Sleep in Adolescents: The Perfect Storm

Mary A. Carskadon, PhD

KEYWORDS
- Adolescence • Circadian rhythms • Sleep regulation
- Homeostatic pressure • Melatonin • Mood

The perfect storm metaphor applies to sleep patterns of adolescents in the sense that developmental trajectories of biopsychosocial factors conspire to limit the quantity of sleep for many adolescents, resulting in a number of negative consequences. A reduction in sleep amount from late childhood through the second decade has long been known; however, the weight of current evidence holds that sleep need does not decline across this span. Nevertheless, parents, pediatricians, and schoolteachers, it seems, long assumed that this sleep decline was an inevitable part of growing up and a normative expectation. This article will describe how the loss of sleep through adolescence is not driven by lower need for sleep but arises from a convergence of biologic, psychological, and socio-cultural influences.

SLEEP PATTERNS OF ADOLESCENTS

Adolescent sleep patterns have been surveyed by investigators in many countries from virtually every continent around the world, and a consistent finding is that the timing of bedtime on school nights gets later across the middle school and high school years (roughly ages 11 through 17 years).1–15 Rise times on school mornings, by contrast, tend to stay relatively consistent except in countries such as the United States where the starting time of school moves to an earlier hour at the transition to high school. Weekend sleep for teenagers tends to delay further, and the difference in amount of sleep reported for school days versus weekends becomes more pronounced as children pass into higher grades (ie, greater reported sleep on weeknights than school nights).

The most recent US poll of sleep patterns in adolescents was reported by the National Sleep Foundation in 2006, and collected self- and parent-reported sleep patterns from grades 6 through 12.16 These data serve as a good example of these general trends, as shown in Table 1. The young people interviewed in this telephone poll reported that average bedtime on the nights before school days were approximately 1.5 hours later from grade 6 to grade 12, and reported weekend bedtime delayed from 10:31 PM to 12:45 AM across this same grade span. Sixth graders...
reported going to bed about an hour later on weekend nights, and for 12th graders, the
weekend bedtime delay was about an hour and 45 minutes. The average reported rise
time on school mornings was 6:42 AM in grade 6 and 6:31 AM in grade 12. The reported
number of hours slept on school nights declined from 8.4 hours in the 6th grade
students to 6.9 hours in the 12th graders; reported weekend sleep was more consis-
tent from grades 6 through 11—about 9 hours—falling to 8.4 hours in grade 12. The
weekend extension of sleep time in this report was nearly an hour in the middle school
children (grades 6–8) and approached 2 hours for grade 11 students.

Amounts of sleep reported by adolescents vary across countries and regions;
however, the overall patterns of later sleep timing and diminished sleep across adoles-
cence is reported by most investigators. Reports of Korean youth, for example, indi-
cate they begin and end grades 6 to 12 with later school night bedtimes and less sleep
than those in the United States.15 Children were assessed with a survey administered
in the classroom querying usual sleep schedule. On average, the children reported:

- School night bedtime: grades 5–6 = 10:42 PM, grades 7–8 = 11:12 PM; grades
  9–10 = 12 AM; grades 11–12 = 12:54 AM
- School morning rise time: grades 5–6 = 7:18 AM; grades 7–8 = 7 AM; grades
  9–10 = 6:48 AM; grades 11–12 = 6:18 AM
- School night hours slept: grades 5–6 = 8.3; grades 7–8 = 7.6; grades 9–10 = 6.6;
  grades 11–12 = 5.4
- Weekend night bedtime: grades 5–6 = 11 PM, grades 7–8 = 11:30 PM; grades
  9–10 = 11:46 PM; grades 11–12 = 11:54 PM
- Weekend morning rise time: grades 5–6 = 8:06 AM; grades 7–8 = 8:54 AM; grades
  9–10 = 9:30 AM; grades 11–12 = 9:18 AM
- Weekend night hours slept: grades 5–6 = 9.0; grades 7–8 = 8.8; grades 9–10 = 9.0;
  grades 11–12 = 8.4.

An apparent difference in the older Korean children from the 11th and 12th graders
in the US poll16 was that the reported weekend bedtime was earlier than on school
nights. The authors note that many students (71.1%) took additional course work in
the evening, with classes lasting until midnight or later for 54.5% of the older students.

