

Survival of Australian women with invasive epithelial ovarian cancer: a population-based study

Satyamurthy Anuradha
MBBS, FRCS, PhD
Research Officer¹

Penelope M Webb
MA, DPhil
Head¹

Penny Blomfield
MBBS, FRANZCOG, CGO
Clinical Associate Professor²

Alison H Brand
MD, FRANZCOG, CGO
Director³

Michael Friedlander
FRACP
Conjoint Professor of
Medicine⁴

Yee Leung
MBBS, FRANZCOG, CGO
Professor⁵

Andreas Obermair
MD, FRANZCOG, CGO
Professor⁶

Martin K Oehler
MD, FRANZCOG, CGO
Director⁷

Michael Quinn
MB ChB, FRANZCOG, CGO
Professor⁸

Christopher Steer
MBBS, FRACP
Medical Oncologist⁹

Susan J Jordan
MBBS, FRACGP, PhD
Senior Research Officer¹

¹Gynaecological Cancers Group, QIMR Berghofer Medical Research Institute, Brisbane, QLD.

²Gynaecologic Oncology, Royal Hobart Hospital, Hobart, TAS.

³Department of Gynaecological Oncology, Westmead Hospital, University of Sydney, Sydney, NSW.

⁴Department of Medical Oncology, Prince of Wales Hospital, Sydney, NSW.

⁵School of Women's and Infants' Health, University of Western Australia, Perth, WA.

⁶Department of Gynaecological Oncology, Royal Brisbane and Women's Hospital, Brisbane, QLD.

⁷Department of Gynaecological Oncology, Royal Adelaide Hospital, Adelaide, SA.

Ovarian cancer is the seventh most common cause of cancer mortality in Australian women, accounting for an estimated 1080 deaths in 2013,¹ but there are no comprehensive national data describing clinical and non-clinical factors associated with survival in Australian women. Available information is derived from cancer registries,^{1,2} which hold little or no information on important prognostic factors like stage of disease, or is state-specific and not contemporary.^{2,3} Furthermore, estimates of conditional survival, which take into account time already survived and are of practical value to clinicians and patients,⁴ are not available in the Australian context.

We used data from a complete national cohort of women diagnosed with invasive epithelial ovarian cancer in Australia in 2005 to describe survival patterns by sociodemographic, clinical and histopathological features, and to calculate conditional survival estimates.

Methods

In 2009, the Australian state- and territory-based cancer registries identified all women aged ≥ 18 years who had been diagnosed with primary invasive epithelial ovarian cancer (including fallopian tube and primary peritoneal cancers) in Australia during 2005. We obtained de-identified clinical information from all the women's medical records. Cancer registries also provided women's ages and postcodes at diagnosis and, where available, the grade and histological subtype of their cancers.⁵

The study was approved by the human research ethics committees of the QIMR Berghofer Medical Research Institute and all participating institutions.

Participant characteristics

Postcode was used to classify women's area of residence using the

Abstract

Objective: To describe survival patterns in a nationally complete cohort of Australian women with epithelial ovarian cancer, by sociodemographic and clinical factors.

Design, setting and participants: All 1192 women diagnosed with invasive epithelial ovarian cancer in 2005 were identified through state-based cancer registries. We obtained detailed information from their medical records in 2009 and updated survival data in 2012.

Main outcome measures: Crude 3-year, 5-year and 7-year survival rates; 3-year and 5-year conditional survival; and hazard ratios (HRs) for the association of participant and cancer characteristics with survival, from multivariable Cox proportional hazards models.

Results: Overall crude 5-year survival was 35% (95% CI, 33%–38%). Conditional survival increased moderately for women who lived beyond a year from diagnosis, although for women alive 2 years after diagnosis, the probability of surviving a further 5 years was still only 53% (95% CI, 49%–57%). Increasing age and disease stage were most strongly associated with poor survival. After adjusting for these, survival was significantly worse for women with carcinosarcomas (HR_{adj}, 2.1 [95% CI, 1.3–3.2]), clear cell (HR_{adj}, 1.7 [95% CI, 1.2–2.3]) and mucinous (HR_{adj}, 2.6 [95% CI, 1.6–4.0]) cancers than for women with serous cancers. Presence of ascites at diagnosis (HR_{adj}, 1.5 [95% CI, 1.3–1.8]), Charlson comorbidity score ≥ 3 (HR_{adj}, 1.5 [95% CI, 1.1–2.1]), relative socioeconomic disadvantage (HR_{adj}, 1.2 [95% CI, 1.1–1.4]) and regional-remote residence (HR_{adj}, 1.2 [95% CI, 1.0–1.4]) were also associated with poorer survival.

Conclusions: Along with expected adverse effects of age and stage, we found survival differences by histological subtype, presence of ascites and comorbidities. Whether geographic and socioeconomic differences relate to treatment access or other factors warrants further investigation. Conditional survival estimates confirm the ongoing poor long-term prognosis for women with ovarian cancer, reinforcing the need for prevention and better treatments.

