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CLINICAL TRIAL REPORT

Pilot study to examine the effects of indoor daylight exposure on depression and other neuropsychiatric symptoms in people living with dementia in long-term care communities

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Keywords: dementia, daylight, depression, memory care communities

Introduction

Depression is one of the most common conditions affecting people living with Alzheimer's disease and related dementias (ADRD). Clinically significant depression occurs in about 20%–30% of people with ADRD.¹ Treatment of depression in people living with ADRD can improve well-being, quality of life, and individual function, even in the presence of ongoing declines in memory and cognition. Exposure to sufficient daylight is a potentially effective non-drug treatment option for depression and other neuropsychiatric symptoms of dementia. Exposure to bright white light at the cornea (2,000–10,000 lux) typically administered in the morning has been shown to be an effective nonpharmacological treatment option for depressed adults with normal cognitive function.² Light therapy has also been shown to be effective in treating sleep disturbances³ and for ameliorating behavioral problems⁴ in people living with ADRD. Riemersma-van der Lek et al⁵ showed that exposure throughout the day to bright white light from large numbers of ceiling-mounted fluorescent lamps (>1,000 lux at the cornea) could increase sleep quality and reduce cognitive decline in people living with ADRD.

However, reliable and well-designed studies examining the efficacy of light therapy in treating depression in people living with ADRD are extremely limited.⁶ Moreover,

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the conventional approach to administering the bright light stimulus, through light boxes containing arrays of bright light fixtures, is problematic for people with ADRD due to reported side effects of headache, eye strain, nausea, and agitation.^{7,8} Furthermore, many light therapy studies were performed prior to scientific understanding of the important role that specific wavelengths of light play in maintaining healthy human biological functions. Research has revealed that the human circadian system is maximally sensitive to short-wavelength (blue) light.^{9,10} This has led to more recent light therapy studies using "blue-enriched" electrical light sources tailored to the spectral response of the circadian system.^{11,12} The circadian system is the body's internal biological clock that regulates most 24-hour behavioral and physiological rhythms, such as sleep/wake cycle, alertness level, hormone suppression/secretion, and core body temperature. A bright light stimulus in the morning is the most powerful timing cue to maintain entrainment with the astronomical 24-hour day.

In institutionalized settings, where light levels can be very low,^{13,14} lack of sufficient exposure to bright light is considered one of the primary contributors to circadian disruption. Circadian disruption, in turn, is associated with depression, sleep disruption, agitated behavior, and cognitive decline.^{15,16} In the present study, we hypothesized that exposure to sufficient daylight indoors can serve as an effective nonpharmacological treatment for reducing depression and other neuropsychiatric symptoms in people living with ADRD. Furthermore, it has the potential for fewer adverse effects and has less practical limitations than specialized electrical lighting devices.

Methods

The pilot study was a nonrandomized clustered trial, where communities were assigned to the daylight intervention or usual care control; all eligible and consenting participants within a community received the same intervention. All study procedures were approved by the University of Southern California (USC) Institutional Review Board (USC UPIRB #UP-16-00487). Participation was voluntary, and participants' legally authorized representatives provided written informed consent and Health Insurance Portability and Accountability Act authorization. <u>ClinicalTrials.gov</u> Identifier: NCT03483896.

Protocol

The research was conducted at eight Silverado Senior Living dementia care communities in Los Angeles and Orange counties using a two-arm (daylight intervention and control) parallel intervention study design. Silverado Senior Living is a memory care/assisted living provider that delivers care for persons with Alzheimer's and other memory-impairing diseases. The study was run over a period of 12 weeks at each community, with staggered start dates ranging from 1 to 3 days per community to enable the research team to visit each site in series to collect baseline and outcome measures.

Participants

Residents were recruited according to the trial inclusion criteria: 1) ADRD diagnosis, 2) no physical comorbidities that precluded participation in the daily group intervention, and 3) a Mini-Mental State Exam score of ≥ 10 . A total of 83 residents who met these criteria were enrolled into the study. Of the 83 participants enrolled, one declined when asked for verbal assent, three passed away during the study, and two were unable to be scored at outcome for the Neuropsychiatric Inventory Nursing Home Version (NPI-NH) (due to absence of regular caregiver at the time of NPI-NH scoring visits).

