

The µMCTQ: An Ultra-Short Version of the Munich ChronoType Questionnaire

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Abstract Individuals vary in how their circadian system synchronizes with the cyclic environment (zeitgeber). Assessing these differences in "phase of entrainment"—often referred to as chronotype—is an important procedure in laboratory experiments and epidemiological studies but is also increasingly applied in circadian medicine, both in diagnosis and therapy. While biochemical measurements (e.g., dim-light melatonin onset [DLMO]) of internal time are still the gold standard, they are laborious, expensive, and mostly rely on special conditions (e.g., dim light). Chronotype estimation in the form of questionnaires is useful in approximating the timing of an individual's circadian clock. They are simple, inexpensive, and location independent (e.g., administrable on- and offline) and can therefore be easily administered to many individuals. The Munich ChronoType Questionnaire (MCTQ) is an established instrument to assess chronotype by asking subjects about their sleep-wake-behavior. Here we present a shortened version of the MCTQ, the µMCTQ, for use in situations in which instrument length is critical, such as in large cohort studies. The µMCTQ contains only the core chronotype module of the standard MCTQ (stdMCTQ), which was shortened and adapted from 17 to 6 essential questions, allowing for a quick assessment of chronotype and other related parameters such as social jetlag and sleep duration. µMCTQ results correspond well to the ones collected by the stdMCTQ and are externally validated by actimetry and DLMO, assessed at home (no measure of compliance). Sleep onset, midpoint of sleep, and the µMCTQ-derived marker of chronotype showed slight deviations toward earlier times in the μ MCTQ when compared with the stdMCTQ (<35 min). The µMCTQ assessment of chronotype showed good test-retest reliability and correlated significantly with phase markers from actimetry and melatonin (DLMO), especially with measurements taken on work-free days. Because of its brevity, the µMCTQ represents an ideal tool to estimate individual internal time in time-critical contexts, from large cohort studies to individualized medicine.

Keywords validation, chronotype, phase of entrainment, DLMO, actigraphy, circadian

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Our daily lives are controlled by at least 2 "clocks," which historically used to be in phase: (1) the sun clock that defines day and night, dawn and dusk, or photoperiod that—depending on latitude—changes over the year and (2) social clocks that represent "local time" and allow us to interact with others (school, work, business hours). Over the course of history, we introduced time zones and Daylight Saving Time (DST), thereby separating local time and sun time. Circadian clocks synchronize to the 24-h day predominately through light and darkness, but the strength of this zeitgeber has greatly decreased during industrialization, as humans live predominantly in buildings throughout the day and artificially illuminate the nights. Everyone's circadian system establishes its own specific phase relationship with the zeitgeber cycle (phase of entrainment [PoE]), evidenced by the difference in timing of biological rhythms in reference to the light-dark cycle between individuals, for example, melatonin, temperature, peak of activity/activity onset, peak of cortisol, or behavioral timing like that of sleep and wake (Roenneberg et al., 2003a). These differences in PoE are commonly called chronotype, ranging from extreme early chronotypes (larks) to extreme late chronotypes (owls). As a consequence of the changed light-dark cycles, the PoE of extreme early types has become even earlier and that of all the other chronotypes has become delayed, greatly widening the difference between early and late chronotypes, especially within urban populations (Roenneberg et al., 2007b; Stothard et al., 2017; Swaminathan et al., 2017; Wright et al., 2013).

Since practically all functions in our body are directly or indirectly organized by the circadian clock, temporal inconsistencies between biological and social timing become problematic, increasing the need to estimate individual internal time (PoE, chronotype, circadian phase) in research or medicine (from diagnosis to treatment). The gold standard for assessing circadian phase is measuring dim-light melatonin onset (DLMO) in samples collected in highly controlled settings, in blood, urine, or saliva (Benloucif et al., 2008; Klerman et al., 2002; Lewy and Sack, 1989). However, these measurements are expensive and cumbersome, involving multiple, well-timed samplings. Although circadian researchers are currently developing methods to assess circadian state with 1 to 2 measurements, these so far still require sampling blood, involving many known complications that limit their application in large-scale studies (Braun et al., 2018; Laing et al., 2017; Wittenbrink et al., 2018). A cost-effective, scalable, and noninvasive solution to this challenge is the use of questionnaires.

The first questionnaire developed to detect individual differences in circadian rhythms, the Morningness-Eveningness Questionnaire, uses temporal preferences to compute a score and classify individuals into chronotypes accordingly (Horne and Ostberg, 1976). The Munich ChronoType Questionnaire (MCTQ), on the other hand, was introduced in 2003 (Roenneberg et al., 2003b) and considers chronotype as the phase relationship between the circadian system of an individual and the zeitgeber cycle: their circadian state. Since it is virtually impossible to assess the phase of all rhythmic processes in humans, the MCTQ uses sleep timing as a phase marker to estimate chronotype. The standard core questions inquire about sleep times separately for work and work-free days. Considering that sleep on work-free days is not as restricted by social constraints, chronotype can be estimated using the midpoint between sleep onset and sleep end on free days (MSF), which is corrected for potential oversleep on free days (to compensate for sleep debt accumulated over the workweek, MSF_{sc}), therefore accounting for the homeostatic process influencing sleep. Other modules assess the use of stimulants or biographic information, for example.

