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Shift work is risk factor for breast cancer among Taiwanese womenC.H. Chu¹, C.J. Chen², G.C. Hsu¹, I.L. Liu³, D.C. Christiani⁴, C.H. Ku³.¹National Defense Medical Center Tri-Service General Hospital, Women's Health Center, Taipei City, Taiwan; ²The Council of Labor Affairs, Institute of Occupational Safety and Health, Taipei County, Taiwan; ³National Defense Medical Center, School of Public Health, Taipei City, Taiwan; ⁴Harvard School of Public Health, Department of Environmental Health Occupational Health Program, Boston, USA**Background:** To assess risk factors for breast cancer among Taiwanese women, we conducted a nested case-control study within a women's health center cohort in a medical center in Taipei city, Taiwan.**Materials and Methods:** All women who came to the women's health center for breast cancer screening or for diagnosis were recruited into the study. Cases were diagnosed as primary breast cancer with pathology confirmation, whereas controls were women who were never diagnosed with breast or other cancers. Informed consent was obtained from each study subject and the study approved by the institutional review board (IRB) of the Tri-Service General Hospital. Risk factors were collected by using a structure questionnaire. Conditional logistic regression was used to assess the hazard ratio (HR) with the adjustment of potential confounders (SAS 9.2 v).**Results:** A total of 2023 study subjects were recruited in the study, including 408 cancer cases and 1615 controls. After adjusting for potential confounders, we found that age (HR = 9.71, 95%CL=7.34–12.84), BMI (HR = 1.09, 95%CL=1.04–1.15), menarche age (HR = 1.16, 95%CL=1.04–1.29), menopause status (yes vs. no, HR = 2.49, 95%CL= 1.51–4.08), shift work (yes vs. no, HR = 2.54, 95%CL=1.37–4.70) were positively associated with breast cancer.**Conclusion:** Among these risk factors, work shift is an occupational exposure. We suggest re-scheduling the timetable for women in workforces with rotating shifts; as well further study on mechanism (e.g. melatonin and other hormone metabolism).

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Parity and age at first childbirth in relation to the risk of different breast cancer subgroupsS. Butt¹, S. Borgquist², L. Anagnostaki³, G. Landberg³, J. Manjer¹.¹Institution for clinical sciences, Dept of Surgery, Malmö, Sweden;²Division of Oncology, Dept of Clinical Sciences, Lund, Sweden; ³Centre for Molecular Pathology, Dept of Laboratory Medicine, Malmö, Sweden

The aim of the present study was to examine parity and age at first childbirth, in relation to the risk of specific breast cancer subgroups. A prospective cohort, The Malmö Diet and Cancer Study, including 17,035 women were followed with linkage to Swedish Cancer Registry until December 31, 2004. A total of 622 incident breast cancers were diagnosed during follow-up and were evaluated regarding invasiveness, tumour size, axillary lymph node status, Nottingham grade, tumour proliferation (Ki67), HER2, cyclin D1 and p27. The tumours were also examined for WHO type and hormone receptor status. Nulliparity was associated with an overall increased risk of breast cancer, although not statistically significant (the relative risk was 1.39 with a 95% confidence interval of 0.92–2.08). Nulliparity was also associated with large tumours (>20 mm) (1.89: 0.91–3.91), high Ki67 levels (1.95: 0.93–4.10), high cyclin D1 levels (2.15: 0.88–5.27), grade III (2.93: 1.29–6.64) and HER2 positive tumours (3.24: 1.02–10.25). High parity was not statistically significantly associated with any specific breast cancer subgroup. Older age at first childbirth (>30) was associated with a slightly increased risk of breast cancer (1.39: 0.94–2.07). There was a statistically significant association between late first childbirth and lobular type (2.51: 1.01–6.28), grade III tumours (2.67: 1.19–6.02), high levels of cyclin D1 (2.69: 1.18–6.12) and low levels of p27 (2.23: 1.15–4.35). We conclude that nulliparity and late first childbirth are associated with relatively more aggressive breast cancer subgroups.

References

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Measuring quality of the process of care in breast-cancer patients: a French perspectiveM. Couralet¹, M. Ferrua¹, S. Morin¹, C. Grenier², C. Gardel³, M.H. Rodde-Dunet⁴, E. Minvielle¹. ¹Projet Compaqh, Inserm U750, Villejuif cedex, France; ²Fnlcc, Qualité/Indicateurs, Paris, France; ³Has, Ipaqss, St Denis La Plaine, France; ⁴Inca, Direction de la Qualité des Soins, Boulogne Billancourt, France**Background:** The objective of the study is to develop a set of quality indicators (QIs), allowing hospitals comparison, measuring the process of care in non-inflammatory non-metastatic invasive breast cancer patients.**Methods:** The COMPAQH project has designed a set of QIs, derived from clinical practice guidelines, in partnership with national agencies, hospitals federations and professional bodies (SFSPM, CNGOF, INCa, FNCLCC, HAS)^a.

