

Original Article  
Ophthalmology



# Effect of Ambient Light Exposure on Ocular Fatigue during Sleep

Young-Woo Suh ,<sup>1</sup> Kun-Hoo Na ,<sup>2</sup> Soh-Eun Ahn ,<sup>3</sup> and Jaeryung Oh <sup>1</sup>

<sup>1</sup>Department of Ophthalmology, Korea University College of Medicine, Seoul, Korea

<sup>2</sup>Department of Ophthalmology, Yeoncheon-gun Health Center and County Hospital, Yeoncheon, Korea

<sup>3</sup>Sungmo Eye Hospital, Busan, Korea

OPEN ACCESS

Received: Nov 22, 2017

Accepted: Apr 4, 2018

Address for Correspondence:

Jaeryung Oh, MD, PhD

Department of Ophthalmology, Korea University Anam Hospital, Korea University College of Medicine, 73 Incheon-ro, Seongbuk-gu, Seoul 02841, Korea.  
E-mail: ojr4991@korea.ac.kr

© 2018 The Korean Academy of Medical Sciences.

This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (<https://creativecommons.org/licenses/by-nc/4.0/>) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

ORCID iDs

Young-Woo Suh   
<https://orcid.org/0000-0003-4857-206X>  
Kun-Hoo Na   
<https://orcid.org/0000-0003-2893-0974>  
Soh-Eun Ahn   
<https://orcid.org/0000-0002-8523-8936>  
Jaeryung Oh   
<https://orcid.org/0000-0002-1036-6562>

Funding

This study is supported by Korea Ministry of Environment as "The Environmental Health Action Program (2012001350010)." The funding organization had no role in the design or conduct of this research.

## ABSTRACT

**Background:** To investigate the influence of nocturnal ambient light on visual function and ocular fatigue.

**Methods:** Sixty healthy subjects (30 men and 30 women) aged 19 through 29 years with no history of ocular disease were recruited. All subjects spent 3 consecutive nights in the sleep laboratory. During the first and second nights, the subjects were not exposed to light during sleep, but during the third night, they were exposed to ambient light, measuring 5 or 10 lux at the eye level, which was randomly allocated with 30 subjects each. The visual function and ocular fatigue were assessed at 7 a.m. on the 3rd and 4th mornings, using best-corrected visual acuity, refractive error, conjunctival hyperemia, tear break-up time, maximal blinking interval, ocular surface temperature, and subjective symptoms reported on a questionnaire.

**Results:** Three men and three women subjects failed to complete the study (4 in the 5 lux; 2 from the 10 lux). For the entire 54 subjects, tear break-up time and maximal blinking interval decreased ( $P = 0.015$ ;  $0.010$ , respectively), and nasal and temporal conjunctival hyperemia increased significantly after sleep under any ambient light ( $P < 0.001$ ;  $0.021$ , respectively). Eye tiredness and soreness also increased ( $P = 0.004$ ;  $0.024$ , respectively). After sleep under 5 lux light, only nasal conjunctival hyperemia increased significantly ( $P = 0.008$ ). After sleep under 10 lux light, nasal and temporal conjunctival hyperemia, eye tiredness, soreness, difficulty in focusing, and ocular discomfort increased significantly ( $P < 0.05$ ).

**Conclusion:** Nocturnal ambient light exposure increases ocular fatigue. Avoiding ambient light during sleep could be recommended to prevent ocular fatigue.

**Keywords:** Asthenopia; Dry Eye Syndrome; Eye; Fatigue; Lighting; Sleep

## INTRODUCTION

As industry and economy progress, the nocturnal exposure to bright light increases. The night is no longer dark, particularly in densely populated areas. The amount of light emitted and entering into bedrooms during nocturnal sleep from nearby buildings, electric billboards, and streetlamps is increasing as well. If this unnecessary nocturnal light causes human or ecosystem harm, it is considered light pollution. Light pollution can affect human

**Disclosure**

The authors have no potential conflicts of interest to disclose.