The stdMCTQ-MSF_{sc} has already been shown to correlate well with data from sleep logs, wrist actimetry, and DLMO (Kantermann et al., 2015; Kitamura et al., 2014; Roenneberg et al., 2007a; Wright et al., 2013). It has been used for 15 years to assess thousands of peoples' sleep behavior (the MCTQ database associated with its online version alone contains \approx 300,000 entries).

Thus, the µMCTQ was developed as an ultra-short version of the stdMCTQ. This questionnaire makes use of the same principles as the stdMCTQ but contains only the essential questions of the MCTQ's core chronotype module. As with the stdMCTQ, MSF_{sc} serves as a marker for chronotype and an approximation for PoE. The µMCTQ therefore allows for a swift assessment of the timing of an individuals' clock, which can be especially useful in larger cohort studies, long durations of investigation, or for efforts of personalized medical practice.

Here we present the validation of the circadian phase assessment by the µMCTQ against assessments by the stdMCTQ (study 1) as well as DLMO and actimetry (study 2). In the supplementary material, we provide additional data supporting the validity of the stdMCTQ for assessing PoE (study 3). Our results show that both the µMCTQ and the stdMCTQ have good validity against commonly used physiological and behavioral markers of PoE.

METHODS

Development of the µMCTQ

The µMCTQ was developed starting from the original MCTQ (Roenneberg et al., 2003b), in the following referred to as standard MCTQ (stdMCTQ).

Munich ChronoType Questionnaire (MCTQ)

I have a regular work schedule (this includes being, for example, a housewife or househusband):
Yes ☐ I work on 1☐ 2☐ 3☐ 4☐ 5☐ 6☐ 7☐ days per week.
No 🗆
Is your answer "Yes, on 7 days" or "No", please consider if your sleep times may <u>nonetheless</u> differ between regular 'workdays' and 'weekend days' and fill out the MCTQ in this respect.
Please use 24-hour time scale (e.g. 23:00 instead of 11:00 pm)!
<u>Workdays</u>
Image 1: I go to bed at o'clock.
Image 2: Note that some people stay awake for some time when in bed!
Image 3: I actually get ready to fall asleep at o'clock.
Image 4: I need minutes to fall asleep.
Image 5: I wake up at o'clock.
Image 6: After minutes I get up.
I use an alarm clock on workdays: Yes ☐ No ☐
If "Yes": I regularly wake up BEFORE the alarm rings: Yes ☐ No ☐
Free Days
Image 1: I go to bed at o'clock.
Image 2: Note that some people stay awake for some time when in bed!
Image 3: I actually get ready to fall asleep at o'clock.
Image 4: I need minutes to fall asleep.
Image 5: I wake up at o'clock.
Image 6: After minutes I get up.
My wake-up time (Image 5) is due to the use of an alarm clock: Yes ☐ No ☐
There are particular reasons why I cannot freely choose my sleep times on free days:
Yes ☐ If "Yes": Child(ren)/pet(s) ☐ Hobbies ☐ Others ☐, for example:
No 🗆

Figure 1. The core module of the standard Munich ChronoType Questionnaire (stdMCTQ) asks questions about sleep timing in relation to the weekly structure. It does so by leading the sleeper in and out of bed in 6 steps, both for workdays and work-free days. In total, participants need to answer 14 to 17 questions (depending on specific answers). On a separate page, participants are given instructions on how to fill in the stdMCTQ, for example, "take an example month" or "according to the past 4 weeks."

The stdMCTQ asks simple questions about sleepwake behavior, separately for work and work-free days. Its core module, which focuses on the estimation of chronotype, contains a total of 17 questions (other optional modules, e.g., regarding the use of stimulants and sociodemographics have varying lengths). The original idea of the stdMCTQ core questions was to accompany people into and out of bed (see Fig. 1 for the complete questionnaire). Since sleep onset (falling asleep) and bed time (going to

µMCTQ

The following section will ask you questions in regards to your sleep and wake behavior on workand work-free days. Please estimate an average of your 'normal' sleep behavior over the past 6 weeks.

I have been a shift- or night-worker in the past three months yes no
Normally, I work days/week.
Please answer all the following questions even if you do not work or work 7 days/week. Please don't forget to circle AM or PM.
On WORKDAYS I normally fall asleep at: AM/PM (this is NOT when you get into bed, but rather when you fall asleep)
I normally wake up at: AM/PM (this is NOT when you get out of bed, but rather when you wake up)
On WORK-FREE DAYS when I DON'T use an alarm clock I normally fall asleep at: AM/PM (this is NOT when you get into bed, but rather when you fall asleep)
I normally wake up at: AM/PM (this is NOT when you get out of bed, but rather when you wake up)

Figure 2. The entire ultra-short version of the Munich ChronoType Questionnaire (µMCTQ) consists of a short explanatory introduction, 2 work-related questions, another short instruction, and 2 questions about sleep timing each for work and work-free days. Thus, participants have to answer 6 questions in total.

bed) are often confused, we asked people questions about every step: (1) going to bed, (2) being busy in bed before deciding (3) to prepare for sleep (e.g., by switching off lights), (4) falling asleep (the last 2 indicate sleep latency), (5) waking up, and (6) finally getting up (the last 2 indicate sleep inertia). The stdMCTQ and more information on it can be found at http://thewep.org/documentations/mctq.

