Background: Life expectancy is increasing continuously, which increases the likelihood of developing dementia or cancer. Both dementia and cancer are serious conditions that give manifold symptoms. The interaction of these conditions is however complex and less explored. Objectives: The aim of this study was to identify the prevalence of cancer and differences regarding neuropsychiatric symptoms (NPS) and medication among nursing home (NH) patients with and without dementia and cancer. Methods: This is a cross-sectional study of Norwegian NH patients (N=1825). Participants were categorized according to degree of dementia (Clinical Dementia Rating > 1) and cancer diagnoses. Differences in NPS and other symptoms, as well as the use of medication, were explored. Results: Eighty-four percent of NH patients had dementia, and 5.5% had comorbid dementia and cancer. Patients with comorbid dementia and cancer received significantly more analgesics compared with patients without cancer but with dementia (P<.05). Compared with patients without dementia but with cancer, patients with comorbid dementia and cancer had significantly more NPS, including sleep disturbances and agitation. Conclusions: Patients with comorbid dementia and cancer receive more analgesics than patients with dementia but still display more agitation and sleep disturbances than patients with cancer and patients with neither dementia nor cancer, suggesting that symptoms may...
Life expectancy is continuously increasing worldwide, and the likelihood of developing dementia is thereby growing dramatically. Dementia, of which the most common cause is Alzheimer’s disease, is an incurable and progressive condition. It affects 35 million people worldwide—a number expected to almost double in the next 2 decades. In general, dementia results in the decline of the person’s cognition and physical function. Up to 90% of people with dementia experience neuropsychiatric symptoms (NPS), such as agitation, depression, and delusion, during the course of their disease. In total, 77,000 Norwegians are affected by this disease, and approximately 14% of Norwegians older than 80 years receive institutional care.

Cancer diseases affect approximately 10 million people in Europe, and 30,000 new cases annually are expected. As a result of the demographic development, the prevalence is expected to double during the next 10 years. By the end of 2014, 242,398 Norwegians were registered with a cancer diagnosis, and the disease was the cause of death for 10,971 people. In Norway, 3 of 4 cancer cases are diagnosed in those older than 60 years, and a quarter of all terminal patients with cancer die in a nursing home (NH). Previous studies indicate that from 14% to 26% of Norwegian NH patients are given a diagnosis of cancer, and international studies show that approximately 5% were receiving active cancer treatment when being admitted to an institution. It is unknown how many develop cancer during the time they live in an NH and how this is discovered and treated.

The likelihood of developing comorbid dementia and cancer increases with age. Cancer symptoms are potentially more onerous for people with both conditions because they are no longer able to give a valid self-report—a prerequisite for adequate treatment. As suggested by symptom science, the management of symptoms needs to be based on an understanding of the individual, the condition, and the context in which the individual resides.

Ideally, the assessment of cancer-related pain and symptoms should be based on the individual’s self-report with subsequent administration of analgesics and appropriate symptom management. However, people with dementia express their pain in ways that are different from those without cognitive impairment, leading to agitation and depression. Furthermore, previous studies show that sleep disturbances are related to the presence of untreated pain and to depression. In light of recent research, which indicates that benzodiazepines have no effect on sleep time between users and nonusers, there is a need for awareness around these circumstances because they can contribute to potentially meaningless treatment and widespread use of psychotropic drugs.

In a recent review, our group summarized the evidence on cancer-related symptoms among NH patients and highlighted 11 studies, which together confirmed high prevalence of pain and reduced analgesic drug prescription in NH patients with cancer. Only 1 smaller study (n=48) included patients with comorbid dementia and cancer, and this study found that people with severe dementia received less opioids and displayed less pain behavior compared with patients with mild cognitive impairment. Another study by Monroe et al found that patients with comorbid dementia and cancer were more likely to receive an opioid if they were enrolled in hospice. However, patients enrolled in hospice were more cognitively intact.

The overall purpose of the study is to investigate comorbid dementia and cancer among NH residents, with a particular emphasis on NPS and the medication thereof. The study has the following research objectives: (1) to investigate the prevalence of cancer among NH patients with and without dementia and (2) to compare patients with comorbid dementia and cancer with those without, with regard to the prevalence of NPS such as agitation, depression, and sleep disturbances and the daily use of medication, including analgesic drug prescription.