**Author Contributions**

Conceptualization: Oh J. Methodology: Oh J, Suh YW. Data curation: Na KH, Ahn SE, Suh YW. Formal analysis: Oh J. Software: Na KH, Ahn SE, Suh YW. Validation: Na KH, Ahn SE, Suh YW. Investigation: Na KH, Ahn SE, Suh YW. Writing - original draft: Suh YW. Writing - review & editing: Oh J.

physiology and disturb the circadian rhythm, which influences mood, vitality, and basal temperature by regulating hormones.<sup>1,2</sup> Reportedly, circadian rhythm disruption affects the neuroendocrine system and induces obesity, diabetes, depression, and cancer.

The light intensity affecting circadian rhythm is perceived by photosensitive retinal ganglion cells (PRGC), while the shape and movement of objects are recognized with retinal photoreceptor cells.<sup>3</sup> The eye is the first sensory organ to perceive the light-dark environment, but the influence of retinal nocturnal light exposure is not thoroughly understood. Nocturnal light exposure induced refractive errors in animal studies<sup>4</sup>; whether ambient light at night also induces refractive error in humans is controversial.<sup>5,6</sup> Bright light at night could disturb normal sleep and subsequently induce systemic and ocular fatigue. However, the effect of dim nocturnal light on ocular fatigue is not well established.

Ocular fatigue decreases reading performance.<sup>7</sup> It also decreases the accuracy of eye movement and saccadic speed.<sup>8</sup> If ambient nocturnal light exposure deteriorates visual function or induces ocular fatigue, then it may cause discomfort and decrease daily work efficiency. The tolerable level of nocturnal light for avoiding ocular fatigue and maintaining visual function is similarly unknown. The present study investigates the influence of nocturnal ambient light at 5 and 10 lux illumination on visual function and ocular fatigue.

**METHODS**

Sixty healthy subjects (30 males and 30 females) aged 19 through 29 years with no history of ocular disease were recruited. Prior to enrollment, best-corrected visual acuity with Snellen chart, and intraocular pressure with non-contact tonometry (CT-80A, Topcon Co., Tokyo, Japan) were measured on the participants. An alternate cover test at far and near distances, slit lamp examination, and fundus examination with fundus photography (TRC-NW8, Topcon Co., Tokyo, Japan) were performed. Subjects with a best-corrected visual acuity less than 20/20, corneal opacity, cataract, ocular hypertension, strabismus, and any fundus photographic abnormality were excluded.

Subjects spent 3 consecutive nights in the sleep laboratory of Korea University Hospital. During the day, subjects were educated to behave normally, taking care to avoid strenuous exercise, napping, excessive caffeine intake, alcohol consumption, and other behaviors potentially affecting sleep quality. During the first and second nights, subjects were not exposed to ambient light during sleep, but during the third night, they were. The light intensity was measured at the subjects' eyes, and the subjects were randomly allocated to one of two groups (30 subjects each), receiving 5 or 10 lux of light during sleep at night. Visual function and ocular fatigue were evaluated at 7 a.m. on the 3rd and 4th mornings. Sleep structure can become distorted in a laboratory setting during the first night of sleep (first-night effect),<sup>9</sup> therefore the visual function and ocular fatigue examinations were not performed after the 1st night (2nd morning). The visual function was evaluated using the best-corrected visual acuity, and refractive error. Refractive error was measured using an autorefractor (RK-F1, Canon, Tokyo, Japan). The objective ocular fatigue tests included the tear break-up time (TBUT), maximal blinking interval, ocular surface temperature, and conjunctival hyperemia. A questionnaire was administered to subjects for subjective assessment of fatigue. For the TBUT measurement, the tear film was stained with a fluorescein paper, and development of the tear film break was observed using the slit lamp;

the elapsed time was measured 3 times and averaged to calculate the TBUT. Maximal blinking interval was obtained by measuring the maximal time that the subject was capable of keeping the eye open without blinking. The central corneal temperature was measured at the ocular surface using the Thermal imaging system (FLIR E50, FLIR Systems, Inc., Wilsonville, OR, USA). Conjunctival hyperemia was determined on a 100-point scale previously described by Schulze et al.<sup>10</sup> All measurements were performed on the right eye. The ocular fatigue questionnaire comprised 10 items that are the most frequently reported symptoms of ocular fatigue.<sup>11</sup> Each item was answered using a 7-point scale (0–6). A score of 0 corresponded to no symptom, and a score 6 corresponded to an extremely severe symptom. This questionnaire had been reported previously to assess subjective ocular fatigue.<sup>12</sup>

