An Open Trial of Morning Light Therapy for Treatment of Antepartum Depression

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Objective: About 5% of pregnant women meet criteria for major depression. No pharmacotherapy is specifically approved for antepartum depression; novel treatment approaches may be welcome. The authors explored the use of morning bright light therapy for antepartum depression.

Method: An open trial of bright light therapy in an A-B-A design was conducted for 3–5 weeks in 16 pregnant patients with major depression. The Hamilton Depression Rating Scale, Seasonal Affective Disorders Version, was administered to assess changes in mood. A follow-up questionnaire was used to assess outcome after delivery.

Results: After 3 weeks of treatment, mean depression ratings improved by 49%. Benefits were seen through 5 weeks of treatment. There was no evidence of adverse effects of light therapy on pregnancy.

Conclusions: These data provide evidence that morning light therapy has an antidepressant effect during pregnancy. A randomized controlled trial is warranted to test this alternative to medication.

About 5% of pregnant women meet the Research Diagnostic Criteria for major depression (1), a suboptimal physiologic milieu for pregnancy. Antepartum depression may be a risk factor for development of preeclampsia (2) and is the strongest predictor of postpartum depression (3). Maternal depressive symptoms during pregnancy may lead to neonatal developmental changes and early childhood irritability (4, 5). Further, an altered intrauterine climate may affect the newborn's neurobehavioral function. Finally, infants of depressed mothers demonstrate lower basal and stimulated frontal EEG activity (6).

Pilot studies have supported the potential value of interpersonal psychotherapy as a form of treatment for antepartum depression (7) and explored the potential value of sleep deprivation or light therapy (8). The safety of pharmacologic treatment of depression in pregnant women is controversial and just beginning to be the subject of controlled investigations (9, 10). Treatment of depressed pregnant women with antidepressants requires skilled decision making based on limited data (11). Exploration of alternative treatments for antepartum depression is therefore critically important.

Daily bright light exposure, alone or combined with antidepressant medications, has been shown in several small studies (12, 13) to be effective in treating patients with nonseasonal major depression, and there are case reports of its potential value in treating postpartum depression (14). Further studies of light therapy in combination with medication are required, however, to ascertain possible interactions and safety (15). Pregnant women have been excluded from past studies of light therapy, however, to simplify study designs and avoid an uncharted dimension of risk assessment. To our knowledge, this study is the first to focus light treatment on this vulnerable population.

Method

Pregnant patients were recruited through the media and referrals from obstetricians. After complete description of the study to the patients, written informed consent was obtained. A diagnosis of major depressive disorder was established according to DSM-IV criteria (16), and a Structured Interview Guide for the Hamilton Depression Rating Scale, Seasonal Affective Disorders Version (SIGH-SAD) (17) score of at least 20 was required. Patients with the following characteristics were excluded from entry: current treatment with psychotropic medication; recent history of suicide attempt; panic or eating disorder unrelated to depression; other axis I diagnosis; current medical illness (including epilepsy, retinal pathology, cataract, or glaucoma); thyroid function test results inconsistent with normal pregnancy; past head injury with neurological or psychiatric sequelae; current use of β-adrenergic blockers, melatonin, or St. John's wort; delayed-sleep-phase syndrome or hypersomnia with habitual sleep onset later than 1:00 a.m. and/or awakening later than 10:00 a.m.; and current use of alcohol or drugs (based on self-report).

The patients were followed for up to 2 weeks (observation period) before beginning treatment at a distance of 33 cm with an ultraviolet-screened diffuse white fluorescent light source (a Day-Light 10,000-lux box; Day-Light Technologies, Inc., Halifax, N.S., Canada) tilted downward toward the head of the seated patients. The patients were instructed to use the light therapy at home for 60 minutes daily, beginning within 10 minutes of awakening for at least 3 weeks. Compliance was documented by having the subjects call an answering machine daily to report their light use. A morning light time was chosen because of preponderant evidence that this interval elicits advances in the circadian rhythm phase that have been highly correlated with the effective treatment of winter seasonal depression (18). The subjects were asked to maintain their habitual sleep schedules but to adjust them to an earlier time if necessary to accommodate a 1-hour morning
Results