Accessibility/Remoteness Index of Australia (ARIA+).⁶ We further categorised women as being from major cities or regional-remote areas. We also used postcode as a measure of socioeconomic status using the Index of Relative Socio-Economic Advantage/Disadvantage, one of the Socio-Economic Indexes for Areas (SEIFA),⁷ which is based on indicators including income and skilled employment. We categorised women as relatively disadvantaged or advantaged by dividing their SEIFA scores at the median. Medical record information was used to derive comorbidity scores based on the Charlson comorbidity index,⁸ with a higher score indicating a larger number or greater severity of comorbidities. Scores were grouped for analysis (0, 1, 2 or ≥ 3).

Cancer characteristics

Histological subtype was classified as serous, mucinous, endometrioid, clear cell or carcinosarcoma; and tumour grade as well differentiated, moderately differentiated or poorly differentiated. Information on stage, according to the International Federation of Gynecology and Obstetrics (FIGO) 1988 system, was abstracted from clinical notes. Women for whom stage information was missing were classified as having advanced disease (called stage III/IV) if they met any of the following criteria: macroscopic residual disease; poorly differentiated cancer with serous subtype; or poorly differentiated cancer without surgical intervention. Women missing data for other cancer characteristics were included in a separate category in our analyses. However, inclusion of women missing subtype data resulted in violation

⁸ Department of Obstetrics and Gynaecology, University of Melbourne, Melbourne, VIC.

⁹ Border Medical Oncology, Wodonga, NSW.

Satyamurthy.Anuradha@qimrberghofer.edu.au

MJA 2014; 201: 283-288
doi: 10.5694/mja14.00132

of proportional hazards assumptions and thus we excluded this group from subtype analyses.

Survival data

As specific cause-of-death information was not available for all women, the outcome was death from any cause. However, among women with this information, over 95% of deaths were from ovarian cancer. We obtained death information from medical records in 2009, and updated data were obtained from the cancer registries in 2012. The New South Wales cancer registry was unable to provide the updated information. However, a third of the women in our study also participated in the Australian Ovarian Cancer Study,⁹ and death information for these women was available until October 2011 through linkage to the National Death Index. Thus, we still had complete 5-year survival information for 74% of women from NSW. Participants not known to have died were censored at the last date of data linkage. Survival time was calculated from date of diagnosis until death or censoring.

Statistical analysis

We estimated crude 3-year, 5-year and 7-year survival rates with 95% confidence intervals (CIs) using the Kaplan–Meier method. If $S(t)$ is taken as the traditional survival at time t , then CS, conditional survival, is the probability of surviving an additional y years, given that an individual has already survived x years, and is expressed as: $CS(y|x) = S(x+y)/S(x)$.⁴ We calculated 3-year and 5-year conditional survival probabilities for women who had already survived 1 and 2 years; and, additionally, 3-year conditional survival for those who had survived 3 years. We calculated hazard ratios (HRs) and 95% CIs for the association of participant and cancer characteristics with survival using multivariable Cox proportional hazards models. Proportional hazards assumptions were tested using scaled Schoenfeld residuals. Analyses were conducted using Stata, version 11.2 (Stata Corporation).

Results

During 2005, 1192 women were registered with a diagnosis of invasive epithelial ovarian, fallopian tube or primary peritoneal cancer. The women's mean age was 65 years (SD, 13 years), and 71% were from major cities. Most (69%) had stage III or IV cancer, and 43% had ascites. More than half the cancers (53%) were serous subtype, and 61% were poorly differentiated (Box 1).

Stage information was not available for 283 women, most of whom did not have surgery. We classified 149 of these women (13%) as having stage III/IV disease. The proportion of data missing for other cancer characteristics ranged from 16% to 24%.

Box 2 shows Kaplan–Meier survival curves for the women by age, FIGO stage, socioeconomic status and area of residence. Box 1 shows 3-year, 5-year and 7-year survival and HRs by participant and cancer characteristics. Advanced age was associated with worse survival. This was mostly explained by stage and comorbidity score but, even after adjustment for these, age remained significantly associated with survival (HR_{adj} 1.3 [95% CI, 1.2–1.4] for each 10-year increment).

The rate of death of women with stage IV disease was 10 times that of women with stage I disease (HR_{adj} 10.2 [95% CI, 6.5–16.1]). After adjusting for other variables, especially stage, the effect estimates associated with grade attenuated and were no longer statistically significant (HR_{adj} 1.5 [95% CI, 0.9–2.5] for moderately differentiated and HR_{adj} 1.4 [95% CI, 0.9–2.4] for poorly differentiated v well differentiated cancers). We explored this relationship further by stage. Among women with stage I or II disease, the HRs associated with grade were 3.2 (95% CI, 0.9–11.3) for poorly differentiated and 1.7 (95% CI, 0.4–6.6) for moderately differentiated cancers; while among women with advanced disease, the HRs were 1.6 (95% CI, 0.9–2.7) and 1.5 (95% CI, 0.9–2.7) for poorly and moderately differentiated cancers, respectively. The interaction term for stage and grade was not, however, statistically significant.

