

The Challenge of Return to Work after Breast Cancer: The Role of Family Situation, CANTO Cohort

Elsa Caumette, Inès Vaz-Luis, Sandrine Pinto, Julie Havas, Thomas Bovagnet, Garazi Ruiz de Azua, Antonio Di Meglio, Anne-Laure Martin, Sibille Everhard, Paul Cottu, et al.

► To cite this version:

Elsa Caumette, Inès Vaz-Luis, Sandrine Pinto, Julie Havas, Thomas Bovagnet, et al.. The Challenge of Return to Work after Breast Cancer: The Role of Family Situation, CANTO Cohort. Current Oncology, 2021, 28 (5), pp.3866-3875. 10.3390/curroncol28050330. hal-03371184

HAL Id: hal-03371184 https://hal.sorbonne-universite.fr/hal-03371184v1

Submitted on 8 Oct 2021

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers. L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.





Article The Challenge of Return to Work after Breast Cancer: The Role of Family Situation, CANTO Cohort

Elsa Caumette ^{1,2}, Inès Vaz-Luis ^{3,4}, Sandrine Pinto ¹, Julie Havas ^{4,5}, Thomas Bovagnet ¹, Garazi Ruiz de Azua ¹, Antonio Di Meglio ⁴, Anne-Laure Martin ⁶, Sibille Everhard ⁶, Paul Cottu ⁷, Laurence Vanlemmens ⁸, Christelle Jouannaud ⁹, Florence Lerebours ¹⁰, Agnès Dumas ^{11,*}, and Gwenn Menvielle ¹

- ¹ IPLESP, Équipe de Recherche en Épidémiologie Sociale, INSERM, Institut Pierre Louis d'Épidémiologie et de Santé Publique, Sorbonne Université, 75012 Paris, France; elsa.caumette@umontpellier.fr (E.C.); sandrine.pinto@iplesp.upmc.fr (S.P.); thomas.bovagnet@iplesp.upmc.fr (T.B.); garazi.ruizdeazua@iplesp.upmc.fr (G.R.d.A.); gwenn.menvielle@inserm.fr (G.M.)
- ² Department of maieutics, University of Montpellier, 34000 Montpellier, France
- ³ Breast Cancer Unit, Department of Medical Oncology, Gustave Roussy, 94800 Villejuif, France; INES-MARIA.VAZ-DUARTE-LUIS@gustaveroussy.fr
- ⁴ INSERM Unit U 981, Gustave Roussy, 94800 Villejuif, France; Julie.HAVAS@gustaveroussy.fr (J.H.); Antonio.DI-MEGLIO@gustaveroussy.fr (A.D.M.)
- ⁵ Clinical Research Department, Gustave Roussy, 94800 Villejuif, France
- ⁶ UNICANCER, 75013 Paris, France; al-martin@unicancer.fr (A.-L.M.); s-everhard@unicancer.fr (S.E.)
- ⁷ Institute Curie, 75005 Paris, France; paul.cottu@curie.net
- ⁸ Center Oscar Lambret, 59000 Lille, France; l-vanlemmens@o-lambret.fr
- ⁹ Institute Jean Godinot, 51100 Reims, France; christelle.jouannaud@reims.unicancer.fr
- ¹⁰ Institute Curie, 92210 Saint Cloud, France; florence.lerebours@curie.fr
- ECEVE UMR 1123, INSERM (National Institute for Health and Medical Research), University of Paris, 75010 Paris, France
- * Correspondence: agnes.dumas@inserm.fr

Abstract: Return to work (RTW) after breast cancer is associated with improved quality of life. The link between household characteristics and RTW remains largely unknown. The aim of this study was to examine the effect of the family situation on women's RTW two years after breast cancer. We used data of a French prospective cohort of women diagnosed with stage I-III, primary breast cancer (CANTO, NCT01993498). Among women employed at diagnosis and under 57 years old, we assessed the association between household characteristics (living with a partner, marital status, number and age of economically dependent children, support by the partner) and RTW. Logistic regression models were adjusted for age, household income, stage, comorbidities, treatments and their side effects. Analyzes stratified by age and household income were performed to assess the association between household characteristics and RTW in specific subgroups. Among the 3004 patients included, women living with a partner returned less to work (OR = 0.63 [0.47-0.86]) and decreased their working time after RTW. Among the 2305 women living with a partner, being married was associated with decreased RTW among women aged over 50 (OR = 0.57 [0.34–0.95]). Having three or more children (vs. none) was associated with lower RTW among women with low household income (OR = 0.28 [0.10–0.80]). Household characteristics should be considered in addition to clinical information to identify vulnerable women, reduce the social consequence of cancer and improve their quality of life.