**Method**

**Design, Setting, and Procedure**

The study builds on secondary analyses of a multicenter, cross-sectional study including 64 NHs, located in urban and rural areas in 5 of 19 counties of Norway. The data set includes standardized interviews with NH staff, as well as information from patients’ medical records. The organization of the setting and the participants of the study have been previously described by Helvik et al wherein this data set was used alongside data from 2004. In brief, Norway has around 34,000 long-term NH patients, and the jurisdiction for public healthcare services is the responsibility of the municipalities. In the process of gaining a representative sample of small, medium-sized, and large municipalities from both urban and rural areas, 49 municipalities were approached, and the sample of NHs was selected.

The study was conducted from June 2010 to November 2011. All patients in the NHs were screened for inclusion, and all patients who had lived in the NH for at least 2 weeks were eligible. Project nurses were responsible for data collection. Before data collection, they participated in a 2-day session on the use of mapping instruments and standardized interviews, as well as on the purpose of the study. Data collection involved registering information from each patient’s medical record, as well as performing standardized interviews with the patient’s primary caregiver, who served as a proxy for the patient, regardless of the patient’s degree of cognitive impairment.
Measurements

We classified the patient’s level of dementia by the Clinical Dementia Rating (CDR) scale (range, 0–3). The instrument consists of 6 categories with scores for no dementia of 0, possible dementia of 0.5, and mild, moderate, and severe dementia of 1, 2, and 3, respectively. The cutoff point for the presence of dementia is CDR of 1 or greater. The CDR scoring algorithm weights memory as the primary domain and the other domains as secondary.31 Good validity and reliability of the scale have been reported (overall $\kappa = 0.62$).32–34

Cancer diagnoses were collected from patients’ records and coded by means of the International Classification of Diseases, Tenth Revision.35 Ongoing medical treatment from patients’ records was coded by the Anatomical Therapeutic Chemical (ATC) System—an international system for the classification of drugs.36 We divided analgesics into 3 groups: (1) peripheral (ATC, N02B), (2) opioids (ATC, N02A), and (3) nonsteroidal anti-inflammatory drugs, namely, anti-inflammatory and anti-rheumatic drugs (ATC, M01). We similarly collected data on the following psychotropic drugs: antidepressants (ATC, N06A), anxiolytic drugs (ATC, N05B), sedatives (ATC, N05C), antidementia drugs (ATC, N06D), and antipsychotics (ATC, N05A).

We assessed NPS related to dementia using the Neuropsychiatric Inventory-Nursing Home version (NPI-NH).37 The NPI-NH maps 12 NPS associated with dementia: delusions, hallucinations, dysphoria/depression, anxiety, agitation/aggression, euphoria, disinhibition, irritability/ability, apathy, aberrant motor behavior, nighttime behavior, and appetite disturbances/eating change (all judged as no/yes). Each symptom was scored for frequency (score, 1–4) and severity (score, 1–3), and a product score (0–12) was calculated thereof for each symptom. In this study, we used a previously identified underlying 4-factor structure,38 in which items are clustered into subsyndromes for agitation (agitation and irritability), affective symptoms (depression and anxiety), and psychosis (hallucination and delusions), whereas apathy is analyzed as a single symptom. In addition to the use of these NPS factors, sleep disturbances were identified using the nighttime behavior item from the NPI-NH. Scores of 1 or higher suggest impaired sleep, and in accordance with previous studies, anyone scoring higher than or equal to 4 on this item was judged as having clinically significant sleep disturbances.39

We used the Cornell Scale for Depression in Dementia (CSDD) to measure depression, an instrument that is found to be a reliable measure for depression in people with dementia (Cronbach’s $\alpha = .84$).40,41 We used the General Medical Health Rating (GMHR) to assess patients’ degree of somatic illness. It is a reliable and valid measure, with 1 item comprising 4 categories, which assesses medical comorbidity (weighted $\kappa = 0.93$).42 The 4 categories were excellent, good, fair, and poor. The rating was performed based on all available medical information. In addition, we used the Physical Self-Maintenance Scale (PSMS), a reliable measure of patients’ level of function with regard to activities of daily living. A high score indicates a lower level of functioning.43,44

Statistical Analyses

We calculated the mean, standard deviation (SD), and range for patient characteristics (number of diagnoses, regular drugs, and analgesics) and calculated the product score for the NPI-NH and its 4 subclusters and sum scores for the CSDD and the PSMS. For the GMHR, we created a binary variable from the 4 categories of the scale simply reflecting good or bad health. Patients who had excellent or good health were recorded as “good,” whereas those with fair or poor health were recorded as “bad.” For all variables, means were compared across patient groups, and the statistical significance of the differences was tested by means of analyses of variance and logistic regressions for continuous and categorical variables, respectively. In the analyses, we divided the final sample into 4 groups, which we refer to throughout the text and tables in the following way: patients with comorbid dementia and cancer (COMORBID), patients with cancer but without dementia (CANCER), patients with dementia but without cancer (DEMENTIA), and patients who have neither dementia nor cancer (NEITHER).