In age-adjusted analyses, mucinous, endometrioid and clear cell subtypes were associated with better survival than serous cancers. However, after adjusting for other variables, especially stage and grade, survival was worse among women with mucinous cancers (HR_{adj} 2.6 [95% CI, 1.6–4.0]) and clear cell cancers (HR_{adj} 1.7 [95% CI, 1.2–2.3]). Carcinosarcomas were associated with poorer survival irrespective of adjustment (HR_{adj} 2.1 [95% CI, 1.3–3.2] v serous cancers). Presence of ascites at diagnosis was also related to poorer survival (HR_{adj} 1.5 [95% CI, 1.3–1.8]).

Higher comorbidity scores were modestly associated with survival, as was socioeconomic status (Box 1). Compared with women from relatively socioeconomically advantaged areas, women from relatively disadvantaged areas had a 21% higher risk of dying during follow-up. A similar difference was seen for women living in regional–remote areas versus major cities. There were no significant survival differences by state after adjustment.

Box 3 shows conditional survival for women who survived up to at least 3 years after diagnosis. Three-year conditional survival was 47% at diagnosis; however, for those who did survive 3 years from diagnosis, the probability of surviving a further 3 years increased to 68% (95% CI, 64%–72%). The 5-year conditional survival rates increased from 35% at diagnosis to 53% (95% CI, 49%–57%) for women who survived 2 years. In general, the greatest increases in conditional survival were seen in groups with the lowest survival probabilities at diagnosis.

Discussion

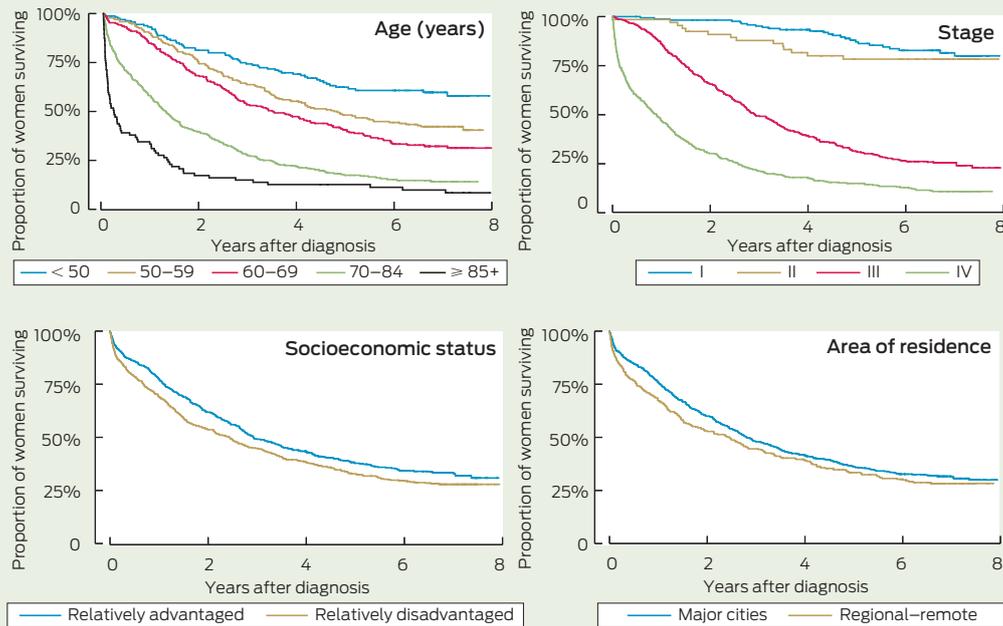
In this national cohort of women with epithelial ovarian cancer, increasing age and disease stage were the factors most strongly associated with poorer survival. In addition, we found significant independent associations with ascites, comorbidities, and regional–remote and relatively disadvantaged areas of residence. Overall survival was poor, with just over a third of women (35%) surviving 5 years from diagnosis. For women who survived

1 Numbers and proportions of women diagnosed with invasive epithelial ovarian cancer in Australia in 2005, and crude 3-year, 5-year and 7-year survival estimates and hazard ratios (HRs), by participant and cancer characteristics

