

Treatment of Breast Cancer in Women Aged 80 and Older: A Systematic Review

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Abstract

Background: The elderly population is growing in the United States. Most clinical trials exclude patients over 80, therefore there is a paucity of data regarding the correct treatment of this group. The purpose of this systematic review was to investigate the treatment patterns for women with primary breast cancer aged 80 years old and older - modalities include surgery, chemotherapy, radiation and hormonal treatment, alone or in combination.

Methods: A formal systematic review was performed with the support of the medical research librarian at the University of California San Diego Biomedical Library. PubMed and Web of Science were the databases used. A patient population of 2,947 was derived from the 16 papers reviewed.

Results: Patients diagnosed over 80 were more likely to be diagnosed by clinical exam. Patients who had standard surgical treatment had an improved disease free survival. Surgical resection and radiation had a low morbidity.

Conclusion: Multimodality treatment is safe in elderly women and is associated with better breast cancer specific survival outcomes.

Keywords: Breast cancer; Elderly breast cancer; Elderly cancer treatments; 80 years and older; Oncologic surgery in the elders

Introduction

According to the latest United States census, there are 16.7 million Americans aged 80 and older, and 65% of these seniors are women [1]. The elderly are the fastest growing segment of the population, and the risk of cancer greatly increases with age. In the US, cancer is the second most common cause of death in women aged 75 years and older [2]. The risk of breast cancer nearly triples for women aged 70 to 80 to a rate of 43 in 1000 women, compared to 15 in 1000 women aged 40-50 [2].

There is no agreed upon standard of care for elderly women with breast cancer. In addition, the screening guidelines for yearly mammograms are vague and contradictory [2-6]. The decision for elderly women to continue breast cancer screening is a controversial issue, [7] and the pursuit of treatment for newly discovered breast cancer is an equally complex decision that is lacking in data. Factors that influence treatment patterns are difficult to measure and report. Though patients apply their own psychosocial factors, physician recommendation was the most influential factor in treatment decision [8]. Given the heterogeneity of the population, providers may not recommend treatment, even though there is evidence that octogenarian women with breast cancer die from their disease as often as younger women [9]. Furthermore, evidence shows that older women tend to face the same as younger women with comparable treatment [10,11].

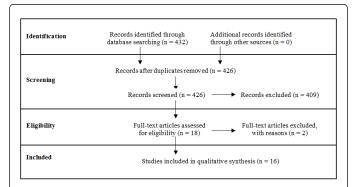
Patients over 65, and especially over 80, are underrepresented in clinical trials and are either overtly excluded or not enrolled [12-14]. The National Surgical Adjuvant Breast and Bowel Project studies, for

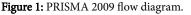
example B-18 and B-32, [15,16] compare treatments in patients younger or older than 50 years of age. Hutchins et al. [17] examined the underrepresentation of patients over 65 years of age in cancertreatment trials, and found that only 9% of breast patients enrolled in clinical trials are over 65. In a trial by Muss et al. [18], which specifically studied the use of chemotherapy in the elderly (over 65), only 4.4% of the patient cohort was over 80. Furthermore, it could be argued that a 65-year cohort is not elderly anymore, as our population continues to age and life expectancy rises. Little to no research exists for the oldest of old, specifically octogenarians. The goal of this review is to investigate the treatment patterns used for women with primary breast cancer aged 80 years old and older-modalities include surgery, chemotherapy, radiation and hormonal treatment, alone or in combination-and their survival.

Methods

A formal systematic review was performed with the support of the medical research librarian at the University of California San Diego Biomedical Library. PubMed and Web of Science were the databases used and a search was conducted following the Preferred Reporting Items for Systematic reviews and Meta-Analysis (PRISMA) Statement, issued 2009 (Figure 1). PICO was used to build the search criteria (Figure 2). The PRISMA Group worked to revise prior reviews and meta-analyses guidelines that had been established called QUOROM (Quality of Reporting of Meta-analyses), and released the PRISMA statement [19]. The goal of these groups was to address poor quality reviews and reporting of randomized data in the format of meta-analyses. The principle is simple – researchers can follow the PRISMA guidelines to develop high quality research questions, capture the relevant studies, and critically appraise the relevant studies. The

PRISMA guidelines include a 27-point checklist, and a four-phase flow diagram that guides researchers through the review process.

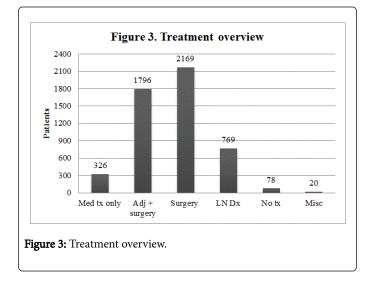




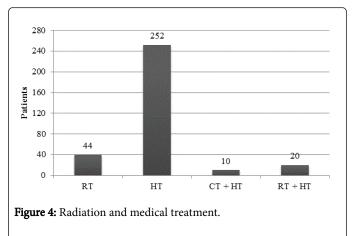
Patient/ population	Elderly women (80yrs+) with breast cancer
Intervention	Treatment patterns including surgery, chemotherapy, radiation, and hormone treatment (in any combination or alone)
Comparison	Elderly women (80yrs+) with breast cancer that are not treated
Outcome	Survival

Figure 2: PICO process.