Finally, we conducted a multiple logistic regression investigating the likelihood of analgesic drug prescription to investigate the characteristics associated with the use of analgesics, while controlling for other variables. The dependent variable was a categorical variable indicating whether the patient received analgesics regularly. The independent variables were the following: cancer, dementia, comorbidity, GHMR, PSMS, CSDD, NPI-NH, regular drugs, use of sedatives, age, and gender. We conducted the statistical analyses in IBM SPSS Statistics 22.

Ethical Approval

We obtained verbal and written information in direct conversation with the patient (if possible) and his/her legal guardian, usually a family member or an advocate, in accordance with local law. Depending on the patient’s ability to give consent, the patient or next of kin made the decision of participation in the study after we explained the aims and protocol of the study. The Regional Committee for Medical and Health Research Ethics (REC West) in Norway approved the study (REC number 2010/1894).

Results

There were 2385 residents eligible for inclusion. From this sample, 560 patients were excluded for the following reasons: the patient or next of kin declined participation (n=423), serious somatic illness or terminal conditions (n=33), death before data collection (n=17), leaving the NH before assessment (n=1), and either missing data or unlisted reasons (n=53). In addition, 33 patients were excluded from the analyses because of the lack of CDR scores. The final sample was composed of 1825 participants. Seventy-one percent of the participants were women, and the mean (SD) age was 85.1 (8.94) years. Baseline clinical characteristics are reported in Table 1.
Prevalence of Dementia and Cancer Among NH Patients

The total sample consisted of 1825 patients. One hundred thirty-three patients (7.3%) had cancer, of which 100 patients had comorbid dementia and cancer (5.5%) and 33 patients had a cancer diagnosis but no dementia (1.8%). Thus, the prevalence of cancer is significantly higher among patients without dementia than among those with dementia (P<.01).

Most common was breast cancer for women (45%) and prostate cancer for men (31%). Colorectal cancer was the second most common cancer diagnosis for both genders (17% of cancer cases overall). Cognitive impairment consistent with dementia (CDR≥1) was found in 1535 patients (84%); 29% had moderate dementia, and 36% had severe dementia. The most frequent diagnoses from the NH medical record were unspecified dementia and Alzheimer’s disease (29% and 12%, respectively), hypertension (27%), and apoplexia cerebri (15%). Regarding our first research objective, our findings reveal higher prevalence of cancer among patients without dementia than among those with dementia. Compared with cancer prevalence in NHs documented in previous research, the prevalence of cancer in this sample is quite low.

Comparison of NPS and Medication Use for the Different Patient Groups

Compared with CANCER patients (n=33), COMORBID patients (n=100) displayed more NPS based on NPI-NH sum score (P<.01) for the agitation cluster score (P<.05) and the apathy item score (P<.05). These individuals also displayed worse physical function (P<.01) and had significantly more sleep disturbances (P<.01). We found no differences between the groups regarding the use of medication or in socio-demographic characteristics (see Table 2).

Compared with DEMENTIA patients (n=1435), COMORBID patients had significantly more diagnoses (P<.01) and received more analgesics (P<.05), in particular, opioids (P<.05). There were no other significant differences between the 2 groups (see Table 2).

Compared with NEITHER patients (n=257), COMORBID patients had significantly reduced physical function and more NPS as measured by the NPI-NH sum score and agitation and psychosis cluster scores (P<.01) (see Table 2). Similarly, they had higher scores for singular behavioral disturbances such as apathy, sleep disturbances, and depression assessed by the CSDD. They also had more comorbidity and worse somatic health according to
NEITHER, Patients with neither dementia nor cancer; NSAIDS, nonsteroidal anti-inflammatory drugs.