Variable	No. (%) [*]	3-year survival (95% CI)	5-year survival (95% CI)	7-year survival (95% CI)	Unadjusted HR (95% CI)	Age-adjusted HR (95% CI)	Fully adjusted HR (95% CI) [†]
Total	1192	47% (44%–50%)	35% (33%–38%)	31% (28%–33%)			
Age (years)							
< 50	166 (14%)	74% (67%–80%)	62% (54%–70%)	60% (51%–67%)	Ref	Ref	Ref
50–59	267 (22%)	64% (58%–69%)	48% (42%–54%)	42% (36%–48%)	1.6 (1.2–2.2)	1.6 (1.2–2.2)	1.5 (1.1–2.0)
60–69	278 (23%)	53% (47%–59%)	40% (34%–46%)	32% (26%–38%)	2.1 (1.5–2.8)	2.1 (1.5–2.8)	1.5 (1.1–2.1)
70–84	394 (33%)	27% (23%–32%)	17% (14%–22%)	14% (10%–18%)	4.2 (3.2–5.6)	4.2 (3.2–5.6)	2.6 (2.0–3.4)
≥ 85	87 (7%)	15% (8%–23%)	13% (7%–21%)	10% (5%–17%)	7.6 (5.4–10.6)	7.6 (5.4–10.6)	3.5 (2.4–4.9)
Stage							
I	171 (14%)	95% (91%–98%)	87% (80%–91%)	82% (74%–87%)	Ref	Ref	Ref
II	67 (6%)	88% (77%–94%)	78% (66%–87%)	78% (66%–87%)	1.4 (0.7–2.6)	1.4 (0.8–2.7)	1.5 (0.8–2.9)
III	489 (41%)	49% (45%–54%)	31% (27%–36%)	26% (22%–30%)	7.0 (4.6–10.1)	6.3 (4.2–9.4)	6.7 (4.3–10.2)
IV	182 (15%)	20% (15%–27%)	13% (8%–19%)	9% (5%–14%)	15.1 (9.7–22.0)	12.9 (8.5–19.5)	10.2 (6.5–16.1)
III/IV [‡]	149 (13%)	19% (13%–25%)	14% (9%–21%)	9% (4%–15%)	16.6 (7.8–19.4)	12.3 (8.0–18.8)	12.0 (7.6–19.1)
Missing	134 (11%)	25% (20%–32%)	18% (12%–25%)	15% (9%–22%)	15.2 (11.2–25.3)	10.2 (6.6–15.7)	6.0 (3.8–9.6)
Grade							
Well differentiated	78 (6%)	83% (73%–90%)	78% (66%–86%)	75% (62–83%)	Ref	Ref	Ref
Moderately differentiated	199 (17%)	60% (53%–66%)	44% (37%–51%)	38% (31%–45%)	3.2 (1.9–5.2)	2.7 (1.6–4.2)	1.5 (0.9–2.5)
Poorly differentiated	723 (61%)	49% (45%–53%)	36% (32%–40%)	30% (27%–34%)	4.1 (2.6–6.6)	3.2 (2.0–5.1)	1.4 (0.9–2.4)
Missing	192 (16%)	14% (9%–19%)	8% (5%–12%)	6% (3%–10%)	14.5 (8.9–23.6)	8.4 (5.1–13.8)	2.0 (1.2–3.6)
Subtype							
Serous	631 (53%)	49% (45%–53%)	34% (30%–38%)	28% (24%–32%)	Ref	Ref	Ref
Mucinous	69 (6%)	58% (45%–69%)	53% (41%–64%)	48% (35%–59%)	0.7 (0.5–1.0)	0.9 (0.6–1.2)	2.6 (1.6–4.0)
Endometrioid	98 (8%)	84% (75%–90%)	75% (65%–82%)	68% (57%–77%)	0.3 (0.2–0.4)	0.3 (0.2–0.5)	0.7 (0.5–1.1)
Clear cell	101 (8%)	68% (58%–76%)	55% (44%–64%)	51% (40%–61%)	0.6 (0.4–0.8)	0.6 (0.5–0.9)	1.7 (1.2–2.3)
Carcinosarcoma	27 (2%)	27% (12%–45%)	16% (5%–32%)	16% (5%–32%)	1.6 (1.1–2.5)	1.4 (0.9–2.1)	2.1 (1.3–3.2)
Other epithelial	64 (5%)	50% (37%–61%)	43% (31%–55%)	40% (27%–51%)	§	§	§
Missing	202 (17%)	11% (7%–16%)	5% (2%–8%)	3% (1%–7%)	§	§	§
Ascites							
No	391 (33%)	66% (61%–71%)	57% (51%–62%)	49% (44%–55%)	Ref	Ref	Ref
Yes	519 (43%)	41% (37%–45%)	26% (22%–30%)	22% (19%–26%)	2.2 (1.9–2.6)	2.1 (1.7–2.5)	1.5 (1.3–1.8)
Missing	282 (24%)	32% (26%–37%)	23% (18%–28%)	20% (16%–25%)	2.9 (2.4–3.5)	2.4 (2.0–2.9)	1.5 (1.2–1.9)
Comorbidity score[¶]							
0	835 (70%)	52% (48%–55%)	40% (36%–43%)	34% (31%–38%)	Ref	Ref	Ref
1	99 (8%)	42% (33%–52%)	31% (22%–40%)	29% (21%–39%)	1.3 (1.0–1.6)	1.0 (0.8–1.3)	0.9 (0.7–1.2)
2	113 (9%)	42% (33%–52%)	33% (24%–42%)	27% (19%–37%)	1.2 (1.0–1.5)	1.0 (0.8–1.2)	1.0 (0.8–1.3)
≥ 3	63 (5%)	32% (21%–43%)	19% (10%–31%)	13% (6%–25%)	1.9 (1.4–2.5)	1.4 (1.0–1.9)	1.5 (1.1–2.1)
Missing	82 (7%)	21% (13%–31%)	12% (6%–21%)	11% (5%–19%)	2.6 (2.0–3.3)	1.7 (1.3–2.2)	1.3 (1.0–1.7)
State							
Victoria	290 (24%)	43% (37%–48%)	33% (28%–39%)	29% (24%–34%)	Ref	Ref	Ref
New South Wales/ACT	454 (38%)	45% (40%–50%)	31% (26%–36%)	**	1.1 (0.9–1.3)	1.1 (0.9–1.3)	1.2 (0.9–1.5)
Queensland	207 (17%)	51% (44%–58%)	38% (32%–45%)	32% (26%–39%)	0.9 (0.7–1.1)	1.0 (0.8–1.3)	1.1 (0.9–1.4)
Western Australia	115 (10%)	57% (48%–66%)	46% (37%–55%)	38% (29%–47%)	0.7 (0.6–1.0)	0.9 (0.7–1.1)	1.0 (0.8–1.3)
South Australia/NT	101 (8%)	50% (40%–59%)	40% (31%–50%)	40% (31%–50%)	0.8 (0.6–1.0)	0.8 (0.6–1.1)	0.8 (0.6–1.3)
Tasmania ^{††}	25 (2%)	40% (21%–58%)	20% (7%–37%)	15% (4%–32%)	1.4 (0.9–2.3)	1.2 (0.8–1.9)	1.4 (0.9–2.3)
Socioeconomic status							
Relatively advantaged	617 (52%)	49% (45%–53%)	38% (34%–42%)	33% (29%–37%)	Ref	Ref	Ref
Relatively disadvantaged	575 (48%)	45% (41%–49%)	33% (29%–37%)	28% (24%–32%)	1.2 (1.0–1.4)	1.3 (1.1–1.5)	1.2 (1.1–1.4)
Area of residence							
Major cities	842 (71%)	48% (45%–51%)	36% (33%–40%)	32% (28%–35%)	Ref	Ref	Ref
Regional–remote	350 (29%)	45% (39%–50%)	33% (28%–39%)	28% (23%–33%)	1.2 (1.0–1.3)	1.2 (1.0–1.4)	1.2 (1.0–1.4)