Eight QIs evaluating delays as well as different steps of the process of care have been defined (table). QIs are evaluated through 3 criteria: feasibility of data collection, metrological quality (reliability, validity) and relevance (discriminative power). For each indicator, data collection is based on a retrospective analysis of 80 randomly selected medical records in each hospital.

QIs are measured in a panel of 70 volunteer hospitals. As data collection is in progress, preliminary results presented here concern 41 hospitals.

Results: QI1 is the delay between patient call for a surgeon consultation and the date of consultation. However, patients' calls are scarcely recorded (<50% of hospitals). This QI cannot therefore allow hospitals comparison: median and extreme values are delivered to hospitals only.

Measures used: Proportion of patients . . .		% , mean (min, max)
QI2	with a delay from first surgeon consultation to first surgery ≤21 days	58 (17, 90)
QI3	with a delay from MRM (multidisciplinary review meeting) to the post-surgery consultation presenting its conclusions ≤14 days	83 (26, 100)
QI4	with a delay from first surgery to first adjuvant treatment ≤30 days if chemotherapy or ≤56 days if radiotherapy	50 (17, 92)
QI5.1	whose case is submitted to a well organized MRM	50 (0, 100)
QI5.2	with a delay from first surgery to MRM ≤14 days	65 (1, 99)
QI6	who receive a complete information before surgery	13 (0, 100)
QI7	where mandatory prognostic factors are specified in medical records	71 (4, 99)

Conclusions: Taking in charge patients without delay has consequences on 5 years survival. The working group decided therefore to maintain the collection of QI1. It should encourage hospitals to improve their information systems.

Large variations in practice are observed on the 7 other QIs. Rather low score values should encourage hospitals to promote quality improvement policies. The very low value of QI6 may be due to the fact that recommendations by health authorities on this topic were made recently.

If these results are confirmed in the whole set of 70 hospitals and after discussion with the working group, a national implementation in every hospital managing breast-cancer patients should occur in 2010.

^aSFSPM, French Senologic Society; CNGOF, French National College of Gynaecologists and Obstetricians; INCa, French National Cancer Institute; FNCLCC, French National Federation of Cancer; HAS, French National Authority for Health.

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A multidisciplinary approach to menopausal symptom management after breast cancerC. Saunders¹, L. Emery¹, J. Gregson², D. Doherty², M. Hickey².¹University of Western Australia, School of Surgery, Crawley, Australia;²University of Western Australia, School of Women's and Infants' Health, Crawley, Australia

Background: Nearly 2.5 million US women are breast cancer (BC) survivors [1]. Breast cancer treatment causes menopausal symptoms in around 60% of patients. While ovarian suppression, endocrine treatments and cessation of HRT all commonly cause menopausal symptoms, few studies have evaluated the severity of symptoms or their impact on quality of life and there is limited information for health care providers or patients about management strategies. While established treatments for menopausal symptoms (estrogen/progestin and tibolone) may increase the risk of breast cancer recurrence [2], relatively little is known about the safety of these treatments after other hormone-dependent cancers [3]. Effective non-hormonal treatments are increasingly available, but there is often poor coordination between health providers in advising and managing menopause after cancer. The multidisciplinary (MD) model of cancer care is now well established and offers many advantages to consumers and health care providers [4–9].

Materials and Methods: A public MD clinic for menopausal symptoms after cancer (MSAC) has been established in Western Australia where

patients are seen by a gynaecologist and management issues referred to the MD team (gynaecologists, BC surgeons, oncologists, endocrinologists, psychiatrist, clinical psychologist and family practitioners) who meet monthly to develop menopause management plan. Data are collected using standardised techniques on menopause symptoms, sexual activity, functional assessment of cancer therapy, and quality of life (QOL).

Results: 720 women were referred to the MSAC clinic (January 2003–December 2008), with 81% BC patients. The most common 'severe' symptoms included hot flashes (40%), night sweats (35%) and loss of interest in sex (30%). Nearly half (42%) of all patients were premenopausal at diagnosis, 37% were previous HRT users and 22% were taking HRT at the time of BC diagnosis.

A written and web based information resource on menopausal symptoms after breast cancer has been developed to assist women with quality of life issues. The resource was highly rated by patients and is now nationally distributed in Australia through the National Breast and Ovarian Cancer Foundation [10].

Conclusions: Menopausal symptoms are a common and distressing symptom of breast cancer treatment. MD care and targeted information resources help address the gap in clinical services for these women.

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Diurnal cortisol rhythm as a predictor of breast cancer survival

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Background: Abnormal circadian rhythms have been observed in patients with cancer, but the prognostic value of such alterations has not been confirmed. We examined the association between diurnal variation of salivary cortisol in patients with metastatic breast cancer and subsequent survival. We explored relationships between cortisol rhythms, circulating natural killer (NK) cell counts and activity, prognostic indicators, medical treatment, and psychosocial variables.

Methods: Salivary cortisol levels of 104 patients with metastatic breast cancer were assessed at study entry at 0800, 1200, 1700, and 2100 hours on each of 3 consecutive days, and the slope of diurnal cortisol variation was calculated using a regression of log-transformed cortisol concentrations on sample collection time. NK cell numbers were measured by flow cytometry, and NK cell activity was measured by the chromium release assay. The survival analysis was conducted by the Cox proportional hazards regression model with two-sided statistical testing.