That is, patients with cancer were almost twice as likely to receive CSDD, and cancer. The highest odds ratio was 1.974 for cancer.

The table shows differences in mean scores between the 4 patient groups (COMORBID, CANCER, DEMENTIA, and NEITHER) for each of the variables outlined in Table 1. For instance, the column “Difference Between COMORBID and CANCER” shows the difference in mean scores between the group with comorbid dementia and cancer (COMORBID) and the group with cancer only (CANCER). Positive values indicate that the former group has the higher mean score, whereas negative values indicate that the latter group has the higher mean score.

Depression refers to differences in the sum score for the Cornell Scale for Depression in Dementia. Level of function (daily activities) refers to differences in the sum score for the Physical Self-Maintenance Scale. Neuropsychiatric symptoms refer to differences in the product score for the Neuropsychiatric Inventory-Nursing Home Edition. Antipsychotics, % 9. Opioids, % 11.3. Peripheral analgesics, % 11.4

Comorbid Dementia and Cancer

To our knowledge, few studies have investigated the comorbidity of dementia and cancer in NH patients. The study by Monroe et al. indicated that patients with severe Alzheimer’s disease and cancer have fewer pain behaviors and found severe Alzheimer’s disease to be negatively associated with total opioid medication.
Another study by Monroe et al.28 found an association between hospice enrolment and more opioid pain treatment among patients with dementia and terminal cancer. However, no patients with severe dementia and cancer were admitted to hospice.28 The study highlights that patients with severe dementia and cancer are at a great risk of having untreated advanced cancer pain because they are no longer able to account for their suffering.

The results of this study indicated that COMORBID patients have increased prevalence of NPS, including sleep disturbances, and reduced level of function with regard to activities of daily living compared with NEITHER and CANCER patients. However, COMORBID patients received more analgesics than DEMENTIA patients but displayed significantly more agitation than did CANCER and NEITHER patients. This could indicate that COMORBID patients experienced more pain compared with CANCER and NEITHER patients. However, there is no significant difference in NPS between COMORBID and DEMENTIA patients, and we do not have a measure of pain in our study. Thus, these findings should be interpreted with caution because it could be that the dementia is the source of the association.

Today, approximately 5550 patients with comorbid dementia and cancer are living in Norway, and the number is expected to increase rapidly. The results of our study provide important information for clinicians in their decision-making process regarding treatment and care for these individuals. Our results may support previous studies that showed that untreated pain is still common among NH patients with cancer.45-47 To meet these challenges, pain management methods of high quality are necessary and should be used. In light of this, it is alarming that recent findings show that, even when pain is identified through proper methods, people with cognitive impairment still do not receive pain treatment.47

Previous studies indicate that approximately 24% of people with cancer die in an NH.11 From the perspective of symptom science, the treatment for such patients should be informed by the patient’s symptom experience, relevant management strategies thereof, and measurable outcomes of the treatments.15 To optimize care for dying adults in the last days of life, the National Institute for Health and Care Excellence recently published guidelines that give an evidence-based approach for all healthcare professionals involved in the care of a person who is nearing death.48 Recommendations highlight the need for advanced care planning and assessment and treatment of pain and distressing symptoms. It is noteworthy that the care and treatment for dying people with dementia are not yet mentioned in this important document.

Interestingly, we did not find differences in NPS between COMORBID and DEMENTIA patients. Patients with comorbid dementia and cancer did however receive more analgesics than DEMENTIA patients, and the lack of differences in NPS could thus indicate that this treatment is somewhat effective in the COMORBID group. However, COMORBID patients expressed more potentially pain-related behavior such as agitation and sleep disturbance compared with CANCER patients. This is of key importance because it highlights the particularly vulnerable position of patients with comorbid dementia and cancer.

Our study reveals a cancer prevalence of 7.3% among NH patients, which is somewhat lower than previous estimates of 14% to 26%.11 Results are closer to reports from the United States, wherein studies revealed cancer prevalence among NH patients varying from 4% to 14%.11 It is beyond the scope of this study to explore the cause of the difference in cancer prevalence between people with and without dementia. However, previous research indicated that frailty status is associated with decreased cancer incidence.49 Another interesting aspect is that cancer and cancer treatment on its own can lead to cognitive impairment, which highlights the need for more knowledge and guidelines about the particularities of this group and how it should be treated.50