### Statistical analysis

Statistical analysis was performed using SPSS 20.0 (SPSS Inc., Chicago, IL, USA). Visual function and ocular fatigue parameter differences between the 3rd and 4th mornings were compared using the Wilcoxon signed-rank test. The magnitude of change in visual function and ocular fatigue between subjects exposed to 5 lux and 10 lux during 3rd nocturnal sleep period were compared using the Mann-Whitney U test. A *P* value of < 0.05 was considered statistically significant.

### Ethics statement

Informed consent was obtained from all study participants after explanation of the nature and consequences of the study, and the study protocol was reviewed and approved by the Institutional Review Board of Korea University Anam Hospital (approval No. ED12261) and adhered to the tenets of the Declaration of Helsinki.

## RESULTS

Six subjects failed to complete the 3-night protocol. Among these 6 subjects, 4 was allocated to receive 5 lux exposure and the other 2 to receive 10 lux exposure. Excluding those subjects, the data of 54 subjects were analyzed. **Table 1** summarizes the post-sleep visual function and ocular fatigue without illumination and at 5 or 10 lux of illumination. The best-corrected visual acuity and refractive error did not change. Nasal and temporal conjunctival hyperemia increased significantly ( $P < 0.001$ ; 0.021 respectively), and TBUT and maximal blinking interval decreased significantly ( $P = 0.015$ ; 0.010, respectively). Among the subjective parameters, eye tiredness and soreness were increased significantly ( $P = 0.004$ ; 0.024, respectively). Difficulty in focusing also increased with marginal significance ( $P = 0.056$ ).

Visual function and ocular fatigue changes of subjects exposed to 5 lux illumination during the 3rd night are summarized in **Table 2**. Nasal conjunctival hyperemia increased significantly ( $P = 0.008$ ). But no other subjective and objective parameters changed.

**Table 3** summarizes the visual function and ocular fatigue after sleep without light exposure and under 10 lux illumination. Both nasal and temporal conjunctival hyperemia increased significantly ( $P = 0.016$ ; 0.041, respectively). TUBT decreased with marginal significance ( $P = 0.061$ ). Based on the questionnaire, eye tiredness, soreness, difficulty in focusing and vision discomfort increased significantly ( $P < 0.05$ ). The amounts of change in visual function and ocular fatigue after sleep under 5 and 10 lux exposure were compared in **Table 4**. There was no statistically significant difference between 5 and 10 lux illumination.

**Table 1.** The post-sleep visual function and ocular fatigue without illumination and at 5 or 10 lux of illumination

Parameters	No illumination		5 or 10 lux illumination		P value
	Mean	SD	Mean	SD	
Objective parameters					
Best corrected visual acuity	1.00	0.00	1.00	0.00	1.000
Refractory errors, diopter	−3.07	2.56	−3.07	2.60	0.904
Conjunctival hyperemia					
Nasal	16.85	8.43	20.93	9.57	< 0.001
Temporal	17.96	8.33	20.93	9.96	0.021
TBUT	4.22	1.78	3.82	1.77	0.015
Maximal blinking interval	12.74	9.85	10.77	6.09	0.010
Ocular surface temperature	34.59	0.82	34.52	0.90	0.259
Subjective parameters					
Tired eyes	1.61	1.42	2.17	1.38	0.004
Sore/aching eyes	0.50	0.84	0.78	1.00	0.024
Irritated eyes	0.98	1.39	1.24	1.48	0.142
Watery eyes	1.15	1.43	1.15	1.28	0.975
Dry eyes	1.69	1.72	1.80	1.45	0.744
Eyestrain	0.43	0.84	0.48	0.84	0.399
Hot/burning eyes	0.28	0.81	0.33	0.75	0.430
Blurred vision	0.56	1.04	0.50	1.00	0.729
Difficulty focusing	0.28	0.74	0.50	0.77	0.056
Vision discomfort	0.35	0.78	0.46	0.77	0.090

SD = standard deviation, TBUT = tear break-up time.