Eighteen outpatients entered the study. Five patients were immediately assigned to treatment. The remaining

18 patients were deemed eligible and were willing to participate in a baseline week of SIGH-SAD assessment and no treatment before beginning treatment; nine of these patients were willing to participate in the 2 baseline weeks of assessment. One week of baseline observation had minimal effect (improvement: mean=13%, SD=22%) on SIGH-SAD scores, while a second week of observation produced no further improvement in the group of nine patients (Figure 1). Two subjects were excluded from analysis. One of them (with a previous stillbirth) experienced another stillbirth 1 week after commencing light therapy and withdrew from the protocol; another patient was noncompliant. The demographic characteristics of the 16 remaining study subjects were as follows: age, mean=34 years (SD=4); all Caucasian; gravity, mean=3 (SD=1); parity, mean=1 (SD=1). Fourteen patients had a past history of major depressive disorder before pregnancy; two of these had winter seasonal depression; and none had a history of dysthymia. Their mean global seasonality score (22) was 8 (SD=6) (two scores were missing), which is indicative only of “slight” seasonal mood variation. Gestation at the start of treatment was 23 weeks (SD=7).

Sixteen patients completed the 3-week protocol, and seven patients completed 2 additional weeks of the study. Twelve patients were observed for a 1-week “withdrawal” period after light treatment. Although the patients were eligible for treatment throughout the year, the timing of patient entry was such that 75% (12 of 16) entered treatment between November 6th and February 6th. Two patients who completed 3 weeks of treatment, we lengthened the study to 5 weeks. For the seven subjects followed

TABLE 1. Outcome Analysis for 16 Pregnant Subjects Undergoing Morning Bright Light Therapy for Antepartum Depression

<table>
<thead>
<tr>
<th>Treatment Period and Outcome</th>
<th>Measure From the Structured Interview Guide for the Hamilton Depression Rating Scale, Seasonal Affective Disorders Version (SIGH-SAD)</th>
</tr>
</thead>
<tbody>
<tr>
<td>3 weeks (N=16)</td>
<td></td>
</tr>
<tr>
<td>SIGH-SAD</td>
<td>Hamilton Depression Rating Scale</td>
</tr>
<tr>
<td>19.1</td>
<td>3, 45                                                               0.0001</td>
</tr>
<tr>
<td>Atypical depression rating subscale 4.0</td>
<td>3, 45                                                               0.0001</td>
</tr>
<tr>
<td>5 weeks (N=7)</td>
<td></td>
</tr>
<tr>
<td>SIGH-SAD</td>
<td>Hamilton Depression Rating Scale</td>
</tr>
<tr>
<td>9.6</td>
<td>5, 30                                                               &lt;0.006</td>
</tr>
<tr>
<td>Atypical depression rating subscale 2.9</td>
<td>5, 30                                                               &lt;0.02</td>
</tr>
<tr>
<td>&lt;0.09</td>
<td></td>
</tr>
</tbody>
</table>

13 patients were deemed eligible and were willing to participate in a baseline week of SIGH-SAD assessment and no treatment before beginning treatment; nine of these patients were willing to participate in the 2 baseline weeks of assessment. One week of baseline observation had minimal effect (improvement: mean=13%, SD=22%) on SIGH-SAD scores, while a second week of observation produced no further improvement in the group of nine patients (Figure 1). Two subjects were excluded from analysis. One of them (with a previous stillbirth) experienced another stillbirth 1 week after commencing light therapy and withdrew from the protocol; another patient was noncompliant. The demographic characteristics of the 16 remaining study subjects were as follows: age, mean=34 years (SD=4); all Caucasian; gravity, mean=3 (SD=1); parity, mean=1 (SD=1). Fourteen patients had a past history of major depressive disorder before pregnancy; two of these had winter seasonal depression; and none had a history of dysthymia. Their mean global seasonality score (22) was 8 (SD=6) (two scores were missing), which is indicative only of “slight” seasonal mood variation. Gestation at the start of treatment was 23 weeks (SD=7).

