Major depression occurs in 10%–30% of older medical inpatients\(^1\)–\(^4\) and appears to be associated with poor functional status,\(^5\) increased use of hospital services\(^6,7\) and reduced survival,\(^6,8–10\) independent of the severity of physical illness. Despite the apparent benefits of treatment,\(^1\) up to 90% of cases of depression are not detected by attending physicians during the course of usual hospital care,\(^3,12\) and few patients receive optimal treatment in hospital or after discharge.\(^12,13\) Because a strategy of systematic detection and treatment has been shown to benefit elderly people with depression living in the community,\(^14–18\) we conducted a randomized clinical trial to evaluate whether systematic detection and multidisciplinary treatment of depression could be effective in reducing symptoms of depression and improving mental and physical health status in elderly medical inpatients. Our secondary objective was to determine the effect of the intervention on basic and instrumental activities of daily living, cognitive status, side-effects profile, mortality and health services utilization (length of hospital stay, and number of readmissions and emergency department visits).

**Methods**

We conducted the study at St. Mary’s Hospital Center, a university-affiliated, primary acute care hospital in Montréal. The trial was approved by the hospital’s Research Ethics Committee.

All patients aged 65 years and over admitted from the emergency department to medical services between Oct. 19, 1999, and Nov. 1, 2002, were screened for eligibility by the research nurse. We excluded patients who (a) were admitted to the intensive care unit or cardiac monitoring unit for more than 48 hours; (b) had an imminently terminal illness; (c) did not speak or understand English or French; and (4) did not live on the Island of Montreal. Eligible patients who scored 4 or less on the Short Portable Mental Status Questionnaire (indicating at most mild cognitive impairment) were assessed using the Diagnostic Interview Schedule.\(^19\) Patients who were found to have major depression (as defined by DSM-IV criteria\(^20\) and consented to participate were enrolled.

Enrolled patients were allocated to the intervention or usual-care groups with the use of block-size randomization and an allocation ratio of 1:1. The block size was varied ran-
domly to improve blinding. The study statistician prepared a series of sealed envelopes containing the treatment allocation. For each enrolled patient, the research nurse opened the next envelope on top of the pile to determine his or her group allocation.

The framework for the intervention was adapted from a methodology to assess geriatric services. The intervention group received systematic treatment for 24 weeks. The treatment was provided in 3 parts: (a) assessment and treatment by a psychiatrist in the hospital's geriatric service; (b) follow-up by the research nurse; and (c) follow-up by the patient's family physician. The psychiatrist assessed each patient and made management recommendations, all recorded on the regular hospital consultation form and signaled in the progress notes. Treatment involved supportive psychotherapy and drug therapy with an antidepressant (e.g., a selective serotonin reuptake inhibitor [SSRI], a tricyclic antidepressant or methylphenidate), prescribed according to clinical practice guidelines. Patients were seen as often as necessary during their hospital stay and after discharge. When the patients were seen by their family physician for follow-up, the psychiatrist was informed of their progress by the research nurse. The research nurse visited the patients at least weekly in hospital and visited or telephoned them weekly after discharge for 24 weeks to monitor their condition, provide supportive psychotherapy, ensure maximum compliance with their treatment and liaise with the family, psychiatrist and family physician. The intervention team (comprising 2 psychiatrists from the geriatric service and the research nurse) met regularly to assure consistency in the diagnosis and management of depression.

Subjects assigned to the control group received usual care before and after discharge. There is no evidence that the proposed intervention improves outcomes of medical inpatients. This study proposed enhanced care for depressed patients and involved only the risks associated with accepted treatments of depression. Subjects in the usual-care group were informed that they had major depression and advised to discuss treatment with their physician, but they received no systematic intervention or follow-up. Hospital staff were not informed when usual-care subjects were enrolled in the study, in order to minimize contamination. Referrals for psychiatric consultation were honoured, consistent with usual practice.