**Table 2.** Visual function and ocular fatigue of subjects exposed to 5 lux illumination during the 3rd night

Parameters	No illumination		5 lux illumination		P value
	Mean	SD	Mean	SD	
Objective parameters					
Best corrected visual acuity	1.00	0.00	1.00	0.00	1.000
Refractory errors, diopter	−2.82	2.66	−2.79	2.76	0.279
Conjunctival hyperemia					
Nasal	16.54	7.45	20.77	7.96	0.008
Temporal	17.69	7.65	20.00	8.49	0.238
TBUT	4.50	1.61	4.15	1.71	0.131
Maximal blinking interval	14.90	13.15	12.29	7.85	0.091
Ocular surface temperature	34.64	0.78	34.58	0.90	0.777
Subjective parameters					
Tired eyes	1.65	1.38	2.12	1.42	0.126
Sore/aching eyes	0.65	0.98	0.88	1.11	0.256
Irritated eyes	1.12	1.53	1.46	1.50	0.124
Watery eyes	1.31	1.52	1.27	1.19	0.964
Dry eyes	1.62	1.50	1.85	1.46	0.430
Eyestrain	0.58	0.99	0.54	0.81	0.874
Hot/burning eyes	0.31	0.97	0.46	0.95	0.157
Blurred vision	0.73	1.19	0.54	0.99	0.344
Difficulty focusing	0.50	0.99	0.62	0.80	0.564
Vision discomfort	0.65	1.02	0.62	0.94	1.000

SD = standard deviation, TBUT = tear break-up time.

## DISCUSSION

Light information during diurnal and nocturnal periods is transmitted to the suprachiasmatic nuclei (SCN) through the PRGC.<sup>13</sup> The SCN is thought to contain the human circadian clock and control the daily pattern of most physiologic processes.<sup>14,15</sup> It regulates hormones including cortisol, serotonin, and melatonin, and it controls the sleep-wake cycle, which is essential for systemic health.<sup>16</sup> When the eye is exposed to light nocturnally, melatonin secretion is suppressed by the SCN.<sup>16,17</sup> Melatonin is considered one of the most

**Table 3.** Visual function and ocular fatigue of subjects exposed to 10 lux illumination during the 3rd night

Parameters	Without illumination		10 lux illumination		P value
	Mean	SD	Mean	SD	
Objective parameters					
Best corrected visual acuity	1.00	0.00	1.00	0.00	1.000
Refractory errors, diopter	−3.31	2.49	−3.35	2.46	0.414
Conjunctival hyperemia					
Nasal	17.14	9.37	21.07	11.00	0.016
Temporal	18.21	9.05	21.79	11.24	0.041
TBUT	3.96	1.91	3.52	1.79	0.061
Maximal blinking interval	10.73	4.66	9.36	3.39	0.060
Ocular surface temperature	34.54	0.86	34.46	0.90	0.165
Subjective parameters					
Tired eyes	1.57	1.48	2.21	1.37	0.013
Sore/aching eyes	0.36	0.68	0.68	0.90	0.029
Irritated eyes	0.86	1.27	1.04	1.45	0.481
Watery eyes	1.00	1.36	1.04	1.37	0.902
Dry eyes	1.75	1.94	1.75	1.46	0.836
Eyestrain	0.29	0.66	0.43	0.88	0.271
Hot/burning eyes	0.25	0.65	0.21	0.50	0.914
Blurred vision	0.39	0.88	0.46	1.04	0.624
Difficulty focusing	0.07	0.26	0.39	0.74	0.024
Vision discomfort	0.07	0.26	0.32	0.55	0.008

SD = standard deviation, TBUT = tear break-up time.