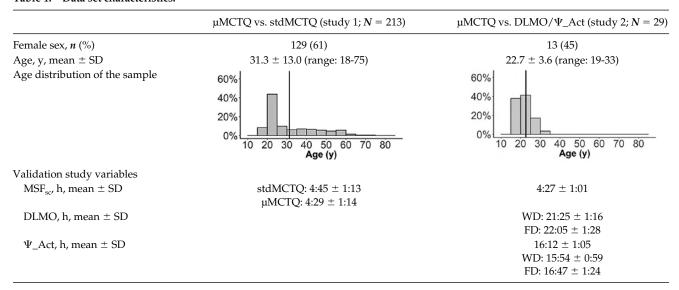
Naturally, in aiming for brevity, the µMCTQ (Fig.2) contains only questions from the core chronotype module of the stdMCTQ and no questions from the optional additional stdMCTQ modules. The core module was then reduced to questions pertaining to the core variables, which were slightly modified but are content-wise identical. Since only sleep onset and sleep end as well as the weekly structure are used to calculate the most important stdMCTQ variables (MSF, MSF_{sc}, sleep duration, and social jetlag), we reduced the 17 questions to 6 questions, probing only these events. We also tried to make participants aware that we do not mean bedtime or rise time but actual time of falling asleep or waking up.

Both the stdMCTQ and the µMCTQ estimate chronotype in 2 steps. First, the midpoint of sleep on work-free days (MSF) is calculated based on sleep onset (SO_f) and sleep end (SE_f) : $MSF = SO_f +$

 $(SE_f - SO_f)/2$, whereby $(SE_f - SO_f)$ provides the sleep duration on work-free days (SD_f). The same variables can be assessed for workdays: midpoint of sleep on workdays (MSW = $SO_w + (SE_w - \overline{SO}_w)/2$) and sleep duration on workdays ($SD_{w} = SE_{w} - SO_{w}$).

The second step to compute the chronotype indicator is to further correct MSF for the potential sleep debt accumulated during the workweek. This linear correction is based on the weighted average of sleep duration across the week (SD_{week}) and on the sleep duration on work-free days (SD_f). The difference of the two is taken as an estimate for how much longer subjects slept on a work-free day (versus if they had no prior sleep debt, when $SD_f \leq SD_w$): if $SD_f \leq SD_w$, $MSF_{sc} = MSF$; if $SD_f > SD_w$, $MSF_{sc} = MSF - (SD_f - SD_f - SD$ SD_{week})/2. Free days are used for chronotyping, as those are the days assumed to be relatively free of constraints on sleep-wake behavior. In the std-MCTQ, participants are asked to specify the need of an alarm clock ("yes" or "no" answer): MSF_{sc} calculations are considered only when the participant does not use/need an alarm clock on work-free days. In the µMCTQ, however, participants are asked to report their wake-up times only on workfree days on which they do not use an alarm clock. Therefore MSF_{sc}, derived from the µMCTQ can be

Table 1. Data set characteristics.



Data sets used in the study for validating the μ MCTQ against the stdMCTQ (column 1) and for validating the μ MCTQ against DLMO and Ψ _Act (column 2). stdMCTQ = standard Munich ChronoType Questionnaire; μ MCTQ = ultra-short version of the stdMCTQ; MSF_{sc} = midpoint between sleep onset and end on free days corrected for potential oversleep on free days to compensate for the sleep debt accumulated over the workweek; DLMO = dim-light melatonin onset; Ψ _Act: center of gravity of activity; WD = workday; FD = free day.

computed for all subjects. A detailed overview of the calculations of the mentioned variables can also be found in the supplementary material (Suppl. Table S2).

Validation of the µMCTQ

The validation of the μ MCTQ against the std-MCTQ was carried out at the University of Padova, Italy, and is further referred to as study 1. The study for the validation of the μ MCTQ against the phase markers from melatonin (DLMO) and actimetry data was carried out at the Ludwig Maximilian University in Munich, Germany, and is referred to as study 2. Data supporting the validity of the stdMCTQ against actimetry are included in the supplementary material and referred to as study 3.

The Italian version of the μ MCTQ, used in study 1, was obtained by the Sackett procedure (i.e., forward translation, expert evaluation, independent backtranslation, pretesting, and definition of the final version). The German μ MCTQ version used in study 2 was a simple translation from the English μ MCTQ version into German. Both translated versions (German and Italian) ask individuals to use the 24-h military time format (e.g., 23:00 h instead of 11:00 p.m.). Table 1 offers an overview of the different sample characteristics in the different studies. Detailed descriptions of the study designs can be found below.

Study 1: Validation of the µMCTQ against the std-MCTQ

Participants. For the validation of the μ MCTQ against the stdMCTQ, we recruited 361 healthy volunteers as part of a series of popular science initiatives (open Padova University event "Veneto Research Night" 2015) during which Padova University scientists provided the general public with information on their ongoing research. Forty-eight subjects were excluded because of significant medical history, shift work, and/or incomplete questionnaire responses. Because of alarm clock use on free days, 87 participants were excluded from all analyses. Thus, the final population included 213 individuals (129 women; age [mean \pm standard deviation]: 31.3 \pm 13.0 years). See Table 1 for further details.