Sixteen patients completed the 3-week protocol, and seven patients completed 2 additional weeks of the study. Twelve patients were observed for a 1-week “withdrawal” period after light treatment. Although the patients were eligible for treatment throughout the year, the timing of patient entry was such that 75% (12 of 16) entered treatment between November 6th and February 6th. Two patients experienced nausea as a side effect, which was largely ameliorated in one case with a reduction of daily light exposure to 45 minutes. No other significant side effects were reported.

In the 16 patients treated for 3 weeks, SIGH-SAD depression ratings had improved moderately (by 49%) after 3 weeks (improvement between weeks 0 and 2: t=4.49, df=15, p<0.001; improvement between weeks 0 and 3: t=6.27, df=15, p<0.001; and improvement between weeks 1 and 3: t=4.76, df=15, p<0.005; all with Bonferroni correction) (Table 1). After observing these responses in the first nine patients who completed 3 weeks of treatment, we lengthened the study to 5 weeks. For the seven subjects followed

FIGURE 1. Depression Ratings From Baseline to 8 Weeks for Pregnant Subjects Undergoing Morning Bright Light Therapy

![Graph showing depression ratings from baseline to 8 weeks for pregnant subjects undergoing morning bright light therapy.](image-url)

a The SIGH-SAD score is the sum of the ratings for “typical” symptoms on the 21-item Hamilton Depression Rating Scale and those on an 8-item scale for “atypical” depressive symptoms. Subjects participated in up to 2 weeks of baseline observation, 3 or 5 weeks of treatment, and 1 week of withdrawal.

light treatment session. At the end of the treatment trial, the patients stopped light therapy for a 1-week period. No patients were treated in the postpartum.

The patients returned to their clinics for weekly evaluations by psychiatrists and were given SIGH-SAD ratings by clinicians (psychiatrists, psychologists, and social workers) who were blind to treatment status and experienced with SIGH-SAD administration. The time of day of the evaluations was consistent among individual subjects but not across patients. Interrater reliability was assessed by consensus review of the rating criteria before, and every month during, the study. Changes in SIGH-SAD scores over time were evaluated by repeated measures analysis of variance with Greenhouse-Geisser correction (19); post hoc t test comparisons of mean scores with Bonferroni correction were performed for each week. Adverse effects were monitored by using the Systematic Assessment for Treatment Emergent Effects (20), and the subjects were also asked to report any distinct occurrence in the week preceding each evaluation session. Symptoms of mania or hypomania were explored by the evaluating psychiatrists weekly. Symptoms of mania or hypomania were explored by the evaluating psychiatrists weekly.

We obtained global seasonality scores (21) to assess the degree of historical seasonal variation of mood and energy. We performed Pearson’s correlations between degree of seasonality, week of gestation, proportion of the baseline depression score made up of atypical symptoms of depression (e.g., hypersomnia and carbohydrate cravings), and the percentage of improvement on the SIGH-SAD at 3 weeks. After delivery we sent each patient a questionnaire to assess birth outcome and the infants’ health status.
during 5 weeks of light treatment, mean scores on the SIGH-SAD for these subjects improved by 59% from baseline (improvement between weeks 0 and weeks 2, 3, 4, and 5: t=4.00–4.40, df=6, p<0.05, with Bonferroni correction). After 3 weeks, eight of the 16 subjects had a decrease in their SIGH-SAD scores by at least 50%. After 5 weeks of treatment, four of seven subjects achieved a decrease in their SIGH-SAD scores by at least 50%, as well as a final SIGH-SAD score less than 8, which we defined as “complete remission.” Of the seven who completed 5 weeks of treatment, three had complete remissions after 3 weeks, and four achieved complete remission by the fifth week. During the 1 week of withdrawal of light therapy, SIGH-SAD scores increased by 5 (SD=10) in the 12 patients we observed. Five of these 12 patients had an increase in their SIGH-SAD scores by at least 5 points.