**Table 4.** Comparison of the amount of changes in visual function and ocular fatigue between 5 and 10 lux light exposure at night

Parameters	5 lux illumination		10 lux illumination		P value
	Mean	SD	Mean	SD	
Objective parameters					
Conjunctival hyperemia					
Nasal	4.23	7.03	3.93	7.86	0.970
Temporal	2.31	9.08	3.57	8.70	0.711
Tear break-up time	−0.35 <sup>a</sup>	1.09	−0.45 <sup>a</sup>	1.21	0.912
Maximal blinking interval	−2.60 <sup>a</sup>	7.32	−1.36 <sup>a</sup>	2.97	0.910
Ocular surface temperature	−0.06 <sup>a</sup>	0.56	−0.08 <sup>a</sup>	0.47	0.471
Subjective parameters					
Tired eyes	0.46	1.39	0.64	1.28	0.485
Sore/aching eyes	0.23	1.14	0.32	0.72	0.706
Irritated eyes	0.35	1.09	0.18	1.56	0.593
Watery eyes	−0.04 <sup>a</sup>	1.61	0.04	1.67	0.993
Dry eyes	0.23	1.39	0.00	1.44	0.789
Eyestrain	−0.04 <sup>a</sup>	1.11	0.14	0.71	0.642
Hot/burning eyes	0.15	0.54	−0.04 <sup>a</sup>	0.79	0.356
Blurred vision	−0.19 <sup>a</sup>	1.13	0.07	1.21	0.160
Difficulty focusing	0.12	0.82	0.32	0.72	0.660
Vision discomfort	−0.04 <sup>a</sup>	0.77	0.25	0.44	0.132

SD = standard deviation, TBUT = tear break-up time.

<sup>a</sup>Decrease after sleep under ambient light exposure.

potent antioxidants, and it has known neuroprotective, anti-aging, immunomodulative, and oncogenic properties.<sup>3</sup> For example, an elevated breast cancer prevalence in night shift-work nurses is linked to disrupted melatonin secretion.<sup>18-20</sup> Symptoms frequently reported by shift-workers such as insomnia, depression, cardiovascular disease, and premature mortality are also thought to be related to melatonin secretion disruption, which is caused by nocturnal light exposure.<sup>21</sup>

The human eye is also influenced by circadian rhythm. Numerous studies report diurnal variation in ocular structures and intraocular pressure. Intraocular pressure in a normal

population is reported to vary 4–5 mm Hg diurnally, and the variation is greater in patients diagnosed with glaucoma.<sup>22,23</sup> Diurnal variation in choroidal thickness and axial length is also reported.<sup>24,25</sup> The circadian structural alterations are thought to control ocular growth, influence emmetropization, and affect refractive error development.<sup>26,27</sup> When the eye is exposed to nocturnal light, the diurnal ocular variation may be altered either by response to the light itself or indirectly through the disrupted circadian rhythm. In animal experiments, disruption of normal diurnal light rhythm changes refractive development in chickens.<sup>4,28</sup> Quinn et al.<sup>5</sup> noted that children exposed to nocturnal ambient light have a higher prevalence of myopia, which suggests that nocturnal light exposure during sleep may be a precipitating factor in myopia development. Reports contradict on the relationship between nocturnal light exposure and myopia development<sup>6</sup> and are inconclusive because of the absence of prospective case-controlled studies, but a potential correlation between nocturnal ambient light and ocular growth cannot be excluded.

This study found that 5 lux or greater light exposure during sleep at night increased ocular fatigue. In most studies, ocular fatigue is evaluated only subjectively using a symptom questionnaire, and an objective measurement of ocular fatigue has not been established. In our previous study,<sup>12</sup> we measured ocular fatigue after 1 hour of computer work both subjectively and objectively, which revealed that TBUT, maximal blinking interval, conjunctival hyperemia, and ocular surface temperature correlated with ocular fatigue. Therefore, we used identical objective and subjective methodology in this study.

