed schedule and composition	Significant increase in sleep latency and reduction in total sleep time Increase in stage N2 and reduction in slow wave sleep Reduced REM sleep
BaHamam et al <sup>30</sup>	8 young healthy adults (age: 31.8±2 years) Location: Saudi Arabia	Case crossover study with repeated measures	Full attended level I in-laboratory PSG	Controlled for sleep schedule, naps, and caloric intake Did not objectively account for the prior sleep/wake pattern before assessing sleep in the laboratory	Significant drop in sleep latency at the end of Ramadan, with no change in total sleep time Reduced REM sleep at the end of Ramadan No significant changes in NREM sleep
BaHamam et al <sup>44</sup>	8 young healthy adults (age: 26.6±4.9 years) Location: Saudi Arabia	Case crossover study with repeated measures	Full attended level I in-laboratory PSG	Controlled for sleep schedule, naps, light exposure, caloric intake, and energy expenditure Assessed the effect of fasting during and outside Ramadan Controlled for sleep/wake and naps for 2 weeks before assessing sleep in the laboratory via actigraphy	Reduced REM sleep during intermittent fasting (during and outside Ramadan) No significant changes in NREM sleep No differences in sleep latency, arousal index, or sleep efficiency
Chamari et al <sup>43</sup>	11 young healthy trained cyclists (age: 21.6±4.8 years) Location: Qatar	Case crossover study with repeated measures	Portable PSG	Unattended PSG Did not control for sleep/wake pattern or sleep duration During Ramadan, volunteers slept during daytime and at night before and after Ramadan During Ramadan, the volunteers slept in the morning after eating a main meal Naps were not controlled for Did not account for the prior sleep/wake pattern before assessing sleep Participants were cyclists at a training camp The portable device used in the study had relative weakness at the level of the number of awakenings Data regarding sleep duration were highly heterogeneous	No change in sleep duration Significant increases in the number of awakenings and light sleep in Ramadan Progressive decrease in duration of sleep and REM sleep stages that became significant 2 weeks after Ramadan

**Abbreviations:** NREM, nonrapid eye movement sleep; PSG, polysomnography; REM, rapid eye movement sleep.



alertness. Table 4 summarizes the studies that have assessed daytime sleepiness during diurnal IF in Ramadan.

## Subjective assessment

Studies that used the Epworth Sleepiness Scale (ESS) score to assess daytime sleepiness reported conflicting results. Some studies reported a significant increase in daytime sleepiness,<sup>21,40</sup> whereas other studies demonstrated no significant changes.<sup>23,30,39,52</sup>

## Objective assessment

In three studies, daytime sleepiness was measured objectively using the multiple sleep latency test (MSLT) under controlled conditions.<sup>30,44,51</sup> The first study demonstrated an increase in daytime sleepiness at the 10:00 and 12:00 naps in the last week of Ramadan.<sup>51</sup> However, the study used an unattended polysomnography recording device, which necessitated the end of the test 20 minutes after the beginning of recording regardless of sleep onset.

To avert the above shortcomings, a subsequent study used a standard attended MSLT.<sup>30</sup> This study measured sleep latency, sleep onset latency, or wake efficiency on the first and third weeks of Ramadan and reported no changes compared with baseline measurements.<sup>30</sup> Furthermore, spectral analysis of electroencephalography waves during each nap revealed no difference between Ramadan and baseline.<sup>30</sup> However, the previous studies did not control for the possibility of sleep deprivation on the nights prior to assessing sleep and daytime sleepiness. A recent study aimed to objectively assess the impact of diurnal IF during Ramadan on daytime sleepiness in eight healthy subjects while providing a fixed sleep–wake schedule, fixed caloric intake, controlled light exposure, and objective monitoring of sleep duration for several nights before the assessment of daytime sleepiness.<sup>44</sup> The assessment was performed on four occasions using attended MSLT and the ESS. During the first visit (adaptation night), wrist actigraphy was provided before the start of the study to monitor sleep pattern while at home to ensure that they had a fixed sleep–wake pattern. In subsequent visits, daytime sleepiness was measured using MSLT and ESS in the following order: 21 days before Ramadan after 1 week of diurnal IF “baseline fasting,” 7 days before Ramadan “nonfasting baseline,” on the 24th day of Ramadan, and 14 days after Ramadan “recovery.” The study revealed no changes in the ESS across the four study times. In addition, the MSLT measurements revealed no differences in sleep latency among the “nonfasting baseline,” “baseline fasting,” “Ramadan,” and “recovery” time points.