There was no significant correlation between improvement in SIGH-SAD scores after 3 weeks and week of gestation (r=0.02, p=0.93, N=16) or between improvement on SIGH-SAD scores and baseline score on the atypical depression portion of the SIGH-SAD (atypical depression score divided by total SIGH-SAD score: r=0.14, p=0.60, N=16). Furthermore, there was no significant correlation between seasonality scores and improvement on the SIGH-SAD after 3 weeks of treatment (r=0.10, p=0.73, N=14; data for two patients were missing).

Fourteen of the 16 patients returned the follow-up questionnaire. One mother had one male and one female twin; the other mothers produced seven boys and six girls. Continued light therapy provided beneficial effect in the treatment responders who used the treatment after the trial and throughout the remainder of the pregnancy. After completion of the study, depression returned during pregnancy for four of the 14 patients, within 3 months of delivery for another four, and later for two more patients. Nine had normal vaginal deliveries; five had Cesarean sections. Mean birth weight was 110 oz (SD=26). Only one of the mothers did not gain weight at a normal pace for her date of pregnancy; her son was born at 30 weeks. Another infant was born prematurely at 32 weeks. In the six children in which it was known, the mean Apgar score at birth was 8 (SD=2). All 15 children were reported to have experienced “normal” weight gain, growth, and attainment of milestones.

Discussion

This pilot study demonstrated that significant improvement occurred with an intervention of active light therapy. Withdrawal of the treatment was associated with an increase in depressive symptoms. The data support the hypothesis that morning bright light therapy has an antidepressant effect on pregnant women with depression. In comparison with light treatment for winter depression, the rate of improvement in this study appeared more gradual than most and more with a slow-onset psychopharma-}

ological effect. The net treatment effect after several weeks with 50% of the patients meeting stringent remission criteria is similar to that seen in a controlled trial of morning light therapy for winter depression (23). Although two of the 18 patients treated with light were excluded from data analysis, robust treatment effects were seen in the women remaining in the trial. No significant safety issues emerged from the trial. We note that nausea also has been previously singled out as an emergent side effect of successful light therapy for treatment of winter depression (24).

The Cochrane Collaboration (25) has concluded that light is effective for the treatment of winter depression, but it also concluded that more studies are needed for the evaluation of light therapy in nonseasonal depression. Our data did not yield any specific factors associated with response to light, although a number of patients without a history of recurrent winter depression clearly had excellent responses. A randomized controlled trial is warranted to ascertain that improvement is specific to bright light exposure in antepartum depression and not to a placebo effect of open treatment.

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References

Brief Report

Åøegaard’s Selection Hypothesis Revisited:
Schizophrenia in Surinamese Immigrants to the Netherlands

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Elizabeth Cantor-Graae, Ph.D.
Joris Slaets, M.D., Ph.D.
René S. Kahn, M.D., Ph.D.

Objective: The incidence of schizophrenia among Surinamese immigrants to the Netherlands is high. The authors tested Ødegaard’s hypothesis that this phenomenon is explained by selective migration.

Method: The authors imagined that migration from Surinam to the Netherlands subsumed the entire population of Surinam and not solely individuals at risk for schizophrenia. They compared the risk of a first admission to a Dutch mental hospital for schizophrenia from 1983 to 1992 for Surinamese-born immigrants to the risk for Dutch-born individuals, using the Surinamese-born population in the Netherlands and the population of Surinam combined as the denominator for the immigrants.

Results: The age- and sex-adjusted relative risk of schizophrenia for the Surinamese-born immigrants was 1.46.

Conclusions: Selective migration cannot solely explain the higher incidence of schizophrenia in Surinamese immigrants to the Netherlands.

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There have been consistent reports of a greater incidence of schizophrenia among certain immigrant groups to Western Europe (1). These groups include, among others, first- and second-generation Caribbean immigrants to the Netherlands.