The questionnaire revealed that eye tiredness, soreness, focusing difficulty and vision discomfort increased in the group with 10 lux light exposure at night. Conjunctival hyperemia also increased, and TBUT and maximal blinking interval decreased. In most previous studies, ocular fatigue was evaluated after visual tasks,<sup>29–31</sup> and, these studies revealed that the blinking rate decreased, the ocular surface was more exposed to air during visual tasks, and the resulting dry eye caused ocular fatigue symptoms.<sup>29,32–34</sup> However, the subjects in our study did not perform any visual tasks; instead, the eyes were closed during sleep. Thus, an explanation other than decreased blinking rate is necessary. There are several potential mechanisms of ocular fatigue induced by nocturnal ambient light. Light may partially enter the eyes through the eyelid, which would induce the light reflex, prolong pupillary constriction, and cause ocular fatigue. In addition, disruption of the circadian rhythm may be a factor in resulting ocular fatigue. The nocturnal light exposure may disrupt the circadian clock and disturb the diurnal variation of ocular structures including choroidal blood flow, resulting in ocular fatigue. Alternatively, the ambient light may deteriorate sleep quality and cause shallow sleep.<sup>35,36</sup> This sleep disturbance can induce systemic fatigue<sup>37</sup> and subsequently ocular fatigue. Sleep disturbance has been also documented to induce dry eye syndrome. Lee et al.<sup>38</sup> reported that sleep deprivation reduced tear secretion, impaired tear film and decreased TBUT. They hypothesized that the alteration in autonomic nervous tone and imbalance of hormone levels induced by sleep deprivation affected tear secretion and tear film stability. Decreased TBUT after nocturnal ambient light exposure in our study could be explained in this aspect. We cannot determine the mechanism of ocular fatigue in the current study, but we surmise that these mechanisms may induce ocular fatigue in varying degrees. Further research of the mechanisms is necessary.

There are several limitations in this study. The number of subjects was small. Although ocular fatigue was induced after light exposure despite the small number of subjects, we think that future study using a larger sample size would reveal additional and more reliable differences



associated with nocturnal ambient light. In addition, daytime activity of the subjects was not controlled. We instructed subjects to avoid activities that may influence ocular fatigue or sleep quality, including vigorous exercise, napping, excessive caffeine intake, and alcohol consumption, and the subjects reporting these activities were excluded. However, daytime activity could not be monitored; therefore, we cannot assume that those activities were perfectly controlled. Further study that systematically controls daytime activity on a larger number of subjects is necessary.

In conclusion, nocturnal light exposure increases ocular fatigue. Ambient light during nocturnal sleep should be prevented to avoid ocular fatigue.