Intervention group

At the four communities in the active light intervention arm, staff increased the daylight exposure of participants by taking them to the perimeter zone of a daylit room from 8:00 to 10:00 AM for socialization over a period of 12 weeks. The perimeter zone was defined to be the region of the room within 3 m from the windows. The intervention was administered each day (7 days/week) over the duration of the study. Randomization at the community level was not feasible in this pilot study because not all of the eight participating communities had a suitably sized perimeter zone space available for use during the study timeframe. Therefore, the four communities assigned to the intervention were those where a suitably sized perimeter zone space was available for use during the intervention period (8:00-10:00 AM). As all communities were operated by Silverado Senior Living, all had similar interior design and furnishing, caregiver staff training, and programmatic activities.

Control group

Participants at the other four communities received the usual care (control arm). During the period from 8:00–10:00 AM each day, the control group were taken to a similar sized area indoors (without daylight) for socialization under typical electrical lighting conditions.

Outcome measures

The Cornell Scale for Depression in Dementia (CSDD) was used to measure depression. The CSDD is designed for the assessment of depression in older people with dementia who

can at least communicate basic needs. Scores >10 indicate a probable major depression. Scores >18 indicate a definite major depression. Scores < 6 are associated with the absence of significant depressive symptoms.¹⁷ The care provider uses the CSDD to evaluate the mental state of higher functioning residents at regular intervals. Consequently, CSDD scores were obtained from the participants' health records. The most recent CSDD score prior to the beginning of the study was used as the baseline score wherever feasible. In some cases, no CSDD was found, or the date of completion was more than 4 months prior to the start of the study. These participants were excluded from the analyses for CSDD, leading to a smaller sample size (n=36 daylight, n=28 control). Due to a delay in the scheduled start of the study, CSDD scores were completed by the care provider on an average of 7.5 weeks prior to the start of the study for the intervention group and 5.9 weeks prior to the start for the control group. However, the group difference in mean weeks prior to start of the study was not statistically significant (p=0.22). The CSDD at outcome was completed, on average, within 3 days of the end of the study.

The NPI-NH is designed for the assessment of people with dementia residing in extended care housing communities.¹⁸ The NPI-NH includes 10 behavioral areas (delusions, hallucinations, agitation/aggression, depression/dysphoria, anxiety, elation/euphoria, apathy/indifference, disinhibition, irritability/lability, and aberrant motor behavior) and two types of neurovegetative changes (sleep and nighttime behavior disorders, and appetite and eating disorders). A summary score is obtained by summing all 12 domain scores, where higher scores indicate greater neuropsychiatric symptomatology.

Light measurements

Lighting measurements were taken periodically during the study to evaluate the fidelity of the daylight intervention. Due to the difference in spectral sensitivity between the visual system and the circadian system, conventional photometric sensors are problematic for assessing human circadian lighting needs.¹⁹ To address this challenge, a digital charge coupled device spectrometer mounted on a mobile cart was used to evaluate the spectral composition of light by taking spot measurements of Spectral Power Distribution (SPD). The spectrometer lens was positioned on a vertical plane at a distance of 1.07 m (42 in.) from the floor to represent views from seated eye height. Use of a calibrated spectrometer for measurement of light exposures follows the consensus research recommendation to record light exposures in the most complete form, that is corneal SPDs.¹⁹ In each

intervention space, SPD measurements were taken during the intervention at distances of 1, 3, and 5 m from windows and in four directions at each distance, beginning with a window-facing view and rotating in the clockwise direction by 90° increments. Each daylight intervention space was measured on five separate visits during the study, with visits spaced 2–3 weeks apart. Due to the uniform and steady-state lighting conditions of the control spaces, each control space was measured once; measurements were taken in four directions at a single location representative of typical eye-level lighting conditions in the space.

Analyses

Light measurements

Each SPD was analyzed to calculate melanopic illuminance (mLux) by applying the melanopic spectral efficiency function to the measured spectral irradiance following the methodology outlined in Lucas et al¹⁹ and Enezi et al.²⁰ To calculate a proxy indicator of the overall light exposure in each daylight intervention space, the average of all measurements taken in each space under clear sky conditions at the 3 m distance in all four view directions was used. For the control, the average of all four view directions in each space was used. This proxy indicator is referred to as "mLux_{AVG}" in the following sections.