ACT = Australian Capital Territory. NT = Northern Territory. * Percentages may not sum to 100% because of rounding. † Adjusted for age at diagnosis, comorbidity score, stage, grade, subtype and presence of ascites. Analysis by state was also adjusted for socioeconomic status and area of residence. ‡ Women with missing stage information were classified as having stage III/IV disease if they met any of the following criteria: macroscopic residual disease; poorly differentiated cancer with no surgery; poorly differentiated cancer of serous subtype. § HR could not be calculated due to violation of proportional hazards assumption. ¶ Weighted score based on the Charlson comorbidity index. ** Estimate could not be calculated due to incomplete 7-year survival data. †† Estimates are imprecise due to small number of women diagnosed with ovarian cancer in Tasmania in 2005. ◆

2 Kaplan–Meier survival curves for women diagnosed with invasive epithelial ovarian cancer in Australia in 2005, by age, FIGO stage of disease, socioeconomic status and area of residence



FIGO = International Federation of Gynecology and Obstetrics. ◆

beyond 1, 2 or 3 years, subsequent 3-year and 5-year survival increased moderately, although 5-year conditional survival at 2 years was still only 53%.

It is important to note that our survival rates are crude and, while they reflect the actual survival for this group, they do not take into account the probability of death from other causes and cannot be used for international comparisons. Contemporary routine Australian statistics suggest a *relative* 5-year survival rate of about 43%,¹ which compares favourably with estimates for similar periods from other Organisation for Economic Co-operation and Development countries (Box 4).

Our findings are comparable to international studies showing that stage is the most important predictor of survival.^{15–17} Similar to some reports,^{15,16} we found that after adjusting for stage, cancer grade had limited independent prognostic significance. In contrast, other studies^{3,17} found that higher grade remained significantly associated with survival after adjusting for stage. There can be considerable intra- and inter-observer variability in determining ovarian cancer grade,¹⁸ which might explain these conflicting results. However,

our results also suggest that the effect of grade may be modified by stage, such that higher grade is most relevant to survival in women with early-stage cancers.