## REFERENCES

1. Stevens RG, Blask DE, Brainard GC, Hansen J, Lockley SW, Provencio I, et al. Meeting report: the role of environmental lighting and circadian disruption in cancer and other diseases. *Environ Health Perspect* 2007;115(9):1357-62.  
[PUBMED](#) | [CROSSREF](#)
2. Fonken LK, Workman JL, Walton JC, Weil ZM, Morris JS, Haim A, et al. Light at night increases body mass by shifting the time of food intake. *Proc Natl Acad Sci U S A* 2010;107(43):18664-9.  
[PUBMED](#) | [CROSSREF](#)
3. Turner PL, Van Someren EJ, Mainster MA. The role of environmental light in sleep and health: effects of ocular aging and cataract surgery. *Sleep Med Rev* 2010;14(4):269-80.  
[PUBMED](#) | [CROSSREF](#)
4. Weiss S, Schaeffel F. Diurnal growth rhythms in the chicken eye: relation to myopia development and retinal dopamine levels. *J Comp Physiol A Neuroethol Sens Neural Behav Physiol* 1993;172(3):263-70.  
[PUBMED](#) | [CROSSREF](#)
5. Quinn GE, Shin CH, Maguire MG, Stone RA. Myopia and ambient lighting at night. *Nature* 1999;399(6732):113-4.  
[PUBMED](#) | [CROSSREF](#)
6. Saw SM, Wu HM, Hong CY, Chua WH, Chia KS, Tan D. Myopia and night lighting in children in Singapore. *Br J Ophthalmol* 2001;85(5):527-8.  
[PUBMED](#) | [CROSSREF](#)
7. Grisham JD, Sheppard MM, Tran WU. Visual symptoms and reading performance. *Optom Vis Sci* 1993;70(5):384-91.  
[PUBMED](#) | [CROSSREF](#)
8. Cohen Y, Segal O, Barkana Y, Lederman R, Zadok D, Pras E, et al. Correlation between asthenopic symptoms and different measurements of convergence and reading comprehension and saccadic fixation eye movements. *Optometry* 2010;81(1):28-34.  
[PUBMED](#) | [CROSSREF](#)
9. Agnew HW Jr, Webb WB, Williams RL. The first night effect: an EEG study of sleep. *Psychophysiology* 1966;2(3):263-6.  
[PUBMED](#) | [CROSSREF](#)
10. Schulze MM, Jones DA, Simpson TL. The development of validated bulbar redness grading scales. *Optom Vis Sci* 2007;84(10):976-83.  
[PUBMED](#) | [CROSSREF](#)
11. Ames SL, Wolffsohn JS, McBrien NA. The development of a symptom questionnaire for assessing virtual reality viewing using a head-mounted display. *Optom Vis Sci* 2005;82(3):168-76.  
[PUBMED](#) | [CROSSREF](#)
12. Suh YW, Kim KH, Kang SY, Kim SW, Oh JR, Kim HM, et al. The objective methods to evaluate ocular fatigue associated with computer work. *J Korean Ophthalmol Soc* 2010;51(10):1327-32.  
[CROSSREF](#)
13. Hannibal J, Fahrenkrug J. Neuronal input pathways to the brain's biological clock and their functional significance. *Adv Anat Embryol Cell Biol* 2006;182:1-71.  
[PUBMED](#)