### Table 1

Sleep patterns reported by adolescent school children: National Sleep Foundation 2006 Sleep in America Poll

<table>
<thead>
<tr>
<th>Grade in School</th>
<th>6th</th>
<th>7th</th>
<th>8th</th>
<th>9th</th>
<th>10th</th>
<th>11th</th>
<th>12th</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>School Nights</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Bedtime (24-h)</td>
<td>2124</td>
<td>2152</td>
<td>2153</td>
<td>2215</td>
<td>2232</td>
<td>2251</td>
<td>2302</td>
</tr>
<tr>
<td>Rise Time (24-h)</td>
<td>0642</td>
<td>0635</td>
<td>0636</td>
<td>0628</td>
<td>0623</td>
<td>0623</td>
<td>0631</td>
</tr>
<tr>
<td>Average Sleep (Hours)</td>
<td>8.4</td>
<td>8.1</td>
<td>8.1</td>
<td>7.6</td>
<td>7.3</td>
<td>7.0</td>
<td>6.9</td>
</tr>
<tr>
<td><strong>Weekend Nights</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Bedtime (24-h)</td>
<td>2231</td>
<td>2305</td>
<td>2326</td>
<td>2353</td>
<td>0003</td>
<td>0025</td>
<td>0045</td>
</tr>
<tr>
<td>Rise Time (24-h)</td>
<td>0853</td>
<td>0912</td>
<td>0921</td>
<td>0954</td>
<td>0954</td>
<td>1006</td>
<td>0951</td>
</tr>
<tr>
<td>Average Sleep (Hours)</td>
<td>9.2</td>
<td>8.9</td>
<td>9.0</td>
<td>8.8</td>
<td>8.9</td>
<td>8.8</td>
<td>8.4</td>
</tr>
<tr>
<td>School Night–Weekend Hours Slept Difference</td>
<td>0.8</td>
<td>0.8</td>
<td>0.9</td>
<td>1.2</td>
<td>1.6</td>
<td>1.9</td>
<td>1.5</td>
</tr>
</tbody>
</table>

This pattern of late night classes on school days likely also explains why the Korean high school students reported earlier weekend than school-night bedtime on average.\textsuperscript{15} It should also be noted that the standard deviation for reported weekend bedtime was 3 to 4 hours, much greater than for other variables.

The issue of determining the sleep need of adolescents is challenging, since the definition of sleep need itself is in dispute. Early work by Carskadon\textsuperscript{4} started from a questionnaire assessment of school children in which the discrepancy between school day and weekend reported amount slept was zero, and the average amount was 10 hours a night. Of interest is the parallel finding of Iglowstein and colleagues\textsuperscript{17} showing a concurrence of weekend and weekday sleep amounts at about 10 hours from reports by mothers of Swiss children.\textsuperscript{17} In the longitudinal study of adolescents by Carskadon, time in bed was fixed at 10 hours a night, and the data showed that youngsters across the span of ages 10 to 17 slept about 9 hours and 20 minutes.\textsuperscript{18} The younger children were more likely to waken spontaneously in the laboratory, whereas the older children took a bit longer to fall asleep and were rarely spontaneously awake at the end of the 10 hours. Furthermore, assessment with the multiple sleep latency test (MSLT, an objective measure of sleepiness) showed that the younger children were more alert across the day, whereas the pubertal and postpubertal adolescents showed a midday trough of alertness in spite of the same amount of nocturnal sleep.\textsuperscript{19} In another study that may bear on the issue of sleep need of adolescents, Carskadon and colleagues\textsuperscript{20} showed that adolescents (ages 11 to 14.6 years) given a 10-hour sleep schedule for 10 to 14 nights and then studied in the laboratory for 3 consecutive 18-hour nights, slept nearly 12.5 hours on average the first night and by night 3 still slept 10.1 hours on average (standard deviation about 1 hour).