Study Design. Subjects completed a personal datasheet to include basic demographic and medical information, height, and weight before they filled out the Italian translations of the µMCTQ followed by the stdMCTQ (always in the same sequence). The researchers G.F., M.S., and S.M. provided assistance and instructions on completion of the questionnaires. Participants provided written, informed consent.

Measurements. Participants filled in the μMCTQ (time span referring to the past 6 weeks) and stdMCTQ (referring to a "regular week").

Statistical Analyses

Bland-Altman plots. The relationship between the µMCTQ and the stdMCTQ was studied using Bland-Altman plots (Bland and Altman, 1986, 1999). The Bland-Altman plot is a graphical method to compare 2 measurements techniques: the differences between the measurements obtained by the 2 techniques are plotted against the average of the measurements. Horizontal lines are drawn at the mean difference and at the 95% limits of agreement, which are defined as the mean difference ±1.96 times the standard deviation of the differences. If these limits do not exceed the maximum "allowable" difference between the methods (i.e., the differences are not yet physiologically or clinically relevant), the 2 measurement methods are considered to be in agreement and can be used interchangeably. Finally, if some degree of correlation exists between the differences and the averages on the Bland-Altman plot, the over- or underestimation of one method versus the other increases/decreases depending on the absolute value or size of the measurement. Since the differences between measurements were not normally distributed, correlations were tested using Spearman's rho.

Correlations. Correlations between variables from the stdMCTQ and the µMCTQ were tested using Pearson's *r*, since all variables were nearly normally distributed (inspection of histograms).

Study 2: Validation of the µMCTQ against Activity and Melatonin Phase (DLMO)

Participants. Thirty participants (15 women, age [mean \pm standard deviation]: 22.7 \pm 3.6 years) were recruited by convenience sampling (mostly students recruited by flyers on campus). Exclusion criteria were irregular work schedules or shift work or a transmeridian flight during a 3-month period prior to study participation. All participants provided informed consent; the study was approved by the Ethics Committee of Ludwig Maximilian University (approval 517-15) and conducted in accordance with the Declaration of Helsinki.

Study Design. Because of reasons of feasibility, the study spanned the autumn DST change. The cohort was divided into 2 groups. Group 1 (G1; 9 women and 6 men) started the study shortly before the DST change in October 2015 and was monitored for 6 weeks. Group 2 (G2; 6 women and 9 men) started a week after the time change and was monitored for 4 weeks. One subject (from G1) did not fill in the questionnaire correctly and had to be excluded.

Participants filled out the µMCTQ once at study onset. Throughout the course of the study, they filled out an online version of the µMCTQ on a daily basis (similar to a sleep-log), but the daily assessments were used only to generate date-type data (work- or work-free day).

Measurements. In addition to the µMCTQ, actimetry and home DLMO data were collected for circadian phase determination. In G1, saliva samples were collected during the first week before the time change, and the $\mu MCTQ$ was filled in 8 days before the time change. In G2, saliva samples were taken in the second week after the time change, and the µMCTQ was answered 5 days after the time change.

Activity phase assessment. Actimeters are wristworn devices that measure locomotor activity by accelerometry. Devices (Daqtometers by Daqtix GmbH) were worn throughout the entire study period (G1: 6 weeks, G2: 4 weeks). Recordings from G1 before the DST change and the first week after the time change were excluded from the analyses. Activity was recorded at 1 Hz, and the average activity counts were stored every 30 s. For analyses, data were averaged into 10-min bins. Participants kept a diary about day types (work or work-free days). We used the software ChronoSapiens (Chronsulting UG; Roenneberg et al., 2015) to assess the daily phase of general locomotor activity measured by the wrist-worn actimeters. For every day, we calculated the acrophase of the best 1-harmonic, 24-h cosine fit (Ψ _Act). The advantage of this phase marker is that it does not rely on any other computations of the actimetry signal such as algorithms identifying sleep. Ψ _Act for every subject was calculated using daily averages across all days. Individual averages were also calculated separately for workdays (Ψ _Act_w) and work-free days (Ψ _Act_f). For the calculation of the general Ψ _Act results, we also included days not specified as work or work free. Data are expressed as local time.

DLMO assessment. DLMO was assessed at home in this study for reasons of feasibility. Home DLMOs have previously been reported to show a good correspondence to lab DLMOs (Burgess et al., 2015; Pullman et al., 2012), especially when participants' compliance is being objectively monitored (which, however, was not the case in our study). Participants had an appointment with study team members, during which they were instructed on how to collect saliva samples. The volunteers were asked to collect 7 saliva samples hourly once on a workday and once on a work-free day starting 6 h before their usual sleep timing. Subjects were told to close the blinds and turn off as many lights as possible 1 h before saliva collection started and were given blue light-blocking sunglasses to wear during the period of collection. They

were also instructed to dim screen lights of electronic devices, not to change their position at least 5 min before taking the sample, and to rinse their mouth with clear water before collection. They were also told not to eat chocolate or bananas or to drink coffee or alcohol during the period of saliva collection. A written version of the instructions was provided as well. Samples were collected using Salivette cotton swabs (Sarstedt AG & Co.). The participants wrote down the times of collection into a log provided by the study team, without further measures of compliance, and they kept the samples in the fridge (\sim 4°C) until bringing them to the laboratory (storage duration <7 days), where they were processed and kept at -20°C until further use (storage duration <3 months). The duration of sample storage was in accordance with manufacturer guidelines.