Similar to others,¹⁹ we found that women with carcinosarcomas had the poorest survival, possibly reflecting aggressive tumour biology or less sensitivity to routinely used chemotherapy.²⁰ We also found differences for women with mucinous and clear cell cancers. These women mostly presented with early-stage disease (data not shown), but our results suggest that for cancers of equivalent stage, survival is worse for women with these subtypes compared with serous cancers. This may reflect the poorer response of mucinous and clear cell cancers to conventional platinum and paclitaxel-based chemotherapy^{21,22} and reinforces the need for new treatments for these cancer subtypes. Consistent with other work,²³ we also found a 50% greater risk of mortality in women with ascites at diagnosis, underscoring the importance of ongoing clinical trials aimed at improving outcomes in these patients.

Age was an important predictor of survival in our analyses, with women aged ≥ 70 years having a two- to

threefold greater risk of death during follow-up. Older women in this cohort were less likely to receive standard chemotherapy,⁵ so this and other age-related treatment variations may be affecting survival times for older women.

The association between pre-diagnosis comorbidities and survival among women with ovarian cancer is inconsistent in the literature.^{24,25} Our results suggest that women with multiple or severe comorbidities do worse. As the type and severity of comorbid diseases affect treatment decisions, some of this effect may be mediated by treatment received.

While it is recognised that cancer mortality is generally higher in socio-economically disadvantaged and regional-remote areas of Australia,^{1,26,27} previous studies have not clearly shown these associations with ovarian cancer survival.^{1,2,27} Our analyses suggest a modest effect of area of residence. We considered the possibility that this might be due to higher rates of deaths from other causes but, among the women for whom we had cause-of-death data, there was no difference in percentage of deaths due to ovarian cancer by area of residence. Possible explanations for socioeconomic and geographic differences in ovarian cancer survival include diagnostic delay and poorer access to recommended treatments.²⁷ Further research is needed to determine the relative contribution of these factors.

Strengths of our study include the large national sample, the comprehensive data collection and completeness of follow-up. To our knowledge, this is also the first study to provide national conditional survival estimates for ovarian cancer in Australia. Conditional survival information can be used to communicate prognosis to patients and their families more accurately along their disease journey. Our estimates confirm the ongoing poor prognosis of the disease and emphasise the need for primary and secondary prevention and better treatments for ovarian cancer to improve long-term outcomes.

In conclusion, this study of all women diagnosed with invasive epithelial ovarian cancer in Australia in 2005 provides population-level evidence on factors that influence

3 Conditional survival for all women who survived at least 1–3 years after diagnosis, by age and cancer characteristics

Variable	3-year conditional survival (95% CI)				5-year conditional survival (95% CI)		
	Years survived after diagnosis				Years survived after diagnosis		
	0	1	2	3	0	1	2
Overall survival	47% (44%–50%)	55% (52%–59%)	61% (57%–65%)	68% (64%–72%)	35% (33%–38%)	44% (40%–47%)	53% (49%–57%)
Age (years)							
< 50	74% (67%–80%)	75% (67%–81%)	77% (68%–83%)	82% (73%–88%)	62% (54%–70%)	66% (57%–73%)	74% (64%–81%)
50–59	64% (58%–69%)	62% (55%–68%)	65% (57%–71%)	70% (62%–76%)	48% (42%–54%)	50% (43%–56%)	57% (49%–64%)
60–69	53% (47%–59%)	56% (49%–62%)	58% (51%–65%)	63% (54%–71%)	40% (34%–46%)	39% (33%–46%)	47% (40%–55%)
≥ 70	25% (21%–29%)	38% (32%–44%)	48% (39%–55%)	58% (47%–67%)	17% (13%–20%)	28% (22%–34%)	38% (30%–47%)
Stage							
I	95% (91%–98%)	94% (90%–97%)	89% (82%–93%)	87% (79%–92%)	87% (80%–91%)	84% (76%–89%)	83% (75%–89%)
II	88% (77%–94%)	81% (69%–89%)	86% (74%–93%)	89% (77%–95%)	78% (66%–87%)	80% (67%–88%)	86% (74%–93%)
III	49% (45%–54%)	45% (41%–50%)	48% (42%–53%)	54% (47%–60%)	31% (27%–36%)	31% (26%–35%)	39% (33%–45%)
IV	20% (15%–27%)	34% (25%–44%)	42% (28%–55%)	56% (37%–72%)	13% (8%–19%)	22% (14%–32%)	30% (18%–43%)
III/IV*	19% (13%–25%)	30% (20%–41%)	46% (31%–60%)	60% (36%–77%)	14% (9%–21%)	23% (14%–35%)	28% (14%–44%)
Missing	25% (20%–32%)	52% (38%–65%)	65% (46%–79%)	65% (44%–79%)	18% (12%–25%)	40% (26%–54%)	54% (35%–69%)
Subtype							
Serous	49% (45%–53%)	49% (44%–53%)	53% (47%–58%)	61% (55%–67%)	34% (30%–38%)	36% (31%–40%)	44% (38%–49%)
Mucinous	58% (45%–69%)	81% (67%–90%)	85% (70%–93%)	85% (68%–94%)	53% (41%–64%)	71% (55%–82%)	76% (59%–87%)
Endometrioid	84% (75%–90%)	83% (74%–90%)	84% (75%–91%)	85% (75%–91%)	75% (65%–82%)	76% (65%–84%)	76% (65%–85%)
Clear cell	68% (58%–76%)	71% (60%–79%)	74% (62%–83%)	75% (61%–84%)	55% (44%–64%)	58% (46%–69%)	69% (55%–79%)
Carcinosarcoma	27% (12%–45%)	26% (10%–47%)	29% (9%–52%)	57% (17%–84%)	16% (5%–32%)	21% (7%–41%)	29% (9%–52%)