14. Gachon F, Nagoshi E, Brown SA, Ripperger J, Schibler U. The mammalian circadian timing system: from gene expression to physiology. *Chromosoma* 2004;113(3):103-12.  
[PUBMED](#) | [CROSSREF](#)
15. Czeisler CA, Klerman EB. Circadian and sleep-dependent regulation of hormone release in humans. *Recent Prog Horm Res* 1999;54:97-130.  
[PUBMED](#)
16. Pandi-Perumal SR, Srinivasan V, Maestroni GJ, Cardinali DP, Poeggeler B, Hardeland R. Melatonin: nature's most versatile biological signal? *FEBS J* 2006;273(13):2813-38.  
[PUBMED](#) | [CROSSREF](#)
17. Skene DJ, Lockley SW, Thapan K, Arendt J. Effects of light on human circadian rhythms. *Reprod Nutr Dev* 1999;39(3):295-304.  
[PUBMED](#) | [CROSSREF](#)
18. Schernhammer ES, Hankinson SE. Urinary melatonin levels and breast cancer risk. *J Natl Cancer Inst* 2005;97(14):1084-7.  
[PUBMED](#) | [CROSSREF](#)
19. Figueiro MG, Rea MS, Bullough JD. Does architectural lighting contribute to breast cancer? *J Carcinog* 2006;5(1):20.  
[PUBMED](#) | [CROSSREF](#)
20. Erren TC, Pape HG, Reiter RJ, Piekarski C. Chronodisruption and cancer. *Naturwissenschaften* 2008;95(5):367-82.  
[PUBMED](#) | [CROSSREF](#)
21. Knutsson A. Health disorders of shift workers. *Occup Med (Lond)* 2003;53(2):103-8.  
[PUBMED](#) | [CROSSREF](#)
22. Liu JH, Kripke DF, Twa MD, Hoffman RE, Mansberger SL, Rex KM, et al. Twenty-four-hour pattern of intraocular pressure in the aging population. *Invest Ophthalmol Vis Sci* 1999;40(12):2912-7.  
[PUBMED](#)
23. Liu JH, Zhang X, Kripke DF, Weinreb RN. Twenty-four-hour intraocular pressure pattern associated with early glaucomatous changes. *Invest Ophthalmol Vis Sci* 2003;44(4):1586-90.  
[PUBMED](#) | [CROSSREF](#)
24. Tan CS, Ouyang Y, Ruiz H, Sadda SR. Diurnal variation of choroidal thickness in normal, healthy subjects measured by spectral domain optical coherence tomography. *Invest Ophthalmol Vis Sci* 2012;53(1):261-6.  
[PUBMED](#) | [CROSSREF](#)
25. Chakraborty R, Read SA, Collins MJ. Diurnal variations in axial length, choroidal thickness, intraocular pressure, and ocular biometrics. *Invest Ophthalmol Vis Sci* 2011;52(8):5121-9.  
[PUBMED](#) | [CROSSREF](#)
26. Nickla DL. Ocular diurnal rhythms and eye growth regulation: where we are 50 years after Lauber. *Exp Eye Res* 2013;114:25-34.  
[PUBMED](#) | [CROSSREF](#)
27. Stone RA, Pardue MT, Iuvone PM, Khurana TS. Pharmacology of myopia and potential role for intrinsic retinal circadian rhythms. *Exp Eye Res* 2013;114:35-47.  
[PUBMED](#) | [CROSSREF](#)
28. Cohen Y, Peleg E, Belkin M, Polat U, Solomon AS. Ambient illuminance, retinal dopamine release and refractive development in chicks. *Exp Eye Res* 2012;103:33-40.  
[PUBMED](#) | [CROSSREF](#)
29. Himebaugh NL, Begley CG, Bradley A, Wilkinson JA. Blinking and tear break-up during four visual tasks. *Optom Vis Sci* 2009;86(2):E106-14.  
[PUBMED](#) | [CROSSREF](#)
30. Cardona G, García C, Serés C, Vilaseca M, Gispets J. Blink rate, blink amplitude, and tear film integrity during dynamic visual display terminal tasks. *Curr Eye Res* 2011;36(3):190-7.  
[PUBMED](#) | [CROSSREF](#)
31. Kim DJ, Lim CY, Gu N, Park CY. Visual fatigue induced by viewing a tablet computer with a high-resolution display. *Korean J Ophthalmol* 2017;31(5):388-93.  
[PUBMED](#) | [CROSSREF](#)
32. Ousler GW 3rd, Hagberg KW, Schindelar M, Welch D, Abelson MB. The ocular protection index. *Cornea* 2008;27(5):509-13.  
[PUBMED](#) | [CROSSREF](#)
33. Toda I, Fujishima H, Tsubota K. Ocular fatigue is the major symptom of dry eye. *Acta Ophthalmol (Copenh)* 1993;71(3):347-52.  
[PUBMED](#) | [CROSSREF](#)



34. Kang BS, Seo MW, Yang HK, Seo JM, Lee S, Hwang JM. Comparison of blinking patterns when watching ultra-high definition television: normal versus dry eyes. *J Korean Ophthalmol Soc* 2017;58(6):706-11.  
[CROSSREF](#)
35. Cho JR, Joo EY, Koo DL, Hong SB. Let there be no light: the effect of bedside light on sleep quality and background electroencephalographic rhythms. *Sleep Med* 2013;14(12):1422-5.  
[PUBMED](#) | [CROSSREF](#)
36. Cho CH, Lee HJ, Yoon HK, Kang SG, Bok KN, Jung KY, et al. Exposure to dim artificial light at night increases REM sleep and awakenings in humans. *Chronobiol Int* 2016;33(1):117-23.  
[PUBMED](#) | [CROSSREF](#)
37. Thomas KS, Motivala S, Olmstead R, Irwin MR. Sleep depth and fatigue: role of cellular inflammatory activation. *Brain Behav Immun* 2011;25(1):53-8.  
[PUBMED](#) | [CROSSREF](#)
38. Lee YB, Koh JW, Hyon JY, Wee WR, Kim JJ, Shin YJ. Sleep deprivation reduces tear secretion and impairs the tear film. *Invest Ophthalmol Vis Sci* 2014;55(6):3525-31.  
[PUBMED](#) | [CROSSREF](#)