Keywords: return to work; breast cancer; family situation; cohort

1. Introduction

Breast cancer is the leading cancer in women [1] but it also has one of the best 5-year survival rates (nearly 90%) [2,3]. The high survival rate may be related to the combination of early diagnosis, novel surgical techniques, and new medications [4,5]. In France, in



Citation: Caumette, E.; Vaz-Luis, I.; Pinto, S.; Havas, J.; Bovagnet, T.; Ruiz de Azua, G.; Di Meglio, A.; Martin, A.-L.; Everhard, S.; Cottu, P.; et al. The Challenge of Return to Work after Breast Cancer: The Role of Family Situation, CANTO Cohort. *Curr. Oncol.* **2021**, *28*, 3866–3875. https:// doi.org/10.3390/curroncol28050330

Received: 14 September 2021 Accepted: 29 September 2021 Published: 1 October 2021

Publisher's Note: MDPI stays neutral with regard to jurisdictional claims in published maps and institutional affiliations.



Copyright: © 2021 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (https:// creativecommons.org/licenses/by/ 4.0/). 2018, 58,400 women were diagnosed with breast cancer, about half of them were between 20 and 59 years of age [6]. A large proportion of women are therefore working when diagnosed, and cancer can have an impact on their work life [7,8]. Furthermore, studies showed that the loss of work after cancer could affect women's quality of life as well as financial situation [9–11] and that return to work (RTW) was seen by women as an important part of their recovery. Understanding all determinants of RTW is thus essential to help patients build a new normality through work and reduce the burden of breast cancer in survivors' life [12]. Many studies showed that clinical characteristics (stage, comorbidities, treatments and their side effects) were major determinants of RTW after cancer. An advanced stage of cancer, poor health at diagnosis, but also some treatments (e.g., chemotherapy and lymph node dissection) and their physical (e.g., lymphedema of the arm, arm pain and fatigue) and psychological (e.g., depression and anxiety) sequelae have a negative impact on women's RTW after breast cancer [13–16]. Favourable working conditions (e.g., non-manual work and support from colleagues) have a positive impact on RTW [17].

Although included in the REWORK-BC model, a theoretical model explaining the RTW process after breast cancer [18], few studies investigated association between household characteristics and RTW. First, living with a partner could be negatively associated with RTW if the partner can ensure the household's financial security alone [19]. This security could be reinforced by marriage [20]. However, living with a partner could also be positively associated with RTW if the partner provides emotional support, which helps the woman returning to work [21,22]. The studies investigating the association between patients' relationship status and RTW reported heterogeneous results [16,23]. Furthermore, the definition of couple differed between studies. Some differentiated between women living with a partner (without taking into account women's marital status) and women not living with a partner [16]. Others including a meta-analysis differentiated between married women and unmarried women (without considering whether the women lived with a partner) [23]. These classifications mixed the concepts of couplehood and marriage and therefore did not enable one to distinguish the impact on RTW of living with a partner from that of being married.

In addition to couplehood, the presence of children, which constitutes a financial burden but also possible support, could impact RTW with possible differences according to the number and age of the children. Several recent studies, including a meta-analysis, showed that having economically dependent children had a positive impact on RTW [23,24]. However, the number of dependent children or their age was seldom taken into account. In particular, no study specifically looked at young children, who could nevertheless be a particular burden because of their low level of autonomy. The literature nevertheless did not suggest any differences, in terms of RTW, between women with or without minor dependent children [25].

Finally, several studies explored family support as a facilitator of RTW, but few specifically evaluated partner's support [22,26].

Age and socioeconomic position (SEP) are associated with RTW, with lower RTW among older [27] and low SEP (low level of education, manual work or low income) [28,29] women. These characteristics could influence the impact of household factors on RTW. Indeed, over the past decades, we observed in western countries delayed age at marriage and increased rate of second couplehood [30,31]. The association between couplehood, marriage and RTW may therefore differ by age. In addition, young children could be a barrier to RTW in particular among low SEP women because of the costs related to child care. Similarly, having a partner could be more closely associated with RTW among low SEP women, as complete wage could be more essential when the household income is low.

Overall, the link between household characteristics and RTW after breast cancer remains largely unknown and most studies on this topic used imprecise variables to characterize household because this was not their main aim. The objective of this study was to examine the effect of family situation on women's RTW two years after breast cancer and to assess whether these associations differed by patient's age or household income.

2. Materials and Methods

We used data from a French prospective cohort of women diagnosed with stage I-III first primary breast cancer (CANTO). Data collection started in 2012 at 26 cancer care centers. The study was approved by the French regulatory authorities (14 September 2011) and the French Ethics Committee (14 October 2011). All the patients enrolled in the study were aged 18 or over and provided written informed consent. Women were enrolled by an oncologist at breast cancer diagnosis. Data were collected prospectively during a clinical examination and through self-reported questionnaires at diagnosis, about 1 year after diagnosis (the visit took place 3 to 6 months after the end of treatment), and 2 years after diagnosis.