**ADOLESCENT DEVELOPMENT AND SLEEP REGULATION**

In describing the biologic regulation of sleep, current thinking uses the 2-process model, first proposed by Borbély.\textsuperscript{21} The 2 processes that comprise the model include a daily (circadian) rhythm of sleep propensity, thought to originate from the suprachiasmatic nucleus of the hypothalamus in mammals, and a sleep–wake pressure (homeostatic) system, for which a neuroanatomical locus has not been identified. In the case of the former process, the signal from the brain’s central clock is thought to affect sleep and arousal systems to help gate the timing of sleep. The sleep homeostatic system favors sleep as the hours of wakefulness are extended and favors waking as sleep is prolonged, thus transducing information about the length and amount of prior sleep or wake, rather than the time of day. The role of these two processes can be distinguished independent from one another under certain experimental conditions,\textsuperscript{22} but work interactively to regulate sleep from day to day.

Developmental changes in the circadian timing system were inferred by some from the delay in the timing of sleep onset, and 2 studies published in 1993 gave initial evidence for the hypothesis that a phase delay occurs in association with puberty. For example, Andrade and colleagues\textsuperscript{23} found in a longitudinal study that later sleep times occurred for adolescents at a more mature Tanner stage\textsuperscript{24} than others. Carskadon and colleagues\textsuperscript{25} reported in a cross-sectional study using self-reported sleep and pubertal stage that 6th grade girls who rated themselves more mature had a more evening type score on a measure of circadian phase preference. A subsequent follow-up cross-sectional study in which adolescents’ sleep–wake schedules were constrained for 2 weeks confirmed that circadian phase as evaluated with the objective measure of dim light melatonin onset (DLMO) phase was positively
correlated with Tanner stage. In other words, the DLMO was later for participants rated more mature, indicating that the onset of the biologic night is later for more mature teens.

Several hypotheses have been put forward to explain the features of the circadian timing system that may underlie this developmental change. A review of adolescent changes in sleep and circadian rhythms by Hagenauer and colleagues26 noted that signs of a juvenile phase delay were found in several mammalian species in addition to humans. Table 2 highlights these findings. The occurrence of a circadian delay around the time of puberty for a number of species lends credence to the hypothesis that this phenomenon arises from intrinsic biologic processes rather than as a response to social/behavioral exposures. Whether or how the phase delay is linked to reproductive development is unknown.

If the adolescent phase delay is inherent, what mechanisms might explain the phenomenon? One hypothesis is that circadian period (ie, the internal day length) may become longer during adolescence. Pubertal male rats were found in 1 study to have longer intrinsic periods than adults,27 and data from people show a longer period in teens than in adults.28 An alternate explanation is that the circadian timing system manifests an altered sensitivity to phase-dependent effects of light; in particular, the pacemaker may become more sensitive to evening light’s phase-delaying effects during adolescence than in childhood.26 One animal study that offers support for this hypothesis showed an exaggerated delay of rhythms to light occurring at the phase-delaying time in pubertal mice compared with adult mice.29 This hypothesis depends on an underlying biologic change that is then affected by behavioral exposures that arise during adolescence (ie, staying awake later, thus experiencing more evening light). A converse to this hypothesis is that adolescents become less sensitive to phase-advancing (morning) light or experience a change in the shape of the phase response curve. Preliminary data from one study of human adolescents provide a bit of evidence supporting this hypothesis.30

The influence of the sleep–wake homeostasis system is well understood, although specific neuroanatomical or neurochemical constituents are not yet fully established, nor has a link between adolescent maturation and this physiologic process been identified. Thus, whether the homeostatic process might be influenced by hormonal input, physical growth, brain maturation, or other factor(s) is unknown. On the other hand,

<table>
<thead>
<tr>
<th>Species</th>
<th>Amount of Delay</th>
<th>Rhythms Delayed</th>
<th>Sex Difference</th>
</tr>
</thead>
<tbody>
<tr>
<td>Human (Homo sapiens)</td>
<td>1–3 h</td>
<td>Sleep, melatonin</td>
<td>Males&gt;females</td>
</tr>
<tr>
<td>Rhesus monkey (Macaca mulatta)</td>
<td>2 h</td>
<td>Activity</td>
<td>Only females studied</td>
</tr>
<tr>
<td>Degu (Octodon degus)</td>
<td>3–5 h</td>
<td>Activity, sleep(?)</td>
<td>Males&gt;females</td>
</tr>
<tr>
<td>Laboratory rat (Rattus norvegicus)</td>
<td>1–4 h</td>
<td>Activity</td>
<td>Males&gt;females</td>
</tr>
<tr>
<td>Laboratory mouse (Mus musculus)</td>
<td>1 h (?)</td>
<td>Activity, corticosterone</td>
<td>Only females studied</td>
</tr>
<tr>
<td>Fat sand rat (Psammomys obesus)</td>
<td>0–3 h to 10–14 h</td>
<td>Oxygen consumption, temperature</td>
<td>Sex unspecified</td>
</tr>
</tbody>
</table>