Another study assessed the impact of diurnal IF during Ramadan on drowsiness under controlled conditions with a fixed light/dark exposure, caloric intake, sleep/wake schedule and sleep duration via measuring total blink duration (measured using infrared reflectance (IR) oculography) and the mean reaction time (MRT).<sup>61</sup> The results of this study showed that diurnal intermittent fasting had no impact on drowsiness or vigilance.<sup>61</sup>

CR has been reported to enhance alertness and to upregulate orexin gene expression in animal models.<sup>53,54</sup> It has been proposed that mammals, when exposed to a negative energy balance, respond with enhanced alertness to improve their capability to find food.<sup>55,56</sup> It has also been shown in animals that orexin (an excitatory neurotransmitter) neurons are important for this adaptive response during fasting.<sup>56,57</sup> Moreover, orexin neurons interact closely with the “hypothalamic–pituitary–adrenal” (HPA) axis,<sup>58</sup> and stimulation of the sympathetic and HPA tone secondary to fasting may further induce arousal through the orexin system.<sup>59</sup> Therefore, a recent study assessed the effects of diurnal IF during the month of Ramadan on plasma orexin while controlling for lifestyle changes that may accompany Ramadan, like sleep duration, bedtime and wake time, energy expenditure, light exposure, and food, in eight healthy young men.<sup>60</sup> The results showed that plasma orexin levels increased during the fasting hours.

## Summary

The impact of diurnal IF on circadian rhythm, sleep, and daytime sleepiness is an exciting topic that deserves future research. Earlier studies have suggested that diurnal IF during Ramadan causes a delay in the circadian rhythm of core body temperature and hormonal secretion. However, more recent studies controlling for lifestyle changes that may influence circadian rhythm showed no effects of diurnal IF on the circadian rhythm.

In free-living, unconstrained environments that do not control for lifestyle changes, studies have revealed abrupt significant delays in bedtime and rise time. Nevertheless, studies that controlled for environmental conditions and sleep–wake pattern revealed no significant changes in sleep architecture, indicating that the earlier reported changes could be secondary to concomitant lifestyle changes rather than the fasting act per se. The only consistent change in all studies that assessed sleep architecture during diurnal IF is a decrease in the amount of REM sleep.

Although earlier subjective studies reported an increase in daytime sleepiness during diurnal IF, subsequent studies that controlled for confounding factors reported no changes.

**Table 4 A summary of the studies that assessed daytime sleepiness and daytime alertness during diurnal intermittent fasting for Ramadan**

Study	Study population	Study design	Assessment tool	Findings	Study details
<b>Subjective assessment</b>					
BaHammam et al <sup>21</sup>	56 healthy medical students (age: 22.6±1.3 years) Location: Saudi Arabia	Case crossover study with repeated measures	ESS	Increase in daytime sleepiness	Subjective assessment Assessment was conducted in an unconstrained environment
Bencheikroun et al <sup>40</sup>	264 young subjects (age: 20–30 years) Location: Morocco	Case crossover study with repeated measures	ESS	Increase in daytime sleepiness	Subjective assessment Assessment was conducted in an unconstrained, free-living environment
BaHammam et al <sup>23</sup>	101 healthy fasting and nonfasting subjects (age: 31.3±2.1 years) Location: Saudi Arabia	Case-control study with repeated measures	ESS	No change in daytime sleepiness	Subjective assessment Assessment was conducted in an unconstrained, free-living environment
Margolis et al <sup>39</sup>	109 healthy medical students (age: 19–23 years) Location: United Arab Emirates	Case crossover study with repeated measures	ESS	No change in daytime sleepiness	Subjective assessment Assessment was conducted in an unconstrained, free-living environment
Nugraha et al <sup>52</sup>	Among young healthy adults Fasting group (n=25) (age: 26.12±0.98 years); nonfasting group (n=25) (age: 26.2±0.98 years) Location: Germany	Case-control with repeated measures	ESS	No difference in daytime sleepiness between the two groups However, the ESS score among the fasting group was significantly lower in the last week of Ramadan than in the first week	Subjective assessment Assessment was conducted in an unconstrained, free-living environment
<b>Objective assessment</b>					
Roky et al <sup>26</sup>	10 healthy young subjects (age: 20–28 years) Location: Morocco	Case crossover study with repeated measures	Visual analog scale MRT CFF All measurements taken at 6 different times of the day: 09:00, 11:00, 13:00, 16:00, 20:00, and 23:00	Decreased daytime alertness MRT increased at the beginning of Ramadan CFF did not change	Controlled for sleep/wake pattern Volunteers slept 1 hour less during Ramadan than at baseline Controlled for meal composition and physical activity