The analysis was based on women included until February, 2017. We limited the analysis to women under 57 years of age who were working at the time of diagnosis (n = 3919). We excluded women who were not treated with curative intent (patients with no surgery) (n = 12), as well as women with evidence of local or distant recurrence (n = 73), who withdrew consent (n = 80), lost to follow-up (n = 23) or deceased (n = 10) two years after diagnosis. Then, we excluded women whose RTW information two years post-diagnosis was unknown (n = 717). Finally, we included 3004 patients (Figure 1).



Figure 1. Flow chart of patient population.

The outcome was RTW two years after BC diagnosis and was collected by an ad hoc self-reported questionnaire. We investigated the association between RTW and various household characteristics self-reported at diagnosis in two populations: among all women and restricted to women living with a partner. In addition, we performed analyses stratified

by the women's age (under vs. over 50 years) and household income (below vs. above 2500 euros). Among all women, we studied the fact of living with a partner (yes/no) and the household structure combining the presence of a partner and economically dependent children (single/partnered women without/with children). Economically dependent children were defined as children aged below 25 living in the household. Among women living with a partner, we studied marital status (married/not married), perceived support by the partner(very strong/a little, not enough, not at all) as well as the number of economically dependent children (0, 1, 2, 3 or more) and their age. The impact of the age of children on RTW was assessed in a supplementary analysis. We identified women having children younger than 7 years of age (yes/no), as this age group is more dependent on adults [32], and women having children aged 18–25 (yes/no), as these could be graduate students with important financial needs.

Multivariate logistic regression analyses were performed. The models were adjusted for age (<40, 40–49 or \geq 50 years), clinical factors, treatment side effects at the first post-treatment visit (1 year after diagnosis) and household income (<2500, 2500–3000, 3000–4000, >4000 euros).

Clinical factors included stage at diagnosis (I, II, III), comorbidities at diagnosis, treatment and their side effects 1 year after diagnosis. Comorbidities at diagnosis were evaluated using the Charlson comorbidity index [33] (0, $1 \ge 2$) and a binary variable assessing the presence of ≥ 3 additional comorbid medical conditions not captured by the Charlson. Information about breast surgery (conservative/mastectomy), lymph node surgery (no or sentinel dissection/axillary dissection), systemic treatment (chemotherapy, hormone therapy, anti HER2 therapy) (yes/no) and radiotherapy (yes/no) was collected. Physical side effects of treatment were collected by a nurse, and defined as the presence of any severe toxicities (grade \geq 3) using version 4 of the Common Toxicity Criteria Adverse Events Scale (CTCAE) [34]. Additional treatment side effects were collected using three subscales of the breast cancer module (QLQ-BR23) (systemic therapy side effects, arm morbidity and breast morbidity as continuous variables) of the European Organization for Research and Treatment of Cancer (EORTC) self-reported quality-of-life questionnaire and the fatigue subscale (as continuous variable) of the core EORTC questionnaire(QLQ-C30) [35]. In addition, anxiety and depression were assessed using the Hospital Anxiety and Depression Scale (HADS) (non-case (0–7), doubtful (8–10), case (11–21) [36].

A sensitivity analysis was realized in the subgroup of women who returned to work and with known working conditions, to investigate the association between household characteristics and change in working time between diagnosis and two years after diagnosis using multinomial logistic regressions (full-time work two years after diagnosis, full-time work at diagnosis and part-time work two years after diagnosis, always part-time work). All analyses were performed with SAS 9.4 (SAS software for Windows version 9.4, SAS Institute Inc., Cary, NC, USA).

3. Results

In this study, the median age of women was 48 years (between 23 and 56 years). Women had 78% of the women lived with a partner and 11% were single mothers (Table 1). Most of the women living with a partner were married (73%) and received very strong support from their partner (85%). Furthermore, among women living with a partner, most women had dependent children (72%), 14% had children younger than 7 years of age and 32% had children aged 18–25.

	N (%) ⁺	% RTW		
Entire study population (<i>N</i> = 3004)				
Living with a partner				
Yes	2305 (78.5)	79.5		
No	632 (21.5)	79.9		
Missing	67			
Household structure				
Single woman with no economically dependent children	284 (9.7)	80.3		
Partnered woman with no economically dependent children	646 (22.2)	76.5		
Single mother	333 (11.4)	79.9		
Partnered woman with economically dependent children	1653 (56.7)	80.7		
Missing	88			
Women living with a partner ($N = 2305$)				
Marital status				
Married	1675 (73.0)	79.0		
Not married	618 (27.0)	81.1		
Missing	12			
Number of economically dependent children				
0	646 (28.1)	76.5		
1	524 (22.8)	79.0		
2	833 (36.2)	82.4		
3 or more	296 (12.9)	79.1		
Missing	6			
Support by the partner				
Very strong	1930 (84.7)	79.4		
A little, not enough, not at all	348 (15.3)	80.5		
Missing	27			

Table 1. Household demographic characteristics and proportion of women who returned to work two years after early breast cancer diagnosis.

N: Number of patients; RTW: Return to work. [†] The percentages do not include the missing data.