*See Hagenauer and colleagues26 for more information about these adolescent phase delays.*
several evaluations of adolescent sleep physiology provide evidence for developmental changes in this sleep regulatory process. The theoretical model of sleep–wake homeostasis has a long history and a significant amount of experimental data supporting the model, based on sleep slow frequency electroencephalogram (EEG) data from animal and human models. As waking is prolonged across a day or longer, slow-frequency EEG amplitude and incidence increase; as sleep unfolds across the night, slow-frequency EEG amplitude and incidence decay. Several studies have shown that adolescent development is not associated with a change in the decay of sleep homeostatic pressure (indexed by EEG slow-wave activity in sleep) but remains stable across early adolescent development (ages about 9–10 through about ages 14–15 years) and up to age 18 years in another study. On the other hand, preliminary data from a recent analysis of sleep EEG slow-wave activity in another adolescent cohort (ages 15–16 through about ages 17–19 years) demonstrate a longer time constant for the decay across the night in several brain regions. One interpretation of the latter findings is that the sleep recovery process is unchanged in early adolescents and that certain parts of the brain may require more sleep to recover in older teens than when younger. This set of findings reinforces the likelihood that sleep need does not decline in adolescents.

Whereas the dissipation of sleep homeostatic pressure may speak to the recovery function of sleep, the other limb of the homeostatic process describes how the pressure to sleep or sleep propensity builds across the waking day. Jenni and colleagues modeled the accumulation of sleep homeostasis in early adolescent humans (ages 9–14 years) who were either pre-/early pubertal or postpubertal. This analysis showed a longer time course for the build up of sleep pressure in the more mature group, indicating that the more mature adolescents may find it easier to stay awake longer than the less mature. This hypothesis was confirmed by analyzing speed of falling asleep on the multiple sleep latency test in a group of pre-/early pubescent versus a group of postpubertal adolescents across 36 hours of extended waking. The less mature adolescents fell asleep faster than the more mature on the tests occurring 14.5 and 16.5 hours after waking.

In summary, current data on the maturation of the biologic regulation of sleep during adolescence show that

- The circadian timing system undergoes a phase delay
- The dissipation of sleep homeostatic pressure does not change until late adolescence when it shows evidence of slowing
- The accumulation of sleep homeostatic pressure slows during puberty.

These findings support the notion that the need for sleep is stable (or increases) across adolescent development and that the delay of sleep timing is supported by the circadian shift as well as the slowing of sleep homeostatic pressure accumulation.

PSYCHOSOCIAL FACTORS AFFECTING ADOLESCENT SLEEP

Many psychosocial factors affect sleep patterns in adolescents and contribute to the phase delay and the decline of time slept. A few examples will follow, as well as an examination of how they interact with these biologic regulatory processes. A primary psychodevelopmental task of adolescents is to achieve independence in many areas of their lives. One area where this striving for autonomy is displayed is the decision of when to go to sleep. One sleep habits survey from the early 1980s showed that a significantly higher percentage of children ages 12 or 13 years reported setting their own bedtime on school nights (eg, 19% of 13-year-olds) than did younger children.
(eg, 50% of 10-year-olds) 10- to 12-year-old children and that reported bedtimes were later for the older children. The other end of school night sleep—waking up—showed an opposite trend: more older children reported needing a parent or alarm to wake them than did the younger children, who were likelier to report that they “just wake up.” Thus, the adolescents who actualize autonomy by staying up late seem to ride the wave of the circadian phase delay and to avail themselves of the relative emancipation from rapid sleep pressure accumulation. Yet, they were sleeping less, and rising early required external intervention to wake up.