Results: Cortisol slope predicted subsequent survival up to 7 years later. Earlier mortality occurred among patients with relatively "flat" rhythms, indicating a lack of normal diurnal variation (Cox proportional hazards, $P=0.0036$). Patients with chest metastases, as opposed to those with visceral or bone metastases, had more rhythmic cortisol profiles. Flattened profiles were linked with low counts and suppressed activity of NK cells. After adjustment for each of these and other factors, the cortisol slope remained a statistically significant, independent predictor of survival time. NK cell count emerged as a secondary predictor of survival.

Conclusions: Patients with metastatic breast cancer whose diurnal cortisol rhythms were flattened or abnormal had earlier mortality. Suppression of NK cell count and NK function may be a mediator or a marker of more rapid disease progression.

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Cognitive behavioral therapy and physical exercise for climacteric symptoms in breast cancer patients experiencing treatment-induced menopause

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Background: Premature menopause is a major concern of younger women undergoing adjuvant therapy for breast cancer. Hormone replace-

ment therapy is contraindicated; non-hormonal medications show side-effects. There is growing evidence that cognitive behavioral therapy and physical exercise can have a positive impact on symptoms in naturally occurring menopause. The purpose of this clinical trial is to evaluate the efficacy of cognitive behavioural therapy (CBT) (A), physical exercise (B), or the combination of these interventions (AB) in alleviating menopausal symptoms, improving sexual functioning and enhancing quality of life.

Material and Methods: In this multicenter study, a minimum of 325 eligible women are being randomised to group A, B, AB or a 'waiting list' control group. For group A, the intervention consists of 6 weekly group CBT sessions of 1.5 hours and a booster session. For group B, the intervention is an individually tailored, 12 week home-based physical exercise program of 2.5–3 hours per week. Group AB receives both the CBT and exercise program. Questionnaires assessing menopausal symptoms, sexuality, body- and self-image, psychological distress and quality of life are being completed at baseline, at 12 weeks and at 6 months follow-up.

Results: As of November 2009, 2688 women have been identified as being potentially eligible for study participation, of whom 1514 completed a screening questionnaire and 662 a postcard indicating they had no interest in the study. 627 of the screened women met eligibility criteria and received a baseline questionnaire. To date, 422 women have returned this questionnaire and have been randomly allocated to the CBT group, ($n=109$), the physical exercise group ($n=104$), the combined intervention group ($n=106$), or the control group ($n=103$). Problems have been experienced in retaining patients in the trial, with dropouts occurring primarily between randomization and start of the intervention. The majority of those who actually participate in the interventions are able to comply with the program. Data collection will continue until mid-2010.

Conclusions: Cognitive behavioral therapy and physical exercise are potentially useful treatments for women with breast cancer who experience treatment-induced, premature menopause. In this conference, the content of the interventions, flow of participants throughout the trial, reasons for dropout and initial experiences with the interventions will be presented.

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Three year experience of a breast cancer family history clinic in a district general hospital in the United Kingdom

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Background: The National Institute of Clinical excellence (NICE) in the United Kingdom issued guidance for the management of women with a family history of breast cancer in 2004 and these were updated in 2006. This led to the setting up of a dedicated family history clinic in this District General Hospital. The three year experience from this risk assessment clinic is presented here.

Methods: The Family History Clinic was set with a research grant. Asymptomatic women with a family history of breast cancer and women with a family history of breast cancer seen in the symptomatic breast clinics were referred into this clinic. All women completed a detailed family history questionnaire and a family pedigree chart was created with the gathered information and using the Progeny software. Using the NICE guidelines, these women were then grouped into near population risk, raised risk or high risk groups and were managed accordingly.

Results: Between Sept 2006 and Aug 2009, 962 women were seen in the clinic. Of these, 244 (25.4%) women were identified to be at near population risk, 327 (34%) raised and 391 (40.6%) were at high risk. All near population risk women and raised risk women over the age of 50 years (298, 31%) were discharged from this clinic. Women between the ages of 40–49 (189; 19.6%) and at raised risk were advised yearly surveillance. Of the 391 high risk women, 360 women consented to be referred to the Regional Genetics Service. Ten women have been identified with BRCA 1 or 2 mutations from this group. In addition, women were offered the opportunity to enter the IBIS II prevention and UKFOCSS Clinical Trials. Eight women entered the IBIS II prevention trial and three women entered the UKFOCSS trial.

Conclusions:

- The Family History Clinic provides a comprehensive service to women with familial breast cancer by one stop onsite consultation, clinical and radiological assessment and appropriate counselling.
- From this study, we have shown a high need for women to be seen in a dedicated clinic (i.e., 962 women).
- There are a group of women clearly at high risk who do require genetic counselling and testing.
- It includes a small group (ten) of women that have shown genetic mutation and now can be identified and managed correctly which includes offering risk reducing prophylactic surgery.