RTW two years after breast cancer diagnosis in relation to clinical characteristics was described in Table S1. About two thirds of the women worked full-time two years after diagnosis (regardless of their work time (full-time or part-time)at diagnosis), 17 % changed from a full-time to a part time job and 18% were still working part-time.

Among all women (Table 2), living with a partner was associated with lower RTW (OR = 0.63 [0.47-0.86]) and with a decrease in working time after RTW (OR = 1.6 [1.00-2.45] compared to full-time work two years after diagnosis (Table S2). RTW was increased for single women without dependent children (OR = 1.62 [1.07-2.47]) compared to women living with a partner and having dependent children.

Table 2. Odds ratios (ORs) and their 95% confidence intervals (CIs) between household demographic characteristics and return to work two years after early breast cancer diagnosis in all women by their age and household income (N = 2347).

	All Women	Analyses by Age Group		Analyses by Monthly Household Income	
	<i>N</i> = 2347	<50 Years of Age N = 1412	\geq 50 Years of Age N = 935	<2500 Euros N = 673	\geq 2500 Euros N = 1674
	OR † [95% CI]	OR ⁺ [95% CI]	OR ⁺ [95% CI]	OR † [95% CI]	OR ⁺ [95% CI]
Living with a partner					
No Yes	1 0.63 [0.47–0.86]	1 0.71 [0.47–1.08]	1 0.55 [0.35–0.88]	1 0.68 [0.47–1.00]	1 0.56 [0.32–0.99]
Household structure					
Single woman with no economically dependent children	1.62 [1.07–2.47]	1.49 [0.82–2.73]	1.77 [0.96–3.25]	1.7 [0.98–2.84]	2.23 [0.88–5.65]

	All Women N = 2347	Analyses by Age Group		Analyses by Monthly Household Income	
		<50 Years of Age N = 1412	\geq 50 Years of Age N = 935	<2500 Euros N = 673	$\geq 2500 \text{ Euros}$ $N = 1674$
	OR ⁺ [95% CI]	OR † [95% CI]	OR ⁺ [95% CI]	OR ⁺ [95% CI]	OR ⁺ [95% CI]
Partnered woman with no economically dependent children	0.90 [0.68–1.20]	1.04 [0.65–1.66]	0.84 [0.57–1.24]	1.22 [0.67–2.21]	0.81 [0.58–1.12]
Single mother with economically dependent children	1.45 [0.98–2.13]	1.38 [0.85–2.25]	1.47 [0.76–2.86]	1.48 [0.90–2.45]	1.37 [0.68–2,76]
Partnered woman with economically dependent children	1	1	1	1	1

Table 2. Cont.

⁺ Models additionally adjusted for age (except in the analyses stratified by age), household income (except in the analyses stratified by household income), stage at diagnosis, health at diagnosis (Charlson score, other medical antecedents), treatment(chemotherapy, hormone therapy, anti HER2 therapy, radiotherapy, breast surgery, lymph node surgery), health one year after diagnosis (fatigue, anxiety, depression, arm morbidity, breast morbidity, systemic therapy side effects, and severe physical toxicities).

Among women living with a partner (Table 3), being married was associated with decreased RTW only in those over age 50 (OR = 0.57 [0.34–0.95]). Having any (versus no) dependent children was associated with lower RTW only in the women with three or more children and in the low household income women (OR = 0.28 [0.10–0.80]). RTW also differed according to the age of economically dependent children. The results suggested lower RTW among women having any (N = 409) (versus no) dependent children under 7 years of age (OR = 0.73 [0.49–1.08]) and higher RTW among women having any (N = 736) (versus no) dependent children aged 18–25 (OR = 1.21 [0.91–1.60]). Perceived support by the partner was not associated with RTW either for the entire sample or in the age- or household income-stratified analyses.

Table 3. Odds ratios (ORs) and their 95% confidence intervals (CIs) between household demographic characteristics and return to work two years after early breast cancer diagnosis in partnered women according to age and household income.

	All Partnered Women	Analyses by Age Group		Analyses by Monthly Household Income	
	<i>N</i> = 1817	<50 Years of Age N = 1100	\geq 50 Years of Age N = 717	<2500 Euros N = 314	\geq 2500 Euros N = 1503
	OR ⁺ [95% CI]	OR ⁺ [95% CI]	OR ⁺ [95% CI]	OR ⁺ [95% CI]	OR [†] [95% CI]
Marital status					
Not married	1	1	1	1	1
Married	0.79 [0.59- 1.06]	0.92 [0.64–1.33]	0.57 [0.34- 0.95]	0.63 [0.34–1.18]	0.88 [0.63–1.23]
Number of economically dependent children					
0	1	1	1	1	1
1	1.02 [0.72–1.44]	0.76 [0.44-1.34]	1.30 [0.80-2.09]	1.03 [0.46-2.28]	1.02 [0.69-1.51]
2	1.25 [0.88–1.77]	1.11 [0.66-1.9]	1.17 [0.69-2.00]	0.65 [0.28–1,48]	1.42 [0.96-2.10]
3 or more	0.9 [0.64–1.55]	0.79 [0.44–1.42]	1.34 [0.48–3.76]	0.28 [0.10-0,80]	1.33 [0.79–2.23]
Perceived support by the partner					
A little, not enough, not at all	1	1	1	1	1
Verystrong	0.83 [0.58–1.19]	0.72 [0.43–1.21]	1.05 [0.62–1.76]	0.52 [0.24–1.11]	0.97 [0.64–1.48]