* Women with missing stage information were classified as having stage III/IV disease if they met any of the following criteria: macroscopic residual disease; poorly differentiated cancer with no surgery; poorly differentiated cancer of serous subtype.

survival from ovarian cancer. While advanced age and stage were the factors associated with the worst survival, histological subtype, ascites and multiple comorbidities also influenced outcome. Further research is needed to explore whether geographic and socioeconomic differences relate to women's access to diagnostic and treatment services or to other factors. Overall, our results show that long-term survival among women with epithelial ovarian cancer remains poor, clearly reinforcing the need for prevention and better treatments.

Acknowledgements: This work was supported by Cancer Australia (formerly National Breast and Ovarian Cancer Centre). Collection of clinical data from the Australian Ovarian Cancer Study was funded by the National Health and Medical Research Council (NHMRC) (400281, 400413); Cancer Councils of Victoria, Queensland, New South Wales, South Australia and Tasmania; and the Cancer Foundation of Western Australia. Susan Jordan and Penelope Webb are funded by fellowships from the NHMRC. We acknowledge the members of the Patterns of Care Study Group; and the research nurses and associated staff for assistance with data collection. We also acknowledge the Australian Ovarian Cancer Study Group for providing clinical information for women who participated in that study. Full membership of the Australian Ovarian Cancer Study Group is listed at <http://www.aocstudy.org>.

Competing interests: No relevant disclosures.

Received 30 Jan 2014, accepted 27 Jun 2014.

- 1 Australian Institute of Health and Welfare, Cancer Australia. Gynaecological cancers in Australia: an overview. Canberra: AIHW, 2012. (AIHW Cat. No. CAN 66; Cancer Series No. 70.) <http://www.aihw.gov.au/publication-detail/?id=10737422954> (accessed Apr 2014).
- 2 Laurvick CL, Semmens JB, Holman CD, Leung YC. Ovarian cancer in Western Australia (1982–98): incidence, mortality and survival. *Aust N Z J Public Health* 2003; 27: 588–595.
- 3 Grossi M, Quinn MA, Thursfield VJ, et al. Ovarian cancer: patterns of care in Victoria during 1993–1995. *Med J Aust* 2002; 177: 11–16.
- 4 Choi M, Fuller CD, Thomas CR Jr, Wang SJ. Conditional survival in ovarian cancer: results from the SEER dataset 1988–2001. *Gynecol Oncol* 2008; 109: 203–209.
- 5 Jordan S, Steer C, DeFazio A, et al. Patterns of chemotherapy treatment for women with invasive epithelial ovarian cancer -- a population-based study. *Gynecol Oncol* 2013; 129: 310–317.
- 6 Australian Institute of Health and Welfare. Rural, regional and remote health: a guide to remoteness classification. Canberra: AIHW, 2004. (AIHW Cat. No. PHE 53; Rural Health Series No. 4.) <http://www.aihw.gov.au/publication-detail/?id=6442467589> (accessed Apr 2014).
- 7 Australian Bureau of Statistics. Information paper: census of population and housing. Socio-Economic Indexes for Areas, Australia 2001. Canberra: ABS, 2003. (ABS Cat. No. 2039.0.) <http://www.abs.gov.au/AUSSTATS/abs@nsf/DetailsPage/2039.02001?OpenDocument> (accessed May 2014).
- 8 Charlson ME, Pompei P, Ales KL, MacKenzie CR. A new method of classifying prognostic

4 Five-year relative survival in selected OECD countries for women diagnosed with ovarian cancer in the period 2003–2008

Country	5-year relative survival
Australia ¹	43%
United States ¹⁰	44%
England ¹¹	43%
Canada ¹²	45%
Japan ¹³	55%
Denmark ¹⁴	37%
Sweden ¹⁴	45%

OECD = Organisation for Economic Co-operation and Development.

comorbidity in longitudinal studies: development and validation. *J Chronic Dis* 1987; 40: 373–383.