Chrono@work (Groningen, the Netherlands) analyzed the samples. Melatonin concentrations were assessed using direct saliva melatonin radioimmunoassay test kits (RK-DSM; Bühlmann Laboratories, Alere Health, Tilburg, the Netherlands). DLMO was calculated by linear interpolation between the time points before and after melatonin concentrations crossed and stayed above the threshold of 3 pg/mL. We opted for the fixed threshold method for sample size reasons (insufficient points to calculate baseline for some individuals), but similar results were seen using the threshold of 2 standard deviations above the baseline (see Suppl Fig. S8). The lower limit detection of the kit was 0.3 pg/mL. The intra-assay variability was 15.9% at low melatonin concentration (mean = 2.0 pg/mL, n = 17) and 13.1% at high melatonin concentration (mean = 24.5 pg/mL, n = 15). The interassay variability was 13.1% at low melatonin concentration (mean = 2.0 pg/mL, n = 16) and 15.0% at high melatonin concentration (mean = 21.4pg/mL, n = 16). DLMO on a workday could be calculated from 24 individuals and DLMO on a work-free day from 25 individuals.

Statistical Analyses. Shapiro-Wilk test and inspection of histograms were used to test the variables for normality, and all variables showed a normal distribution. Correlations between MSF_{sc} (as assessed by the μ MCTQ) and DLMO as well as the different Ψ _Act values were tested using Pearson's correlation. An alpha level of 0.05 was chosen. SPSS 24 and Graph-Pad Prism 6 were used for statistical analyses. Graphs were plotted using the R package ggplot2 (Wickham, 2016).

Test-Retest Reliability. We performed test-retest measurements in 37 subjects to test chronotype reliability (µMCTQ-MSF_{sc}) over 2 different time frames. We correlated assessments taken 56 to 63 or 14 to 18 days apart. Twenty students filled in the English version of the μ MCTQ (age [mean \pm SD]: 23.8 \pm 3.3 years; 40% female; interval between assessments: 56-63 days). Eighteen subjects recruited through snowball sampling filled in the German or English online versions of the μ MCTQ (age [mean \pm SD]: 33.5 \pm 7.8 years; 39% female; interval between assessments: 14-18 days).

RESULTS

Study 1: Validation of the µMCTQ against the std-**MCTQ**

Generally, sleep timing and MSF_{sc} corresponded well between the µMCTQ and stdMCTQ but showed systematic deviations toward earlier times in the uMCTQ in most of the assessed variables.

The µMCTQ estimated sleep onset on workdays and work-free days (SO_w and SO_f) earlier than the stdMCTQ (mean difference \pm SD: SO $_{\rm w}$ 24.7 \pm 28.4 min and $SO_f 30.8 \pm 37.7$ min; Fig. 3), with limits of agreement of less than 120 min: -30.9 to 80.3 min for workdays and -42.1 to 104.7 min for work-free days.

In contrast, the 2 tools produced similar estimates for sleep end on both workdays and workfree days (mean difference \pm SD: SE $_{\rm w}$ –2.7 \pm 19.0 min, $SE_f 2.0 \pm 28.7$ min; Fig. 4), with a limit of agreement of less than 60 min: SE_w –39.9 to 34.6 min, SE_t -54.3 to 58.3 min.

The average estimate of mid-sleep on workdays (MSW) and on work-free days (MSF) produced by the µMCTQ was less than 20 min earlier than that of the stdMCTQ (MSW mean difference \pm SD: 11.0 \pm 17.1 min; MSF mean difference \pm SD: 16.4 \pm 24.7 min; Fig. 5). MSF_{sc} was estimated as less than 20 min earlier by the $\mu MCTQ$ than by the stdMCTQ (MSF $_{sc}$ mean difference \pm SD: 16.3 \pm 27.1 min; Fig. 5). Their limits of agreement ranged from about 30 to 70 min: MSW -22.5 to 44.5 min, MSF -32.0 to 64.9 min; MSF_{sc} -36.8 to 69.4 min.

Correlations between the µMCTQ and stdMCTQ in terms of sleep onset (workdays and work-free days), sleep end (workdays and work-free days), MSF, MSW, and MSF_{sc} were all statistically significant and produced coefficients ranging from 0.89 to 0.95 (Suppl. Fig. S1). The same correlations are shown in Supplementary Figures S2 to S4 differentiated into age categories.

Study 2: Validation of the µMCTQ against Activity (Ψ _Act) and Melatonin Phase (DLMO)

MSF_{sc}, as an indicator of chronotype, showed a moderately positive correlation with Ψ _Act and Ψ _

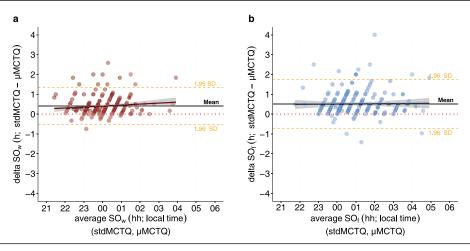


Figure 3. Bland-Altman plots of sleep onset on workdays (a) and work-free days (b) (method- differences on the y-axis versus method averages on the x-axis). The mean ± 1.96 SD limits of agreement, together with the regression line and pertinent confidence interval, are also depicted in the plot. Of the sample, (a) 6% and (b) 7% were out of the limits of agreement, respectively. Correlation analysis: (a) Spearman's rho 0.11 (ns); (b) 0.01 (ns). SO_w = sleep onset on workdays; SO_f = sleep onset on work-free days. N = 213 (study 1: validation of the ultra-short version of the Munich ChronoType Questionnaire [µMCTQ] against the standard MCTQ [stdMCTQ]).