⁺ Models additionally adjusted for age (except in the analyses stratified by age), household income (except in the analyses stratified by household income), stage at diagnosis, health at diagnosis (Charlson score, other medical antecedents), treatment(chemotherapy, hormone therapy, anti HER2 therapy, radiotherapy, breast surgery, lymph node surgery), health one year after diagnosis (fatigue, anxiety, depression, arm morbidity, breast morbidity, systemic therapy side effects, and severe physical toxicities) and the other two sociodemographic variables.

4. Discussion

The aim of this study was to examine the effect of family situation (living with a partner and his support, being married, having children) on women's RTW and work time two years after breast cancer diagnosis.

There is no general consensus in the literature on the association between the presence of a partner and RTW after breast cancer [16,23,37]. In this study, at equal household income at diagnosis, living with a partner appeared to decrease RTW but also working time after RTW. This points to the importance of the financial dimension and the partner financial support for RTW through more freedom and flexibility in the process of RTW.

In addition to couplehood, we were able to study the specific effect of marriage on RTW after breast cancer in women living with a partner, which is almost never done in the literature [22]. In this study, among women living with a partner, marriage was generally not associated with RTW, but married women returned less to work than non-married women in the women over 50 years of age. Among women living with a partner and over 50 years of age most were married (79%) and married women had a statistically longer couple duration than the unmarried ones (median 30 years vs. 15 years, difference statistically significant). This is consistent with what is observed in the general population. Unmarried women over 50 years of age are therefore more likely to be in a recent relationship and consequently may have more psychological and financial independence, which would explain a higher rate of RTW among these women [38].

In the literature, there is a positive association between having dependent children and RTW after breast cancer [23]. The results of this study only partly support these findings. First, this analysis on the household structure and RTW among all women emphasizes the lack of association between the presence of economically dependent children and RTW. As regards partnered women, we found that having dependent children had a negative effect on RTW, but only among the women with at least three children and a low household income. In the general population, a threshold effect is observed in the association between the number of children and the employment rate: women work much less from three children upwards [39]. The negative effect of having more than three children and a low SEP on RTW could thus sum up and constitute an important barrier to RTW.

In the scarce literature taking into account the children's age, a study did not find any association between having minor dependent children and RTW [25]. A German cohort study nevertheless observed that having children decreased RTW among women between 18 and 44 years of age compared to older women [24], suggesting that young children may be a barrier to returning to work. This is consistent with our findings. Although based on small numbers, we observed a tendency towards lower RTW among women with children below age 7 and increased RTW among women having children aged 18–25. The latter finding may be related to the need for money associated with the fees of graduate studies.

Theoretical models such as the REWORK-BC model already suggested the importance of household characteristics in RTW process after breast cancer [18] but this study provides new insights into the role of household characteristics on RTW thanks to the large sample size, the detailed information collected on household variables, and the inclusion of a large number of confounding factors (SEP, age, clinical factors, treatment and their side-effects). Then, while most studies only investigate RTW, we also studied change in working time after RTW. Nevertheless, several limitations should be discussed. First, perceived support by the partner was collected only at the time of diagnosis whereas it can vary during the course of the disease and should therefore be better assessed over time. This may explain why we did not find partner's moral support to be associated with RTW. Indeed, in the literature, the qualitative [40] and quantitative [22] studies that mentioned partner's support as a facilitator of RTW after breast cancer measured this parameter over the long term (e.g., 3 to 4 years after diagnosis). Secondly, 19% of the eligible participants did not answer to the question on RTW, which could have biased the results of the study. Missing information on RTW was significantly less frequent for women living with a partner, after

40 years old, with a higher household income, and treated by hormonotherapy. Therefore, the proportion of women living with a partner could be slightly overestimated.

For most patients, cancer is a physical and emotional earthquake in their life [41]. The RTW period is often seen as the first step towards a new normal life [42]. The results of this study identified populations who were less likely to RTW. These patients could be at risk for poorer quality of life and psychosocial wellbeing. It is therefore essential that clinicians, as the first contact with patients, identify and refer them to social workers early in the disease process, therefore reducing the burden of breast cancer on women's life.

5. Conclusions

RTW is a complex social process where clinical, demographic and social variables interact. In addition to women's clinical characteristics, demographic household characteristics are important determinants for RTW after breast cancer diagnosis and working time after RTW. These associations differ by age and household income. Further studies investigating the reasons to stop working would help better understand the barriers for RTW.