- 9 Merritt MA, Green AC, Nagle CM, Webb PM; Australian Cancer Study (Ovarian Cancer); Australian Ovarian Cancer Study Group. Talcum powder, chronic pelvic inflammation and NSAIDs in relation to risk of epithelial ovarian cancer. *Int J Cancer* 2008; 122: 170–176.
- 10 Siegel R, Naishadham D, Jemal A. Cancer statistics, 2012. *CA Cancer J Clin* 2012; 62: 10–29.
- 11 Office for National Statistics. Cancer survival rates, cancer survival in England, patients diagnosed 2005–2009 and followed up to 2010. London: ONS, 2011. <http://www.ons.gov.uk/ons/publications/re-reference-tables.html?edition=tcm%3A77-239726> (accessed May 2014).

- 12 Canadian Cancer Society's Advisory Committee on Cancer Statistics. Canadian cancer statistics 2013. Toronto: Canadian Cancer Society, 2013. <http://www.cancer.ca/-/media/cancer.ca/CW/cancer%20information/cancer%20101/Canadian%20cancer%20statistics/canadian-cancer-statistics-2013-EN.pdf> (accessed Jun 2014).
- 13 Monitoring of cancer incidence in Japan – survival 2003-2005 report. Tokyo: Center for Cancer Control and Information Services, National Cancer Center, 2013. http://ganjoho.jp/pro/statistics/en/table_download.html (accessed Jun 2014).
- 14 Engholm G, Ferlay J, Christensen N, et al. NORDCAN: cancer incidence, mortality, prevalence and survival in the Nordic countries. Version 6.1 (25.04.2014). Copenhagen: Association of the Nordic Cancer Registries, Danish Cancer Society 2014. <http://www.ancre.nu> (accessed May 2014).
- 15 O'Malley CD, Cress RD, Campleman SL, Leiserowitz GS. Survival of Californian women with epithelial ovarian cancer, 1994-1996: a population-based study. *Gynecol Oncol* 2003; 91: 608-615.
- 16 Tingulstad S, Skjeldestad FE, Halvorsen TB, Hagen B. Survival and prognostic factors in patients with ovarian cancer. *Obstet Gynecol* 2003; 101 (5 Pt 1): 885-891.
- 17 Yang L, Klint A, Lambe M, et al. Predictors of ovarian cancer survival: a population-based prospective study in Sweden. *Int J Cancer* 2008; 123: 672-679.
- 18 Shimizu Y, Kamoi S, Amada S, et al. Toward the development of a universal grading system for ovarian epithelial carcinoma: testing of a proposed system in a series of 461 patients with uniform treatment and follow-up. *Cancer* 1998; 82: 893-901.
- 19 Rauh-Hain JA, Diver EJ, Clemmer JT, et al. Carcinosarcoma of the ovary compared to papillary serous ovarian carcinoma: a SEER analysis. *Gynecol Oncol* 2013; 131: 46-51.
- 20 Brown E, Stewart M, Rye T, et al. Carcinosarcoma of the ovary: 19 years of prospective data from a single center. *Cancer* 2004; 100: 2148-2153.
- 21 Chan JK, Teoh D, Hu JM, et al. Do clear cell ovarian carcinomas have poorer prognosis compared to other epithelial cell types? A study of 1411 clear cell ovarian cancers. *Gynecol Oncol* 2008; 109: 370-376.
- 22 Pignata S, Ferrandina G, Scarfone G, et al. Activity of chemotherapy in mucinous ovarian cancer with a recurrence free interval of more than 6 months: results from the SOCRATES retrospective study. *BMC Cancer* 2008; 8: 252.
- 23 Puls LE, Duniho T, Hunter JE, et al. The prognostic implication of ascites in advanced-stage ovarian cancer. *Gynecol Oncol* 1996; 61: 109-112.
- 24 Tetsche MS, Nørgaard M, Jacobsen J, et al. Comorbidity and ovarian cancer survival in Denmark, 1995-2005: a population-based cohort study. *Int J Gynecol Cancer* 2008; 18: 421-427.
- 25 Maas HA, Kruitwagen RF, Lemmens VE, et al. The influence of age and co-morbidity on treatment and prognosis of ovarian cancer: a population-based study. *Gynecol Oncol* 2005; 97: 104-109.
- 26 Coory MD, Ho T, Jordan SJ. Australia is continuing to make progress against cancer, but the regional and remote disadvantage remains. *Med J Aust* 2013; 199: 605-608.
- 27 Australian Institute of Health and Welfare, Australasian Association of Cancer Registries. Cancer survival in Australia 1992-1997: geographic categories and socioeconomic status. Canberra: AIHW, 2003. (AIHW Cat. No. CAN 17; Cancer Series No. 22.) <http://www.aihw.gov.au/publication-detail/?id=6442467451> (accessed Apr 2014). □