(Continued)

Table 4 (Continued)

Study	Study population	Study design	Assessment tool	Findings	Study details
BaHammam et al <sup>42</sup>	16 fasting and nonfasting subjects 8 fasting (age: 36.25±4.46 years) 8 nonfasting (age: 34.75±3.33 years) Location: Saudi Arabia	Case-control with repeated measures	Johns Drowsiness Scale to assess sleepiness Visual reaction time test ESS	No decrease in alertness No change in daytime sleepiness	Assessed sleep duration objectively using the SenseWear Pro Armband™. There was a significant reduction in sleep duration during Ramadan in the fasting group Assessment was conducted in an unconstrained environment
BaHammam et al <sup>41</sup>	8 healthy young male subjects (age: 25.3±2.9 years) Location: Saudi Arabia	Case crossover study with repeated measures	Johns Drowsiness Scale IR for total blink duration and a visual reaction time test	No decrease in alertness	Controlled for sleep/wake schedule, sleep duration, caloric intake, energy expenditure, and light exposure Actigraphy to assure adequate sleep duration in days prior to the study
Chamari et al <sup>43</sup>	11 young healthy trained cyclists (age: 21.6±4.8 years) Location: Qatar	Case crossover study with repeated measures	CANTAB, RTI, and RVP tests	No decrease in alertness RTI was not affected by Ramadan intermittent fasting or time of day Overall, RVP accuracy increased during and after Ramadan compared with baseline; in the last week of Ramadan, accuracy was highest at the end of the day	Sleep duration was not assessed during the 24 hours Tests were conducted at different times during and outside Ramadan
Roky et al <sup>41</sup>	8 healthy young subjects (age: 20–28 years) Location: Morocco	Case crossover study with repeated measures	Portable MSLT Visual analog scale	Increase in daytime sleepiness Subjective alertness decreased at 12:00 on day 11 of Ramadan but did not change on day 25 On the MSLT, sleep latency was decreased on day 11 and day 25 of Ramadan	Portable device used; programmed to end test after 20 minutes of recording Sleep duration was significantly lower during Ramadan than at baseline Meals during baseline and Ramadan followed a fixed schedule and composition Did not rule out possible sleep deprivation in nights prior to study

(Continued)

Table 4 (Continued)

Study	Study population	Study design	Assessment tool	Findings	Study details
BaHamman et al <sup>30</sup>	8 healthy young subjects (age: 31.8±2 years) Location: Saudi Arabia	Case crossover study with repeated measures	ESS and standard MSLT	No change in daytime sleepiness	Did not rule out possible sleep deprivation in nights prior to study MSLT was preceded by an overnight in-laboratory PSG
BaHamman et al <sup>44</sup>	8 healthy young subjects (age: 26.6±4.9 years) Location: Saudi Arabia	Case crossover study with repeated measures	ESS and standard MSLT Actigraphy to assess sleep duration in days prior to the study	No change in daytime sleepiness during Islamic intermittent fasting	Sleepiness was assessed while the volunteers were performing intermittent fasting during and outside Ramadan Controlled for sleep duration in nights prior to study and when at home by objective measurements (actigraphy)
Tian et al <sup>62</sup>	18 male athletes (age: 17–29 years) Location: Singapore	Case crossover study with repeated measures	Computerized neuropsychological testing	Performance in functions requiring sustained rapid responses was best in the morning and declined in the late afternoon Performance in nonspeed-dependent accuracy measures was more resilient	Standardized meals were provided Participants' lifestyle and training in-between test sessions were not controlled for Sleep duration was significantly shorter during Ramadan than at baseline Previous night's sleep and daytime naps, as well as the time of awakening, were not controlled for in this study Participants answered questionnaires regarding diet and sleep duration in the previous 24 hours

**Abbreviations:** CANTAB, Cambridge Neuropsychological Test Automated Battery; CFF, critical flicker fusion; ESS, Epworth Sleepiness Scale; IR, infrared reflectance; MRT, movement reaction time; MSLT, multiple sleep latency test; PSG, polysomnography; RTI, Reaction Time Index; RVP, Rapid Visual Information Processing.

Larger studies that control for potential confounding factors, such as environmental and cultural conditions (the delay in the start of work and school, light exposure, exercise, and meal composition), are warranted to assess the impact of diurnal IF on circadian rhythms, sleep, and daytime sleepiness.

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## Disclosure

The authors report no conflicts of interest in this work.

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