Outcome measures

Mean values of the trial variables, CSDD and NPI-NH, were analyzed by treatment group and by community at baseline and at the end of study. The change in each variable (end – baseline) was examined. Between- and within-group comparisons (daylight vs usual care) in CSDD and NPI-NH scores were analyzed by linear mixed effects models to reflect the clustered (ie, community level) assignment to interventions. A random effect at the community level was specified to model correlated outcomes among participants within communities. Intraclass (within community) correlations in the change in CSDD and NPI-NH were calculated using estimates of variability (mean squares) within and between communities from analysis of variance. The linear association of mLux-AVG measurements with changes in trial outcome measures was also analyzed with linear mixed effects models as above; rather than intervention group, the primary independent variable was each participant's mLux_{AVG} measurement.

Results

While mean values of the CSDD and NPI-NH trial outcomes varied markedly between communities (Table 1), most of

Table I Comparisons by community

	I	2	3	4	5	6	7	8	p-value
N	18	11	18	9	7	6	5	3	
Treatment	Daylight	Control	Daylight	Control	Daylight	Control	Control	Daylight	
mLux	230*	20	100	56	75	57	49	120	
Baseline									
NPI-NH	18.4 (0, 96)	22.7 (2, 54)	11.5 (0, 66)	11.6 (0, 46)	20.6 (3, 43)	9.7 (1, 29)	18.6 (9, 32)	18.3 (0, 31)	0.64
CSDD	9.4 (0, 20)	8.7 (2, 16)	4.1 (0, 17)	0.9 (0, 2)	1.6 (0, 4)	1.3 (0, 3)	4.8 (0, 8)	1.7 (0, 4)	0.0003
End									
NPI-NH	20.3 (0, 93)	26.8 (5, 62)	10.7 (0, 39)	13.6 (0, 44)	6.4 (0, 18)	6.3 (0, 22)	29.2 (2, 59)	14.7 (0, 30)	0.08
CSDD	4.8 (0, 17)	10.9 (2, 24)	1.9 (0, 7)	1.0 (0, 2)	1.8 (0, 4)	1.8 (0, 5)	6.4 (0, 11)	1.3 (0, 4)	0.0001
Change									
NPI-NH	I.9 (–47, 42)	4.I (-9, 36)	-0.8 (-27, 20)	2.0 (-5, 16)	-14.1 (-25, -3)	-3.3 (-28, 13)	10.6 (-7, 41)	-3.7 (-17, 6)	0.13
CSDD	-4.6 (-15, 9)	2.2 (-8, 15)	-2.6 (-10, 3)	0.1 (-2, 2)	0.2 (0, 1)	0.2 (0, 1)	1.6 (0, 3)	-0.3 (-1, 0)	0.1

Notes: Numbers in table are mean (minimum, maximum); *p*-value from analysis of variance; participants with baseline CSDD >4 months before enrollment excluded; *median value.

Abbreviations: mLux_{AVG}, average melanopic illuminance at 3 m from windows; NPI-NH, Neuropsychiatric Inventory Nursing Home Version; CSDD, Cornell Scale for Depression in Dementia.

these community means (baseline, endpoint, and change) did not significantly differ between communities. However, significant differences between communities were observed in the CSDD mean values at baseline (p=0.0003) and endpoint (p=0.0001). Mean NPI-NH measures did not significantly differ across communities. Numbers of participants at communities ranged from 3 to 18. The mLux_{AVG} measures at each community reflected the daylight versus control interventions (Table 1).

Eligible and enrolled participants who completed the study included 46 participants at daylight intervention communities and 31 participants at control intervention communities (Table 2). Baseline demographic variables did not differ between the groups. Overall, the sample had a mean age of 85.3 years with a SD of 7.0 years. Subjects had resided in their communities for an average of 1.0 (1.1) years. The great majority of the sample was white (92.2%), female (72.7%), with diagnoses of either Alzheimer's disease (32.5%) or dementia not otherwise specified (41.7%). The baseline NPI-NH (n=77) and CSDD (n=66) means did not differ significantly between the control and intervention groups (Table 2).