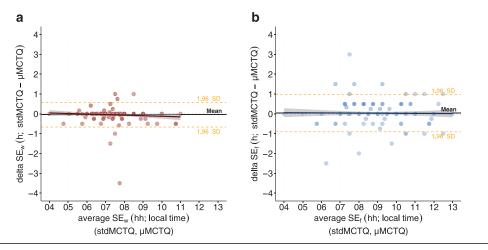


Figure 4. Bland-Altman plots of SE_w (a) and SE_f (b) (method differences on the y-axis versus method averages on the x-axis). The mean ± 1.96 SD limits of agreement, together with the regression line and pertinent confidence interval, are also depicted in the plot. Of the sample, (a) 3% and (b) 8 were out of the limits of agreement, respectively. Correlation analysis: (a) Spearman's rho -0.10 (ns); (b) -0.04 (ns). $SE_w = \text{sleep}$ end on workdays; $SE_t = \text{sleep}$ end on work-free days. N = 213 (study 1: validation of the ultra-short version of the Munich ChronoType Questionnaire [µMCTQ] against the standard MCTQ [stdMCTQ]).

 Act_f and a tendency to correlate with Ψ_Act_w (Fig. 6). Correlations of MSW and Ψ_{-} Act_w as well as MSF and Ψ _Act_f are provided in the supplementary material (Suppl. Fig. S6).

MSF_{sc} was positively associated with DLMO_f but not with DLMO_w (Fig. 7). Correlations of MSW and DLMO_w as well as of MSF and DLMO_f are provided in the supplementary material (Suppl. Fig. S7).

The µMCTQ showed good test-retest reliability within different time frames (see the Methods section: \sim 60 days: Pearson's r = 0.77, p < 0.001; \sim 14 days: Pearson's r = 0.79, p < 0.001; Suppl. Fig S9).

DISCUSSION

Our results show that both the µMCTQ and the stdMCTQ are valid instruments to assess PoE and that the 2 questionnaires show good correspondence with each other. The indicator of chronotype (MSF_{sc}), as measured by the µMCTQ, correlates with the timing of both melatonin (DLMO) and activity (Ψ Act; center of gravity of best fit). We also replicate previous findings, showing the stdMCTQ to correspond with actimetry measures (see the supplementary material).

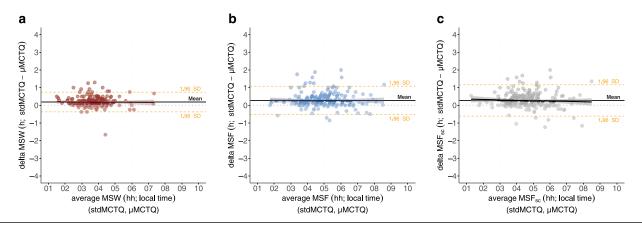


Figure 5. Bland-Altman plots of (a) MSW, (b) MSF, and (c) MSF_{sc} (method differences on the y-axis versus method averages on the x-axis). The mean \pm 1.96 SD limits of agreement, together with the regression line and pertinent confidence interval, are also depicted in the plot. Of the sample, (a) 7%, (b) 6%, and (c) 7% were out of the limits of agreement, respectively. Correlation analysis: (a) Spearman's rho -0.01 (ns); (b) -0.01 (ns); (c) -0.07 (ns). MSW = midpoint of sleep on workdays; MSF = midpoint of sleep on work-free days; MSF_{sc} = indicator of chronotype, midpoint of sleep on work-free days corrected for sleep debt accumulated over the workweek. N = 213 (study 1: validation of the ultra-short version of the Munich ChronoType Questionnaire [µMCTQ] against the standard MCTQ [stdMCTQ]).

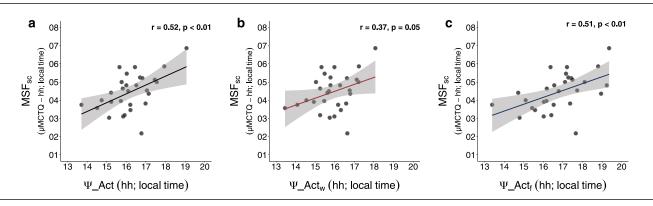


Figure 6. Associations between MSF_{sr} from the ultra-short version of the Munich ChronoType Questionnaire (μMCTQ) and actimetry phase (Ψ _Act). MSF_{sc} correlates significantly with Ψ _Act (a) as well as with Ψ _Act on work-free days (Ψ _Act, c), but tended to be associated with Ψ_{-} Act only measured on workdays (Ψ_{-} Act_w, b). The gray-shaded area around the regression line represents the 95% confidence interval. Results of Pearson correlations are provided in each graph. Data are expressed in local time for both variables. N = 29 (study 2: validation of the µMCTQ against actimetry).