### Table 3: Multiple Logistic Regression Predicting the Likelihood of Using Analgesics

| OR     | SE     | P     |
|--------|--------|-------|
| Cancer | 1.974  | 0.219 | .002* |
| Dementia | 1.000  | 0.158 | .998  |
| Comorbid | 1.001  | 0.034 | .973  |
| General health | 1.206  | 0.127 | .141  |
| Level of function (daily activities) | 1.065  | 0.012 | .000* |
| Depression | 1.035  | 0.016 | .000* |
| Neuropsychiatric symptoms | 1.004  | 0.003 | .352  |
| Regular drugs | 1.261  | 0.021 | .000* |
| Use of sedatives | 0.776  | 0.124 | .41b  |
| Age, y | 1.047  | 0.007 | .000* |
| Gender, male | 0.624  | 0.120 | .000* |

The table shows a multiple logistic regression predicting the likelihood of using analgesics. The binary variable analgesics (yes/no) is the dependent variable. The independent variables are listed in the first column. The model was statistically significant: $\chi^2(11, N = 1825) = 300.4, P < .001$, Nagelkerke $R^2 = 21.9\%$. Significant $P$ values are shown in boldface, with $P < .01$ indicated by the footnote letter a and $P < .05$ with the footnote letter b.

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2. Cancer Nursing®
Furthermore, COMORBID patients had more sleep disturbances than did CANCER and NEITHER patients. This indicates that dementia is also the source of this association and implies that having comorbid dementia and cancer potentially can give sleep-related symptoms that are more burdensome. Our results also show a negative association between the use of sedatives and the use of analgesics, indicating that, if patients receive sedatives, they are less likely to be treated with analgesics. This is of interest because recent research indicates that intense anxiety and agitation are associated with sedative-hypnotic use. Because agitation may be an expression of untreated pain among people with dementia, our results may indicate that some people with comorbid dementia and cancer are given sedatives instead of pain treatment.

Limitations

The main limitation of this study is the lack of a validated pain assessment instrument, which could have been used before and after pain treatment was initiated. Thus, we are not able to demonstrate the efficacy of analgesic treatment on pain intensity or to analyze how pain and symptom management affected related symptoms. For this reason, further research should be initiated, or alternatively, longitudinal data should be collected that allow for research on the relationship between treatment and patient characteristics. Another challenge is that there is potential for systematic differences between patients declining inclusion and the final sample, but our data do not allow us to investigate this. There is also a significant difference in sample sizes between the 4 patient groups. A third limitation is that the patients do not have a confirmed diagnosis of dementia. However, good validity and reliability of the CDR scale have been reported, and it therefore serves as a good substitute for a diagnosis. Our data do not reveal the cancer stage and actual cancer treatment and do not distinguish between current and historical diagnoses of cancer, that is, patients who already have received successful treatment and it therefore serves as a good substitute for a diagnosis. Our study reveals relatively low cancer prevalence in our sample, yet significantly higher prevalence of cancer among patients with dementia than among those without. Although we find higher levels of NPS among COMORBID patients than among CANCER and NEITHER patients, there are no significant differences in NPS between COMORBID and DEMENTIA patients. Similarly, we find significantly higher use of analgesics for COMORBID patients than for DEMENTIA and NEITHER patients but no significant difference between COMORBID and CANCER patients.

The fact that COMORBID patients receive more analgesics than DEMENTIA patients, but still display more agitation and sleep disturbances than CANCER and NEITHER patients, raises the question of whether they receive adequate treatment. The lack of pain assessment in our study renders us unable to provide further evidence on this relationship.

Because COMORBID patients are often no longer able to give a valid self-report, a prerequisite for adequate treatment, our results suggest that healthcare professionals working at NHs should pay close attention to this patient group. Furthermore, our results highlight the importance of identifying patients at risk for developing sleep disturbances and other types of NPS. This could include screening of NPS using validated tools, in addition to the active use of pain assessment tools to optimize treatment.

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Conclusions and Implications for Practice

Our study reveals relatively low cancer prevalence in our sample, yet significantly higher prevalence of cancer among patients with dementia than among those without. Although we find higher levels of NPS among COMORBID patients than among CANCER and NEITHER patients, there are no significant differences in NPS between COMORBID and DEMENTIA patients. Similarly, we find significantly higher use of analgesics for COMORBID patients than for DEMENTIA and NEITHER patients but no significant difference between COMORBID and CANCER patients.

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