Data were collected at baseline and at 3 and 6 months later by research assistants blind to group allocation. Measures at baseline included 3 measures of physical illness (Clinical Severity of Illness, the Charlson Comorbidity Index, and an acute physiology score, derived from 12 physiologic variables of APACHE II). Alcohol abuse was determined with the use of the CAGE questionnaire. Measures at baseline and follow-up included the Diagnostic Interview Schedule (symptoms were counted toward the diagnosis of depression regardless of their origin [physical illness or depression]); the 21-item Hamilton Depression Rating Scale (HAMD), the

Fig. 1: Flow of patients through the trial. SPMSQ = Short Portable Mental State Questionnaire, ICU = intensive care unit, CMU = cardiac monitoring unit, DIS = Diagnostic Interview Schedule, R = randomization. *Three patients withdrew and 1 patient died before the intervention.
Medical Outcomes Study 36-item Short Form (SF-36);\textsuperscript{29} the Mini-Mental State Examination (MMSE);\textsuperscript{30} the Older Americans Resources and Services (OARS) questionnaire to assess basic and instrumental activities of daily living (OARS-ADL and OARS-IADL);\textsuperscript{31} and the Rating Scale for Side Effects (RSSE)\textsuperscript{32} (modified to include the side effects of SSRIs and methylphenidate). Measures at follow-up included mortality, suicide and suicide attempts, and health services utilization (length of hospital stay, and number of readmissions and emergency department visits). Information about antidepressant use was collected from 3 sources: the hospital chart (drug use at admission and discharge), the research assistant’s notes (drug use at the 6-month follow-up) and the provincial drug database (prescriptions filled 1 month before admission, 2 months after discharge and 5–7 months after enrolment). The psychiatrists and the research nurse recorded their contacts with the patients, families, family physicians and each other.

The interrater agreement (kappa value) between the psychiatrist and research nurse and between the 2 research assistants for major versus no major depression was 1.0 (n = 9) and 0.78 (n = 28) respectively. The research assistants’ intra-class correlation coefficient (ICC) was 0.99 (n = 28) for the HAMD and 1.0 (n = 18) for the SF-36. For the other measures, the ICC ranged from 0.99 to 1.0 (n = 14–28).

Intervention and control groups were compared at baseline. Patients who completed both baseline and follow-up interviews were compared with patients who withdrew from the study or died.

To evaluate the effect of the intervention, we estimated the difference between the intervention and control groups in the change in each primary outcome variable (i.e., HAMD, and SF-36 summary scores for the physical and mental health components) between baseline and the 6-month follow-up. An intention-to-treat approach was used. This analysis was based on all observed data with the assumption that data were missing at random.

Using the HAMD, we defined improvement in major depression as a decrease in the HAMD score of 50% or greater at 6 months compared with baseline; we defined remission of major depression as a HAMD score of less than 7 points at 6 months. We estimated the difference between the intervention and usual-care groups in the proportion of patients who showed improvement, remission or major depression (according to DSM IV criteria) at 6 months. We also estimated the difference between the 2 groups in the following outcomes: change from baseline to 6 months in the OARS-ADL, OARS-IADL, MMSE and RSSE scores, length of hospital stay, and incidence during the 6 months after baseline of admission to hospital, suicide or suicide attempts, visits to the emergency department and death.

For both primary and secondary outcomes, we compared continuous variables using the t test and discrete variables using the Fisher’s exact test. To compare length of stay during the index admission between the 2 groups, we used Wilcoxon’s rank sum test because of the non-normal distribution of this variable. We report 95% confidence intervals for the differences in all outcomes. For differences in proportions, we calculated bootstrap confidence intervals because of the small samples. We repeated the analysis of both primary and secondary outcomes while adjusting for baseline characteristics of patients and process-of-care variables.

Although not planned in the protocol, we carried out additional analyses by including the 3-month data for the primary outcomes. A mixed-effects regression model was fit that adjusted for the dependence between multiple observations for each patient.

The dropout rate was high, and the resulting sample of patients with complete data had an underrepresentation of severely ill subjects. Therefore, we repeated our analysis of both primary and secondary outcomes by imputing missing data, with the last measure carried forward. This allowed us

### Table 1: Baseline characteristics of elderly medical inpatients with major depression in the intervention and usual-care groups