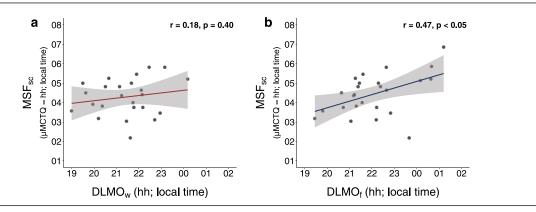


Figure 7. Association between MSF_{sc} from the ultra-short version of the Munich ChronoType Questionnaire (µMCTQ) and dim-light melatonin onset (DLMO). MSF_{sc} correlates significantly with DLMO measured on work-free days (DLMO₁, b) but not with DLMO measured on workdays (DLMOw, a). The gray-shaded area around the regression line represents the 95% confidence interval. Results of Pearson correlations are provided in each graph. Data are expressed in local time for both variables. n = 24 to 25 (study 2: validation of the µMCTQ against DLMO). DLMO was collected at home with no objective measures of compliance.

Correspondence between µMCTQ and stdMCTQ

Overall, all measurements derived from the µMCTQ and the stdMCTQ are in good agreement with each other. The µMCTQ yielded slightly earlier results for sleep onset and thus also for MSW, MSF, and MSF_{sc}. A likely reason for this systematic difference in onset timing is that the µMCTQ lists and asks about less detail on the "going-to-bedand-falling-asleep process." The µMCTQ does not lead people in and out of bed but asks directly for the time that participants usually fall asleep. Some participants may interpret this as the time when they were prepared to sleep and would therefore indicate an earlier time than actual sleep onset depending on their sleep latency. Alternatively, some people may misinterpret the detailed descriptions in the stdMCTQ explaining the different stages from going to bed to getting up (Fig. 1). For example, different people may have different concepts of what "sleep preparation" means. Does sleep preparation include sleep latency or not? Since the stdMCTQ calculates sleep onset by adding sleep latency to the time people indicate for preparing for sleep, latency might occasionally be added twice. The average sleep latency in the MCTQ database is 15.4 ± 15.2 min for free days and 18.8 ± 17.5 minutes for workdays. Notably, sleep end was not different between µMCTQ and std-MCTQ. An earlier estimate of sleep onset also influences MSW, MSF, and MSF_{sc}. A difference of less than 20 min for MSF_{sc} is lower than the median intraindividual variance of $\Psi_{-}Act_f$ observed in our study (MCTQ study median: 110 min, Q1-Q3: 40-250 min; µMCTQ study median: 126 min, Q1-Q3: 76-192 min) and is well within the variance of both questionnaires and other instruments.

Furthermore, correlations of the stdMCTQ with Ψ_{Act} (Suppl. Fig. S10) were not significantly stronger than those of the µMCTQ with the same measurement (Ψ _Act_f vs. stdMCTQ-MSF_{sc} and Ψ _Act_f vs. μ MCTQ-MSF_{sc}: Fisher's Z test, z = 0.26, ns; Ψ _Act vs. stdMCTQ-MSF_{sc} and Ψ _Act vs. μ MCTQ-MSF_{sc}: Fisher's Z test, z = -0.06, ns).

Both the µMCTQ and the stdMCTQ chronotype showed good test-retest reliability (see Suppl. Figs. S9 and S12). Regarding the stability of our chronotype estimation, we expect the indicator variable to vary across time. This is because we are estimating chronotype as circadian state. Transient state constructs, in contrast to more enduring trait dispositions, are expected to be susceptible to influences from the environment (Boyle et al., 2015). Yet, circadian phase (estimated both using MSF and DLMO) has been shown to be fairly reproducible over months (Kantermann and Eastman, 2018).

Validation of the µMCTQ against Actigraphy and **DLMO**

 μ MCTQ-MSF_{sc} correlated with DLMO and Ψ _Act on work-free days but was not significantly correlated with either of these markers on workdays. Since it is assumed that MSF_{sc} reflects an estimation of PoE less affected by social constraints, a weaker correlation with parameters gathered on workdays (DLMO_w, $\Psi_{Act_{w}}$) was expected. DST, which occurred during the participation of half the subjects, might have also contributed to the observed result. The µMCTQ assesses sleep behavior in the past 6 weeks, and the second group filled out the questionnaire and measured DLMO 1 week after the time change. The release from DST causes a delay in Ψ _Act and in sleep timing that is more gradual on workdays than on work-free days. It was also shown that the process of adjustment to the time change and how long this process takes is chronotype specific, with late types delaying more readily (Kantermann et al., 2007). The transition accentuates the misalignment between internal time and sleep on workdays, which may be reflected in our data.

Although DST might have influenced our results, µMCTQ-MSF_{sc} was still significantly correlated with DLMOf, similarly to what has been shown in other publications using the stdMCTQ. Even stronger correlations between MSF_{sc} (as per the stdMCTQ) and DLMO (home or lab) have been previously reported (Facer-Childs et al., 2019; Kantermann et al., 2015; Wright et al., 2013). However, when comparing the strength of correlations found in our study (µMCTQ validation) to the ones cited, a significant difference is detectable only between our data and data from Facer-Childs et al. (Fisher's z = -2.1, p < 0.05). Their data, however, show a wider range in DLMO and MSF_{sc}, as Facer-Childs et al. selected for extreme chronotypes. Correlations are known to be sensitive to data distribution (Goodwin and Leech, 2006), and therefore, we propose the observed differences in correlation strength to likely be attributed to this.