Supplementary Materials: The following are available online at https://www.mdpi.com/article/ 10.3390/curroncol28050330/s1, Table S1: Clinical characteristics and proportion of women who returned to work two years after early breast cancer diagnosis, Table S2: Odds ratios (ORs) and their 95% confidence intervals (CIs) between household demographic characteristics and evolution in working time between diagnosis and two years after early breast cancer diagnosis in all women and in partnered women.

Author Contributions: Conceptualization/Methodology: E.C., A.D. and G.M.; Resources: I.V.-L. and G.M.; Data curation, formal analysis: E.C., A.D., I.V.-L., T.B., J.H., A.D.M., S.P. and G.M.; Writing: E.C., I.V.-L., S.P., J.H., T.B., G.R.d.A., A.D.M., A.-L.M.; S.E., P.C., L.V., C.J., F.L., A.D. and G.M.; Validation: E.C., I.V.-L., S.P., J.H., T.B., G.R.d.A., A.D.M., A.-L.M.; S.E., P.C., L.V., C.J., F.L., A.D. and G.M.; Supervision: G.M. and A.D. All authors have read and agreed to the published version of the manuscript.

Funding: The study was supported by French Foundation for Cancer Research Grant No. ARC, PGA1 RF20170205420, and by Susan G. Komen Career Catalyst Research Grant No. CCR17483507 to I.V.L. and the Philanthropic Odyssea Gustave Roussy Program. The CANTO study is supported by the French Government under the Investment for the Future program managed by the National Research Agency, Grant No. ANR-10-COHO0004. Elsa CAUMETTE's work research was financed by department of maieutics, university of Montpellier.

Institutional Review Board Statement: The study was approved by the French regulatory authorities (14 September 2011) and the French Ethics Committee (14 October 2011).

Informed Consent Statement: Informed consent was obtained from all subjects involved in the study.

Data Availability Statement: Deidentified participant data available under request to control the results of the study (contact the corresponding author, agnes.dumas@inserm.fr)The CANTO data are open to the scientific community. Any project can be submitted. Please contact the promotor Unicancer for further information: s-everhard@unicancer.fr.

Conflicts of Interest: I.V.-L. have personal financial interests: Novartis, Amgen, AstraZeneca (Public Speaking), Kephren (Writing engagements), P.C. received grants, personal fees and non-financial support from Novartis; personal fees, non-financial support and travel support from Roche; personal fees from Pfizer; personal fees and travel support from AstraZeneca.

References

- Sung, H.; Ferlay, J.; Siegel, R.L.; Laversanne, M.; Soerjomataram, I.; Jemal, A.; Bray, F. Global Cancer Statistics 2020: GLOBOCAN Estimates of Incidence and Mortality Worldwide for 36 Cancers in 185 Countries. CA Cancer J. Clin. 2021, 71, 209–249. [CrossRef] [PubMed]
- Bray, F.; Ferlay, J.; Soerjomataram, I.; Siegel, R.L.; Torre, L.A.; Jemal, A. Global Cancer Statistics 2018: GLOBOCAN Estimates of Incidence and Mortality Worldwide for 36 cancers in 185 countries. *CA Cancer J. Clin.* 2018, 68, 394–424. [CrossRef] [PubMed]
- 3. Société Canadienne du Cancer. Statistics on Breast Cancer. Available online: https://www.cancer.ca:443/fr-ca/cancerinformation/cancer-type/breast/statistics/?region=on (accessed on 15 December 2019).