| Characteristic | Intervention group (n = 78) | Usual-care group (n = 79) |
|---------------|-----------------------------|--------------------------|
| Age, mean (SD), yr | 77.5 (6.7) | 78.5 (6.6) |
| Female sex, % | 69.2 | 69.6 |
| Living arrangement, % | | |
| Home alone | 48.7 | 51.3 |
| Home with spouse | 18.0 | 20.5 |
| Home with family | 16.7 | 12.8 |
| Other* | 16.7 | 15.4 |
| Past history of depression, % | 14.5 | 15.4 |
| Duration of current depression ≥ 2 yr, % | 24.4 | 30.4 |
| Alcohol use (CAGE score > 0), % | 12.5 | 10.2 |
| Has a confidant, % | 74.6 | 84.2 |
| Death of someone close in past year, % | 46.8 | 42.9 |
| Drug use, % | | |
| Psychotropic | 46.2 | 53.2 |
| Antidepressant | 25.6 | 27.9 |
| Severity-of-illness score, mean (SD) | 4.3 (0.9) | 4.2 (1.1) |
| HAMD score, mean (SD) | 21.3 (5.5) | 20.1 (5.9) |
| SF-36 score, mean (SD) | | |
| Mental component | 37.4 (10.9) | 34.3 (13.2) |
| Physical component | 32.8 (7.2) | 33.9 (11.7) |
| MMSE score, mean (SD) | 24.9 (3.5) | 23.8 (4.9) |
| OARS-ADL score, mean (SD) | 11.5 (2.4) | 11.2 (2.5) |
| OARS-IADL score, mean (SD) | 9.6 (2.9) | 9.6 (2.5) |
| RSSE score, mean (SD) | 18.3 (6.5) | 18.2 (6.7) |
| Acute physiology score, mean (SD) | 2.5 (2.3) | 2.2 (2.0) |
| Charlson Comorbidity Index, mean (SD) | 2.1 (1.9) | 1.7 (1.7) |

Note: SD = standard deviation, HAMD = Hamilton Depression Rating Scale, SF-36 = Medical Outcomes Study 36-item Short Form, MMSE = Mini-Mental State Examination, OARS-ADL and OARS-IADL = Older Americans Resources and Services questionnaire assessing basic and instrumental activities of daily living, RSSE = Rating Scale for Side Effects.

*Includes foster home, senior residence and nursing home.
to include data for all patients who had completed the baseline interview, which reduced the proportion of missing data from 50% to about 29%. We also examined the sensitivity of the results by repeating the analyses after excluding (a) patients in the intervention group who did not receive a consultation and those in the control group who did receive a consultation, and (b) patients taking antidepressants at baseline.

The sample-size calculations were based on the formula for comparison of 2 means for one principal outcome, the HAMD. In a study involving elderly ambulatory patients, we had found a mean difference in HAMD scores between the intervention and usual-care groups of 5–7 points. Among physically ill and disabled elderly medical inpatients, we anticipated that the difference would be less (i.e., 4 points). On the basis of an estimated standard deviation of change in the HAMD score of 8.0, we determined that 85 subjects would be required in each group to detect a change of at least 4 points in the HAMD score. However, because the attrition rate was higher than expected (60%), the sample-size calculation was repeated midway through the study. On the basis of a standard deviation of change in the HAMD score of 5.6 (derived from the data collected), we determined that 31 subjects in each group would be required to detect a change of at least 4 points in the HAMD score with 80% power and a 2-sided significance level of 0.05. Allowing for an attrition rate of 60%, we needed to recruit 78 patients in each group.

**Results**

The flow of patients through the study is shown in Fig. 1. Of the 1500 inpatients screened, 225 (15.0%) were found to have major depression. There were no deviations from the study protocol. Of the 157 patients who consented to participate in the study, 78 were allocated to the intervention group and 79 to the usual-care group. At randomization, there were no clinically significant differences between the groups (Table 1).
Sixty-four subjects (40.8%) completed the follow-up to 6 months; 57 (36.3%) withdrew (almost all before 3 months), and 36 (22.9%) died before the 6-month follow-up (Table 2). Those who withdrew had more severe physical illness than those who completed the 6-month follow-up. There were statistically significant differences in sex, severity of illness and Charlson Comorbidity Index scores between those who died and those who completed the 6-month follow-up.

Seventy-four patients (94.6%) in the intervention group received a consultation, which occurred a mean 1.2 days after enrolment; of the remaining 4 patients, 3 withdrew and 1 died. The mean number of follow-up contacts per patient by the psychiatrists and research nurse are presented in Table 3. Twenty-one patients (26.6%) in the usual-care group received a consultation, which occurred a mean 2.4 days after enrolment.

The rates of antidepressant use in the intervention and usual-care groups are presented in Table 3. There were differences in the rates by source of information (chart v. provincial drug database v. self-report). Antidepressants were not recommended for 8 (25%) of the intervention patients because their depressive symptoms were mild or because potential risks outweighed potential benefits.

At 6 months, there were no statistically significant differences between the intervention and usual-care groups in the primary outcomes (Table 4). There were no differences in the rates of improvement (≥ 50% decrease in HAMD score) or in the rates of remission (HAMD score < 7) of depression. There was a potentially clinically important (but not statistically significant) difference in the rates of major depression (intervention 45.5% v. control 56.7%). The results were similar when 3-month data were included.