Study Limitations

The different age ranges in the cohorts should be noted, as study 2 was conducted in young students with a narrow age range. Nevertheless, correspondence between the questionnaires was good across different age categories (as shown in Suppl. Figs. S2 to S4). As the std-MCTQ was shown to be valid against Ψ _Act in a sample with a wide age range (18-81 years old; see supplementary material), and the µMCTQ corresponds well with the stdMCTQ

- in all aspects, we do anticipate similar validations at higher ages. Furthermore, the Korean stdMCTQ has already been shown to be valid against actimetry and sleep logs in a large sample of older adults (\geq 65 years, N =192, Ryu et al., 2018).
- In study 2, we acquired a small, homogeneous sample consisting of mostly young university students, which could potentially affect generalizability, and a release from DST occurred in half the subjects while recording actimetry data. Still, MSF_{sc} was significantly correlated to both DLMO and Ψ _Act when they were taken on work-free days, despite the small sample size.
- The µMCTQ was administered at the beginning of the study and refers to the previous 6 weeks, not corresponding with the time monitored by actimetry. The questionnaire was administered at the beginning to avoid a bias in subjects' responses, since daily assessments of sleep timing were also implemented.
- The µMCTQ in study 2 was administered in a translated German version without backtranslation into English (original language).
- The µMCTQ was reduced from the stdMCTQ in an intuitive manner, rather than using more objective dimension reduction techniques.
- DLMO assessment in study 2 was done at home with no measures of compliance, and self- reported collection times in DLMOs assessed at home can be rather inaccurate (Kudielka et al., 2003). However, studies have compared lab-based and home-measured DLMOs and found significant correlation between the 2 conditions regardless of measuring compliance (Burgess et al., 2015; Pullman et al., 2012). Furthermore, Pullman et al. (2012) considered the at-home assessments of DLMO to be satisfactorily accurate (compared with the corresponding lab measurements) in 62.5% to 75% of the cases.

Comparison between µMCTQ and stdMCTQ

As a shortened questionnaire, the µMCTQ naturally collects less information about peoples' sleep behavior than the stdMCTQ. The µMCTQ does not enable the estimation of sleep latency or inertia. Furthermore, in contrast to the stdMCTQ, the µMCTQ does not inquire about reasons for externally induced waking on work-free days other than an alarm clock (e.g., children, hobbies). Nonetheless, the µMCTQ also offers an advantage when compared with the stdMCTQ: it allows for the assessment of the circadian phase of people who mostly use alarm clocks on

work-free days by asking them to consider only those work-free days on which they do not use an alarm clock. We use sleep behavior on free days as a proxy for chronotype because it is less confounded by social constraints and therefore is a closer reflection of entrained phase. Only for this conceptual reason, we usually do not compute MSF_{sc} when subjects report using alarm clocks on free days in the stdMCTQ (Pilz et al., 2018; Roenneberg et al., 2012; data from study 1). However, several studies have computed chronotype based on people who use an alarm clock on free days. While this is theoretically possible, we strongly recommend stating it clearly in the publication. Individuals who use alarm clocks on work-free days show slightly later mid-sleep and MSF_{sc} than subjects who do not (Suppl. Fig. S5).

Time Frames of the Questionnaires

The stdMCTQ has been used for more than 15 years in studies with varying research questions and designs. Originally, it asked people about their sleep behavior "in a regular week." Depending on the specific study design and question, the time frame assessed by the stdMCTQ has been modified to fit the demands of the investigation. Here, we suggest using 6 weeks with the µMCTQ to obtain a stable assessment that can still accommodate, for example, seasonal changes. stdMCTQ ("a regular week") and µMCTQ (past 6 weeks) in study 1 used different time frames for assessment but still corresponded well. Only the µMCTQ was used in study 2.

CONCLUDING REMARKS

We have developed and validated an ultra-short version of an already well-established instrument, the stdMCTQ. Both the stdMCTQ and the µMCTQ can be used to estimate PoE and can serve as good alternatives to time-consuming and more expensive measurements such as sleep logs, actimetry, and DLMO. They allow the calculation of quantitative, not qualitative, variables and thereby permit a range of statistical operations impossible to conduct with categorical data.

The µMCTQ, in alignment with the stdMCTQ, represents a valuable tool to assess information about the human clock in a concise manner. It uses relevant questions established by the stdMCTQ but might offer a better alternative for large samples and longer study durations, which would benefit from shorter questionnaires that offer relevant information about sleep-wake behavior.

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AUTHOR CONTRIBUTIONS

T.R. and E.W. developed the µMCTQ. N.G., L.K.P., S.M., and T.R. designed the studies. L.K.P., N.G., D.L., and S.M. collected and organized the data for the µMCTQ validation and C.V., T.R., and E.W. did the same for the stdMCTQ validation. N.G., L.K.P., S.M., G.F., M.S., R.C., E.W., G.Z., C.V., and T.R. analyzed the data. N.G., L.K.P., S.M., and T.R. wrote the first draft of the article. All authors read, extensively revised, and approved the final article.

CONFLICT OF INTEREST STATEMENT

The authors have no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

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NOTE

Supplementary material is available for this article online.

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