- 4. Di Cosimo, S.; Appierto, V.; Pizzamiglio, S.; Silvestri, M.; Baselga, J.; Piccart, M.; Huober, J.; Izquierdo, M.; de la Pena, L.; Hilbers, F.S.; et al. Early Modulation of Circulating MicroRNAs Levels in HER2-Positive Breast Cancer Patients Treated with Trastuzumab-Based Neoadjuvant Therapy. *Int. J. Mol. Sci.* **2020**, *21*, 1386. [CrossRef] [PubMed]
- Von Minckwitz, G.; Procter, M.; de Azambuja, E.; Zardavas, D.; Benyunes, M.; Viale, G.; Suter, T.; Arahmani, A.; Rouchet, N.; Clark, E.; et al. Adjuvant Pertuzumab and Trastuzumab in Early HER2-Positive Breast Cancer. N. Eng. J. Med. 2017, 377, 122–131. [CrossRef]
- 6. Santé Public France. National Estimates of Cancer Incidence and Mortality in Metropolitan France between 1990 and 2018—Volume 1: Solid Tumors: Study Based on the Cancer Registers of the Francim Network. Available online: https://www.santepubliquefrance.fr/maladies-et-traumatismes/cancers/cancer-du-sein/documents/rapport-synthese/ estimations-nationales-de-l-incidence-et-de-la-mortalite-par-cancer-en-france-metropolitaine-entre-1990-et-2018-volume-1tumeurs-solides-etud (accessed on 7 January 2020).
- 7. Butow, P.; Laidsaar-Powell, R.; Konings, S.; Lim, C.Y.S.; Koczwara, B. Return to work after a cancer diagnosis: A meta-review of reviews and a meta-synthesis of recent qualitative studies. *J. Cancer Surviv.* **2020**, *14*, 114–134. [CrossRef]
- 8. Cocchiara, R.; Sciarra, I.; D'Egidio, V.; Sestili, C.; Mancino, M.; Backhaus, I.; Mannocci, A.; De Luca, A.; Frusone, F.; Di Bella, O.; et al. Returning to work after breast cancer: A systematic review of reviews. *Work* **2018**, *61*, 463–476. [CrossRef]
- 9. Ganz, P.A. Current issues in cancer rehabilitation. *Cancer* **1990**, *65*, 742–751. [CrossRef]
- Azzani, M.; Roslani, A.C.; Su, T.T. The perceived cancer-related financial hardship among patients and their families: A systematic review. *Support. Care Cancer* 2014, 23, 889–898. [CrossRef]
- Duijts, S.F.A.; Van Egmond, M.P.; Gits, M.; Van Der Beek, A.J.; Bleiker, E.M. Cancer survivors' perspectives and experiences regarding behavioral determinants of return to work and continuation of work. *Disabil. Rehabil.* 2016, 39, 2164–2172. [CrossRef]
 Waddell, G.; Burton, A.K. *Is Work Good for Your Health and Well-Being?* The Stationery Office: London, UK, 2006.
- Dumas, A.; Luis, I.V.; Bovagnet, T.; El Mouhebb, M.; Di Meglio, A.; Pinto, S.; Charles, C.; Dauchy, S.; Delaloge, S.; Arveux, P.; et al. Impact of Breast Cancer Treatment on Employment: Results of a Multicenter Prospective Cohort Study (CANTO). J. Clin. Oncol. 2020, 38, 734–743. [CrossRef]
- 14. Ekenga, C.C.; Pérez, M.; Margenthaler, J.A.; Jeffe, D.B. Early-stage breast cancer and employment participation after 2 years of follow-up: A com-parison with age-matched controls. *Cancer* **2018**, *124*, 2026. [CrossRef] [PubMed]
- 15. Lee, M.K.; Kang, H.S.; Lee, K.S.; Lee, E.S. Three-Year Prospective Cohort Study of Factors Associated with Return to Work after Breast Cancer Diagnosis. *J. Occup. Rehabil.* **2016**, *27*, 547–558. [CrossRef] [PubMed]
- 16. Schmidt, M.E.; Scherer, S.; Wiskemann, J.; Steindorf, K. Return to work after breast cancer: The role of treatment-related side effects and potential impact on quality of life. *Eur. J. Cancer Care* **2019**, *28*, e13051. [CrossRef]
- 17. Mehnert, A.; De Boer, A.; Feuerstein, M. Employment challenges for cancer survivors. Cancer 2013, 119, 2151–2159. [CrossRef]
- 18. Porro, B.; Durand, M.-J.; Petit, A.; Bertin, M.; Roquelaure, Y. Return to work of breast cancer survivors: Toward an integrative and transactional conceptual model. *J. Cancer Surviv.* **2021**, 1–14. [CrossRef]
- 19. Salonen, P.; Rantanen, A.; Kellokumpu-Lehtinen, P.-L.; Huhtala, H.; Kaunonen, M. The quality of life and social support in significant others of pa-tients with breast cancer—A longitudinal study. *Eur. J. Cancer Care* **2014**, *23*, 274–283. [CrossRef]
- 20. Zissimopoulos, J.M.; Karney, B.; Rauer, A.J. Marriage and economic well-being at older ages. *Rev. Econ. Househ.* 2013, 13, 1–35. [CrossRef]
- 21. Suwankhong, D.; Liamputtong, P. Social Support and Women Living with Breast Cancer in the South of Thailand. *J. Nurs. Sch.* **2015**, *48*, 39–47. [CrossRef]
- 22. Fantoni, S.Q.; Peugniez, C.; Duhamel, A.; Skrzypczak, J.; Frimat, P.; Leroyer, A. Factors Related to Return to Work by Women with Breast Cancer in Northern France. *J. Occup. Rehabil.* **2009**, *20*, 49–58. [CrossRef]
- Wang, L.; Hong, B.Y.; Kennedy, S.A.; Chang, Y.; Hong, C.J.; Craigie, S.; Kwon, H.Y.; Romerosa, B.; Couban, R.J.; Reid, S.; et al. Predictors of Unemployment after Breast Cancer Surgery: A Systematic Review and Meta-Analysis of Observational Studies. *J. Clin. Oncol.* 2018, *36*, 1868–1879. [CrossRef] [PubMed]
- Heuser, C.; Halbach, S.; Kowalski, C.; Enders, A.; Pfaff, H.; Ernstmann, N. Sociodemographic and disease-related determinants of return to work among women with breast cancer: A German longitudinal cohort study. *BMC Health Serv. Res.* 2018, 18, 1000. [CrossRef]
- 25. Johnsson, A.; Fornander, T.; Olsson, M.; Nystedt, M.; Johansson, H.; Rutqvist, L.E. Factors associated with return to work after breast cancer treatment. *Acta Oncol.* 2007, *46*, 90–96. [CrossRef]
- Nilsson, M.; Olsson, M.; Wennman-Larsen, A.; Petersson, L.-M.; Alexanderson, K. Return to work after breast cancer: Women's experiences of encounters with different stakeholders. *Eur. J. Oncol. Nurs.* 2011, 15, 267–274. [CrossRef]
- De Boer, A.G.; Torp, S.; Popa, A.; Horsboel, T.; Zadnik, V.; Rottenberg, Y.; Bardi, E.; Bultmann, U.; Sharp, L. eLong-term work retention after treatment for cancer: A systematic review and meta-analysis. *J. Cancer Surviv.* 2020, 14, 135–150. [CrossRef] [PubMed]
- Landeiro, L.C.G.; Gagliato, D.M.; Fêde, A.B.; Fraile, N.M.; Lopez, R.M.; da Fonseca, L.G.; Testa, L.; Hoff, P.M.; Mano, M.S. Return to work after breast cancer diagnosis: An obser-vational prospective study in Brazil. *Cancer* 2018, 124, 4700–4710. [CrossRef] [PubMed]