Two recent studies highlight the importance of parental-set bedtimes. Gangwisch and colleagues examined data in a large epidemiologic data set gathered from adolescents (grades 7–12) in the United States during the mid-1990s. The analysis showed that young people whose parent set their bedtime at midnight or after relative to those with bedtimes set at 10 PM or earlier, were significantly more likely to suffer from depression or suicidal ideation. The authors also found that this association was mediated by total sleep time; thus, those with earlier bedtimes reported sleeping more and were less likely to be depressed or experience suicidal ideation. A smaller, focused study of South Australian adolescents found that adolescents who reported a parental-set bedtime versus those without a set bedtime reported earlier bedtimes, more sleep, and less daytime fatigue experienced. These 2 studies provide evidence that parental intervention by setting a bedtime results in extended sleep and that the combination of earlier bedtime and extended sleep contribute to more positive outcomes for depression, sleepiness, and fatigue. These findings may also indicate that such behavioral interventions as having a fixed, earlier bedtime may help mitigate the circadian phase delay associated with adolescent maturation, likely through preventing light to the circadian timing system at the delay-sensitive evening phase.

Screen time, technology use, and social engagement in the evening become more available as youngsters pass through adolescence. The US National Sleep Foundation poll of adolescents reported that electronic devices (ie, electronic music devices, television, electronic/video games, cell phone, telephone, computer, Internet access) in the bedroom become more common for students from grade 6 to grade 12; the median number of such devices in bedrooms of younger adolescents was 2 versus a median of 4 in the older teens. A recent review by Cain and Gradisar notes that the preponderance of studies report shorter, later, and/or more disrupted sleep, as well as such daytime consequences as sleepiness or disruptive behavior, for children and adolescents as TV watching, computer/Internet/electronic games use, or mobile phone use in the evening before bedtime is greater. These activities are arousing in and of themselves and usually more easily accessed by the older adolescents, taking advantage of increased accessibility of technology and of the changes to the sleep regulatory systems that make it easier to stay away later. Indeed, to the extent that the activities involve light exposure—perhaps particularly blue-spectrum light exposure to which the circadian clock may have greater sensitivity—evening light has the phase-specific effect of delaying circadian rhythms, thus pushing sleep timing later.

The impact of school schedules of adolescents is most often to restrict sleep by requiring an early rising time. As noted previously, however, school schedules can delay sleep further, such as described for Korean adolescents enrolled in evening classes to improve preparation for academic placement tests. Evening homework assignments often carry a similar challenge for adolescents in the United States. More attention has been directed toward school starting times because of the tendency for many school systems in the United States to start the school day earlier as children get older, thus requiring teens to arise at an early hour relative to their typical bedtimes, circadian phases, and need for sleep.
One study examined sleep, sleepiness, and circadian rhythms in teens for whom the transition from 9th grade to 10th grade involved an advance of the school start time from 8:25 AM to 7:20 AM.44 Table 3 highlights findings from this study. Actigraphically monitored sleep data across 2 weeks confirmed that, although the students woke up significantly earlier on school days in tenth than ninth grade, they did not go to sleep earlier and consequently experienced less sleep on average. The general level of sleepiness in these students is shown with the daytime sleep latency data, showing a moderate level of sleepiness in 9th grade, and a significant change in 10th grade to a level considered severely sleepy for the sleep latency at 0830. The 0830 assessment was also associated with the occurrence of rapid eye movement (REM) sleep within 10 or 15 minutes of sleep onset, (a finding often used to aid in the diagnosis of narcolepsy) in about one half of the participants. The study also showed that the time of the DLMO was nearly 30 minutes later on average in the students when assessed in 10th grade; indeed, the DLMO time in the 10th graders was significantly later (2146 vs 2036) for those with the short sleep latency and REM sleep in the morning nap, interpreted as evidence that the circadian timing system of these teens was phase delayed relative to the early start time and favored sleep during the morning hours they were scheduled to attend school.44 The association of a circadian phase delay, excessive morning sleep tendency, and REM sleep in morning trials for one-half of the adolescents studied is remarkable and raises the level of concern for early morning classes.