In the secondary analyses, we found no important relations between the baseline patient characteristics (age, sex, living arrangement, severity of illness, Charlson Comorbidity Index, OARS-ADL, OARS-IADL or HAMD scores, or duration of depression), process-of-care variables (number and type of contacts, use of antidepressant medication) and outcomes.

The results of the sensitivity analysis of primary and secondary outcomes were similar to those of the main analysis.

**Interpretation**

We proposed to determine the effectiveness of a strategy of systematic detection and multidisciplinary care of depression in elderly medical inpatients. We were unable to demonstrate that the intervention was beneficial, perhaps because of the high patient attrition rate, the low number of contacts between patients and psychiatrists (mean 4.5), the suboptimal compliance with antidepressant medication or possible contamination of the usual-care group (patients in both groups were managed on the same units by the same attending physicians).

Our results are similar to those of a recently published trial of care management of anxiety and depression among elderly veteran inpatients.39 That intervention involved assessment by a multidisciplinary team and assignment of a case coordinator who engaged patients in treatment and follow-up. The rate of attrition through withdrawal and death was as high as our rate (60%). As in our study, there were no significant benefits of the intervention.

Our study has several potential limitations. It was difficult to maintain the participation of physically ill and depressed subjects (57 withdrew, and 36 died); even though there were no important differences between the groups in rates of withdrawal or death and no clinically important differences in baseline characteristics between those who completed the study and those who withdrew, this attrition rate may have affected the results. A quarter of enrolled patients had been depressed for 2 or more years; these patients may have been less likely to respond to the intervention; however, we found no relation between duration of depression and outcomes. A quarter of enrolled patients were taking antidepressant medication on admission, which may have reduced the effect of the intervention; however, there were no differences in outcomes when those taking antidepressants at baseline were excluded from the analysis. Arguably, the intervention group

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**Table 3: Process of care of subjects who completed the 6-month follow-up**

| Variable                                      | Intervention group n = 33 | Usual-care group n = 31 | p value |
|-----------------------------------------------|---------------------------|-------------------------|---------|
| Contacts by treating psychiatrist, mean no.   |                           |                         |         |
| With the patient                              | 4.5                       | 0.3                     |         |
| With the intervention nurse                   | 9.4                       | 0.0                     |         |
| With the family                               | 0.3                       | 0.0                     |         |
| With the family physician                     | 0.1                       | 0.0                     |         |
| Contacts by research nurse, mean no.          |                           |                         |         |
| With the patient                              | 16.7                      | 0.0                     |         |
| With the family                               | 2.9                       | 0.0                     |         |
| With the family physician                     | 0.8                       | 0.0                     |         |
| Mean no. of psychiatry outpatient visits      | 2.5                       | 0.3                     | < 0.001 |
| Antidepressant use, no. (%) of patients       |                           |                         |         |
| At admission                                  |                           |                         |         |
| Chart                                         | 11 (33.3)                 | 9 (29.0)                | 0.71    |
| RAMQ                                          | 9 (27.3)                  | 11 (35.5)               | 0.48    |
| At discharge                                  |                           |                         |         |
| Chart                                         | 19 (57.6)                 | 11 (35.5)               | 0.08    |
| RAMQ                                          | 20 (60.6)                 | 14 (45.2)               | 0.22    |
| At 6-mo follow-up                             |                           |                         |         |
| Self-report                                   | 19 (57.6)                 | 11 (35.5)               | 0.08    |
| RAMQ                                          | 17 (51.5)                 | 14 (45.2)               | 0.61    |

*Psychological treatment received by patients in the intervention and usual-care groups is not included here because we were unable to collect this information from the hospital chart or from the patients at follow-up.
could have received interpersonal or cognitive-behavioural therapy (in addition to the antidepressant medication), but these therapies were not feasible in our study. The low number of follow-up contacts between the treating psychiatrists and the patients (mean 4.5), ostensibly owing to difficulties in attending clinic after discharge, may also have reduced the effect of the intervention; however, the number of contacts between the research nurse, patients, psychiatrists, family members and family physicians facilitated implementation of the intervention. Patients in both groups were managed on the same units by the same attending physicians; consequently, there may have been contamination of usual care by the intervention program, or there may have been a Hawthorne effect, whereby implementation of the study resulted in improved detection and care of all patients with depression. In the usual-care group, the rates of request for psychiatric consultation and antidepressant therapy prescribed at discharge were 26.6% and 35.5%–45.2%, respectively, much higher than reported in descriptive studies.\textsuperscript{3,12,13} Controversibly, the rate of antidepressant use in the usual-care group was high (29%–35.5%) on admission. These results are disappointing, but they should not discourage future trials of innovative interventions for the detection and management of depression in elderly inpatients. Such interventions might include extensive patient and family education, more aggressive pharmacotherapy, physical rehabilitation to improve activities of daily living, enhanced social supports or modified cognitive-behavioural therapy for the helplessness and disempowerment often associated with declining physical health. Finally, it may be necessary to use cluster randomization or a nonrandomized study design instead of randomization, to avoid contamination of the usual care.