The mLux_{AVG} measures at daylight communities were significantly higher than in control communities (p < 0.0001, Table 2) confirming the effectiveness of the daylight intervention. Changes in NPI-NH and CSDD varied substantially between communities (Figures 1 and 2). Participants in the daylight intervention showed an average decrease over the trial in both NPI-NH and CSDD, while the control participants showed an average increase in both NPI-NH and CSDD (Table 2). The group differences in outcome changes achieved statistical significance for CSDD (p=0.01) but not for NPI-NH (p=0.17). Group differences in the CSDD change were also evident among nine participants with baseline CSDD >10 (Table 3, p=0.03 between groups). The mLux_{AVG} measures were significantly inversely correlated with the change in CSDD over the trial (Spearman r=-0.37, p=0.002), but not with change in NPI-NH (Table 3, Figure 3).

Table 2 Baseline and endpoint comparisons by treatment

	Daylight	Control	p-value
			between
			treatments
N	46	31	
Female	31 (67.4%)	25 (80.6%)	0.30
Age (years)	85.6 (1.0)	84.7 (1.3)	0.56
White race	42 (91.3%)	29 (93.5%)	1.00
Clinical group			0.69
AD	17 (37.0%)	8 (25.8%)	
Frontotemporal dementia	I (2.2%)	I (3.2%)	
Lewy body dementia	0	I (3.2%)	
Vascular dementia	2 (4.3%)	I (3.2%)	
Dementia NOS	19 (41.3%)	17 (54.8%)	
Mild cognitive impairment	6 (13.0%)	2 (6.5%)	
Not specified	I (2.2%)	I (3.2%)	
Years in community	1.2 (0.2)	0.6 (0.3)	0.14
Baseline outcomes			
NPI-NH	16.2 (3.2)	16.1 (2.7)	0.97
CSDD	4.2 (1.9)	3.9 (1.9)	0.93
Endpoint outcomes			
NPI-NH	13.4 (4.2)	19.1 (4.4)	0.35
CSDD	2.7 (2.1)	5.3 (2.0)	0.39
Change (endpoint minus base	line) outcomes		
NPI-NH	-2.8 (2.9)	3.1 (3.2)	0.17
p-value within group	0.33	0.33	
CSDD	-2.0 (0.9)	1.5 (1.0)	0.01
p-value within group	0.025	0.13	
mLux _{AVG}	159.3 (13.8)	42.3 (3.1)	<0.0001

Notes: Numbers in table are mean (SEM); for age, years in community, and trial outcome measures, *p*-values between treatments are by mixed effects linear models, specifying a random effect for community to account for within-community clustering. Sample size for years in community (n=46 daylight, n=24 control); sample size for CSDD: (n=36 daylight, n=28 control), intraclass correlation (ICC) for within-community change in outcomes: ICC =0.065 for change in NPI-NH; ICC =0.03 for change in CSDD.

Abbreviations: AD, Alzheimer's disease; mLux_{AVG}, average melanopic illuminance at 3 m from windows; NOS, not otherwise specified; NPI-NH, Neuropsychiatric Inventory Nursing Home Version; CSDD, Cornell Scale for Depression in Dementia; SEM, standard error of the mean.



Figure I CSDD change by community. Note: Intraclass correlation of Cornell change (within-community correlation) =0.03. Abbreviation: CSDD, Cornell Scale for Depression in Dementia.



100

Figure 3 Scatter plot of changes in CSDD.

20

10

0

-20

0

CSDD change

Abbreviations: CSDD, Cornell Scale for Depression in Dementia; mLux_{AVG} average melanopic illuminance at 3 m from windows.