THE PERFECT STORM: SLEEP BEHAVIOR AND TROUBLING OUTCOMES

Fig. 1 illustrates the confluence of factors that combine for adolescents in the 21st century to reduce time spent sleeping on school nights below a healthful amount, with waking and expected school performance timed to occur at an inappropriate circadian phase. As reviewed previously, bioregulatory and psychosocial forces collude to push sleep onset later, yet schools are timed to begin earlier across adolescence, and sleep time is compressed as a consequence. The list of negative outcomes associated with insufficient sleep is lengthy and includes sleepiness and mood disturbances noted previously, as well as inattention, poor grades, behavior problems, substance use, driving crashes, overweight, and immune system compromise. For some

<table>
<thead>
<tr>
<th>School Nights</th>
<th>Weekend Nights</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Actigraph Sleep Data</strong></td>
<td><strong>Actigraph Sleep Data</strong></td>
</tr>
<tr>
<td>Sleep onset</td>
<td>2242 (36)</td>
</tr>
<tr>
<td>Sleep offset</td>
<td>0626 (28)</td>
</tr>
<tr>
<td>Time slept</td>
<td>429 (47)</td>
</tr>
<tr>
<td><strong>Daytime Sleep Latency</strong></td>
<td><strong>Daytime Sleep Latency</strong></td>
</tr>
<tr>
<td>0830</td>
<td>10.9 (6.5)</td>
</tr>
<tr>
<td>1030</td>
<td>10.7 (5.9)</td>
</tr>
<tr>
<td>1230</td>
<td>11.0 (5.5)</td>
</tr>
<tr>
<td>1430</td>
<td>11.5 (6.1)</td>
</tr>
<tr>
<td>Melatonin onset time</td>
<td>2024 (57)</td>
</tr>
</tbody>
</table>

* Statistically significant difference between 9th and 10th grade
adolescents, the issue can present as sleep-onset insomnia that may be associated with the circadian phase delay. When combined with lack of motivation, depressed mood, and fatigue, a depressive disorder is often the initial assumption. An approach that targets sleep timing and phase adjustment is likely to be ameliorative.

At the end of the day, the pediatrician can have positive impact by intervening in any of a number of levels:

- Encourage local school districts to consider starting the school day later for adolescents
- Recommend that the local school district also limit late-evening activities
- Galvanize schools to provide instructional information about sleep and circadian rhythms
- Support and encourage parents to identify and set an appropriate bedtime
- Encourage teens to avoid light and stimulating activities in the evening and to get light exposure in the morning
- Empower adolescents to make informed choices about their sleep schedules
- Remind families of the utility of a relaxing presleep ritual.

Fig. 1. This figure illustrates the timing of sleep from preadolescence through adolescent development, highlighting the factors that affect sleep as described in the text. Thus, sleep is relatively long and timed at an early hour for preadolescents, but maturational changes to intrinsic bioregulatory factors—the circadian phase delay arising from the circadian timing system and a slowed rise of sleep pressure stemming from sleep-wake homeostasis—push for a delay of the timing of sleep. Such psychosocial factors as self-selected bedtimes, response to academic pressure, and the availability and use of technology and social networking in the evening also push for a delay in the timing of sleep. Note that the length of sleep is not affected by these processes. Societal pressures that push for an early rise time—most notably an early start to the school day—are the forces that limit amount of time available for sleep. As a consequence, adolescents sleep too little and are asked to be awake at an inappropriate circadian phase.

adolescents, the issue can present as sleep-onset insomnia that may be associated with the circadian phase delay. When combined with lack of motivation, depressed mood, and fatigue, a depressive disorder is often the initial assumption. An approach that targets sleep timing and phase adjustment is likely to be ameliorative.

At the end of the day, the pediatrician can have positive impact by intervening in any of a number of levels:

- Encourage local school districts to consider starting the school day later for adolescents
- Recommend that the local school district also limit late-evening activities
- Galvanize schools to provide instructional information about sleep and circadian rhythms
- Support and encourage parents to identify and set an appropriate bedtime
- Encourage teens to avoid light and stimulating activities in the evening and to get light exposure in the morning
- Empower adolescents to make informed choices about their sleep schedules
- Remind families of the utility of a relaxing presleep ritual.
ACKNOWLEDGMENTS

I thank the many colleagues, fellows, staff, and families who have contributed to the research from my group that is summarized in this paper. In addition, I acknowledge support from the National Institutes of Health, including MH45945, MH52415, MH58879, HL71120, MH076969, MH079179.

REFERENCES