- Carlsen, K.; Jensen, A.J.; Rugulies, R.; Christensen, J.; Bidstrup, P.E.; Johansen, C.; Madsen, I.E.H.; Dalton, S.O. Self-reported work ability in long-term breast cancer survi-vors. A population-based questionnaire study in Denmark. *Acta Oncol.* 2013, 52, 423–429. [CrossRef] [PubMed]
- INSEE. Nuptiality—Demographic Report 2019. Available online: https://www.insee.fr/fr/statistiques/1892240?sommaire=1912 926 (accessed on 11 August 2020).
- Bureau UC. Historical Marital Status Tables. The United States Census Bureau. Available online: https://www.census.gov/data/ tables/time-series/demo/families/marital.html (accessed on 14 September 2020).
- CDC. Child Development: Middle Childhood (6–8 Years Old) | CDC Centers for Disease Control and Prevention. 2021. Available online: https://www.cdc.gov/ncbddd/childdevelopment/positiveparenting/middle.html (accessed on 28 July 2021).
- 33. Charlson, M.E.; Pompei, P.; Ales, K.L.; MacKenzie, C. A new method of classifying prognostic comorbidity in longitudinal studies: Development and validation. *J. Chronic Dis.* **1987**, *40*, 373–383. [CrossRef]
- 34. US Department of Health and Human Services. *Common Terminology Criteria for Adverse Events (CTCAE)*; Version 4.0.; US Department of Health and Human Services: Washington, DC, USA, 2009; p. 196.
- Giesinger, J.M.; Kuijpers, W.; Young, T.; Tomaszewski, K.; Friend, E.; Zabernigg, A.; Holzner, B.; Aaronson, N.K. Thresholds for clinical importance for four key domains of the EORTC QLQ-C30: Physical functioning, emotional functioning, fatigue and pain. *Health Qual. Life Outcomes* 2016, 14, 1–8. [CrossRef]
- 36. Zigmond, A.S.; Snaith, R.P. The Hospital Anxiety and Depression Scale. Acta Psychiatr. Scand. 1983, 67, 361–370. [CrossRef]
- Bouknight, R.R.; Bradley, C.J.; Luo, Z. Correlates of Return to Work for Breast Cancer Survivors. J. Clin. Oncol. 2006, 24, 345–353. [CrossRef]
- 38. Domínguez-Folgueras, M.; Lesnard, L. Family and social changement. L'Ann. Sociol. 2018, 68, 295–314.
- 39. Eurostat. Conciliation between Work and Family Life in the EU25 in 2003. Available online: https://ec.europa.eu/eurostat/web/ products-euro-indicators/-/3-12042005-ap (accessed on 14 March 2011).
- 40. Tamminga, S.J.; de Boer, A.G.E.M.; Verbeek, J.H.A.M.; Frings-Dresen, M.H.W. Breast cancer survivors' views of factors that influence the return-to-work process—A qualitative study. *Scand. J. Work Environ. Health* **2012**, *38*, 144–154. [CrossRef] [PubMed]
- Liamputtong, P.; Suwankhong, D. Breast cancer diagnosis: Biographical disruption, emotional experiences and strategic management in Thai women with breast cancer. *Sociol. Health Illn.* 2015, *37*, 1086–1101. [CrossRef] [PubMed]
- 42. Trusson, D.; Pilnick, A.; Roy, S. A new normal? Women's experiences of biographical disruption and liminality following treatment for early stage breast cancer. *Soc. Sci. Med.* **2016**, *151*, 121–129. [CrossRef]