Table 4: Effect of intervention on outcome measures

| Outcome measure | Intervention group n = 33 | Usual-care group n = 31 | Difference between groups (95% CI) |
|-----------------|--------------------------|-------------------------|-----------------------------------|
| Primary         |                          |                         |                                   |
| Difference in mean score from baseline to 6-mo follow-up |                          |                         |                                   |
| HAMD            | -6.3                     | -5.0                    | -1.3 (-4.9 to 2.2)               |
| SF-36, mental component | 9.4 | 9.2 | 0.2 (8.7 to 8.9) |
| SF-36, physical component | -2.9 | -2.7 | -0.2 (5.4 to 5.0) |
| Secondary       |                          |                         |                                   |
| Improvement (≥ 50% decrease in HAMD score at 6 mo) | 28.1 | 20.0 | 8.1 (-13.3 to 29.3) |
| Remission (HAMD score < 7 at 6 mo) | 15.6 | 16.7 | -1.1 (-19.4 to 17.3) |
| Major depression (DSM-IV criteria) at 6 mo | 45.5 | 56.7 | -11.2 (-35.8 to 13.3) |
| Difference in mean score from baseline to 6-mo follow-up |                          |                         |                                   |
| OARS-ADL        | -1.0                     | -0.8                    | -0.2 (-1.6 to 1.3)               |
| OARS-IADL       | -1.4                     | -1.0                    | -0.4 (-2.0 to 1.3)               |
| MMSE            | 1.5                      | 0.9                     | 0.6 (-1.1 to 2.2)                |
| RSSE            | -3.3                     | -4.0                    | 0.7 (2.6 to 4.0)                 |
| Length of hospital stay (index admission), median, d | 12.0 | 10.0 | 2.0 (6.5 to 6.5) |
| Event during 6-mo follow-up, % |                          |                         |                                   |
| Readmission     | 39.4                     | 29.0                    | 10.4 (21.3 to 23.5)              |
| Suicide or suicide attempt | 3.2 | 3.3 | 0.1 (9.8 to 9.4) |
| Any visit to emergency department | 45.5 | 41.9 | 3.6 (24.5 to 24.7) |
| Mortality at 6-mo follow-up, % | n = 78 | n = 79 | 0.3 (12.5 to 13.1) |

Note: CI = confidence interval, HAMD = Hamilton Depression Rating Scale, SF-36 = Medical Outcomes Study 36-item Short Form, MMSE = Mini-Mental State Examination, OARS-ADL and OARS-IADL = Older Americans Resources and Services questionnaire assessing basic and instrumental activities of daily living, RSSE = Rating Scale for Side Effects.
Contributors: Martin Cole, Jane McCusker and Nandini Dendukuri contributed to the conception and design of the study, the analysis and interpretation of the data, and the drafting and revision of the article. Michel Elie contributed to the acquisition, analysis and interpretation of the data and the revision of the article. Eric Latimer and Eric Belzile contributed to the analysis and interpretation of the data, and the revision of the article. All of the authors approved the final version for publication.

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Editor’s note

• Most elderly medical inpatients have serious chronic diseases, and many have a limited life expectancy because of their disease burden. It is not surprising that major depression is prevalent among these patients; however, the condition goes undetected and untreated in most cases. Will a strategy of systematic detection and multidisciplinary management of depression help these patients?

• Consecutive elderly patients admitted to medical services in a primary care hospital who were found to have major depression were randomly assigned to receive the intervention or usual care. The intervention involved assessment and treatment by a psychiatrist and follow-up by a nurse and the patient’s family physician.

• No benefit from the intervention could be shown. The finding is limited by the power of the study and the fact that some of the patients in the usual-care group received treatment for depression.

Clinical implications: No additional benefit was found for more intensive surveillance and treatment of major depression in medical inpatients. More study is needed, but any unintended benefit is likely to be small.