200

mLux_{AVG}

300

Predicted CSDD change

400

Discussion

Few, if any, studies of indoor daylight exposure have been conducted in dementia care communities. Our results indicate that an intervention that increases exposure to daylight in the morning (8:00–10:00 AM) may be used to reduce depression in people living with dementia. Overall, the mean mLux_{AVG} level measured in the daylight intervention spaces (159.3) was 3.8 times greater than the mean value for the control spaces (42.3) (Table 2). The mLux_{AVG} measures were found to be significantly inversely correlated with the change in CSDD over the trial (Spearman r=–0.37, p=0.002), suggesting that greater reduction in depression symptoms (Table 3). This finding has the potential to be clinically significant because it can inform the daily programming of dementia care communities, for example, by prioritizing the use of existing daylit spaces



Figure 2 NPI-NH change by community. Note: Intraclass correlation of NPI-NH change (within-community correlation) =0.065. Abbreviation: NPI-NH, Neuropsychiatric Inventory Nursing Home Version.

during the morning (ie, 8:00–10:00 AM) as a nonpharmacological treatment alternative for people living with dementia and depression.

Among the subset of participants with a probable major depression (CSDD >10) at the beginning of the trial (n=5 daylight, n=4 control), the daylight intervention was found to reduce the average CSDD score by over 11 points, from 16.8 at baseline to 5.6 at the end of the trial, which is below the threshold score of six used to indicate the absence of significant depressive symptoms (Table 4). In comparison, the average CSDD score for the control group with CSDD >10 increased slightly (from 12.7 to 13.5, Table 4). While this outcome should be interpreted with caution due to the small number of study participants (n=9 in the analysis), it suggests that increased light exposure has the potential to be an effective treatment for residents with probable major depression, producing a reduction in symptoms similar to the reduction that would be expected with an effective pharmacological treatment.

There are currently no minimum requirements for light exposure in dementia care environments to ensure the effectiveness of the indoor environment as a nonpharmacological treatment option. Recommended practices such as Lighting

	Table 3	Correlations	among	mLux	and	change	in	outcomes
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	Spearman	Beta coefficient
	Correlation	(SE) [p-value]
	(p-value)	from mixed model
NPI-NH	0.002 (0.99)	0.012 (0.026) [0.64]
CSDD	-0.37 (0.002)	-0.016 (0.01) [0.054]

Notes: Sample size: n=77 in NPI-NH analysis; n=64 in CSDD analysis. **Abbreviations:** NPI-NH, Neuropsychiatric Inventory Nursing Home Version; CSDD, Cornell Scale for Depression in Dementia; mLuxAVG, average melanopic illuminance at 3 m from windows; SE, standard error. Table 4Treatment group comparisons on behavioral anddepression change: baseline Cornell >10

	Daylight	Control	p-value between treatment
NPI-NH			
Baseline	14.3 (12.0)	34.7 (15.5)	0.34
End of study	25.4 (9.3)	39.0 (10.4)	0.37
Change	10.6 (15.7)	4.3 (19.7)	0.81
CSDD			
Baseline	16.8 (1.3)	12.7 (1.5)	0.09
End of study	5.6 (2.6)	13.5 (2.9)	0.09
Change	-11.2 (2.7)	0.7 (3.1)	0.03

Notes: p-values from linear mixed effects models, with random effect for community. Numbers in table are model-estimated mean (SE) change in trial outcomes. N=9 in analysis (n=5 daylight, n=4 control).

Abbreviations: NPI-NH, Neuropsychiatric Inventory Nursing Home Version; CSDD, Cornell Scale for Depression in Dementia; SE, standard error.

and the Visual Environment for Seniors and the Low Vision Population²¹ and Lighting for Hospitals and Healthcare Communities²² have focused primarily on factors of safety and visual performance and have only recently begun to incorporate language addressing non-visual needs for light; yet, there remain no specific, measurable lighting requirements. Findings from this pilot study represent a first step toward the development of evidence-based lighting guidance to support the design and operation of dementia care environments that have the potential to reduce the need for pharmacological treatments for depression and other health conditions.

The lighting measurements recorded in the daylight intervention spaces should not be interpreted as a definitive "prescription" for effective indoor light stimulus. While our reported findings may be due to chance in light of multiple hypothesis testing in this pilot trial, the consistency of the mean changes in daylight versus control participants across the communities (Table 1) provides some assurance that these findings are real and estimates can be used for the design of a large informative clinical trial. Further studies are needed to build a body of evidence for the appropriate timing, duration, wavelength, and intensity of light exposure for adults with dementia, as well as to determine how patterns of light and dark should be orchestrated over 24-hour periods to suit the biological needs of each individual.

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Disclosure

The authors report no conflicts of interest in this work.

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