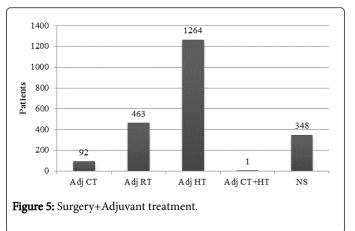
An updated search on PubMed and Web of Science was conducted using MeSH terms (Medical Subject Headings) in PubMed, and keywords in both PubMed and Web of Science. The search terms used include the following: "Breast Neoplasms"[MeSH] OR "Breast Neoplasms" AND "elderly"; then keywords "treatment", "surgery", "chemotherapy", "hormone therapy" and the associated MeSH term for each treatment modality. Lastly "survival" or "recurrence" search terms were added, AND NOT "metastatic" NOT "advanced"; filter: "aged 80 and over" was applied. Eligible literature included randomized trials and retrospective studies on the treatment of elderly women. The papers had to be written in English, full-text peer-reviewed, and published after 1/1/1997.



Breast Can Curr Res, an open access journal



The year 1997 was chosen because this is the year that anastrazole was approved, thus including publications that would have modern era hormonal treatments available. Selection criteria included primary breast cancer, early stage breast cancer, and patients 80 years old and older. Case reports, reviews, book chapters, patients with metastases, and patients with inflammatory cancer were excluded.



The literature review was performed in July of 2015. 432 papers were identified by JR. After duplicates were removed, 426 papers were screened using the abstracts in parallel by JR and AW. Early on during abstract review, it was recognized that there were very few papers that reported specifically on the over 80 population and early stage breast cancer. Thus the inclusion criteria was expanded to include all stages of breast cancer, except metastatic. 18 papers fit the above inclusion criteria and were selected to undergo full-text review. Every manuscript and patient cohort was closely examined. One study was excluded from the final review [20] because the patient population was identical to a prior study by the same author group [21]. A second manuscript was excluded because it is a SEER based study, which includes all patients in the United States, which inherently overlaps the trials or institutional retrospective patient cohort studies [9].

Two independent reviewers reviewed the remaining 16 papers, and data was separately tabulated [21-36]. Data extracted included number of patients \geq 80 years old, stage of disease at time of treatment, tumor characteristics, the different types of treatment plans, and survival (when available). Data integrity and any discordance were settled by consensus between AW and JR. Data was pooled and presented.

Page 3 of 9

GRADE analysis was then performed to determine the quality of evidence, and strength of recommendation.

Results

A patient population of 2,947 was derived from the 16 papers reviewed. A summary of the papers reviewed is shown in Tables 1-3. At time of treatment, 93 (3.2%) women had non-invasive disease, 1,736

(58.9%) women had early stage breast cancer (stage 1 or stage 2), 527 (17.9%) women had late stage breast cancer (stage 3 or stage 4), and 169 (5.7%) women had an unknown or unreported stage of disease. Table 4 shows stage, tumor size and hormone status when reported. See Supplemental Table 1 for the complete list and summary of the papers reviewed. The significant heterogeneity of treatments provided and outcomes studied is displayed.

Paper	Year	Median follow- up	# of pts	Treatment	Overall survival	Breast cancer specific survival	Study design	
Mery et al. [18]	2014	39 weeks	44	Radiation	88%	NR	Retrospective cohort study	
Bouchardy et al. [15]	2003	5 years	48	None	5-yr: 11%	5-yr: 46%	Retrospective cohort study	
ai. [15]			132	Hormone tx	5-yr: 18%	5-yr: 51%	conort study	
			28	BCS alone	5-yr: 27%	5-yr: 63%		
			55	Mastectomy	5-yr: 52%	5-yr: 82%		
			78	Mastectomy +adjuvant tx	5-yr: 44%	5-yr: 62%		
			57	BCS +adjuvant tx	5-yr: 67%	5-yr: 91%		
			9	Misc	5-yr: 22%	5-yr: 42%		
Cortadellas et al. [16]	2013	65 months	30	Unspecified surgery + chemo	NR	Early stage cancer: 108 mos (95%Cl 101-115);	Retrospective cohort study	
	42 Unspecified surgery + ra					Late stage cancer: 76 mos (95% CI 62-89)		
			140	Unspecified surgery + hormone tx				
			58	Hormone tx		Early stage cancer: 50 mos (95%Cl		
			10	Chemo + hormone tx		39-61) Late stage cancer: 68 mos (95% CI		
			16	Radiation + hormone tx		49-86)		
Dialla et al.	2012	3 years	15	None	1-yr RS: 93.5% (95%Cl	NR	Retrospective	
[19]			33	Unspecified surgery alone	91-96)* 5-yr RS: 78.2% (95%CI		cohort study	
			62	Hormone tx	72-83)*			
			94	BCS + unspecified adjuvant tx				
			119	Mastectomy + unspecified adjuvant tx				
			8	Other				

Table 1: Studies that included nonsurgical treatment arms. *Relative Survival. These numbers were not stratified by age group or treatment type.NR=Not reported, TX=Treatment, BCS=Breast Conserving Surgery.

Four papers included non-surgical treatment strategies alone as a treatment arm, representing 1,078 patients. One paper had radiation alone as the only treatment modality. Two papers had hormone therapy alone, surgery alone, or surgery plus adjuvant treatment as the three treatment arms. Lastly, one paper reported hormone therapy, hormonal and chemotherapy, hormonal and radiation therapy, and surgery plus adjuvant treatments as the treatment arms. One paper reported overall and breast cancer specific survival, two papers reported overall survival only, and one reported breast cancer specific survival only. The longest follow up was 65 months. All studies were retrospective in nature.

Forty-four women (1.5%) underwent radiation as their only form of treatment.

There were multiple treatment modalities, as shown in Figure 3. Three hundred and twenty-six (11.1%) patients received only medical management. Of these women, 252 (77.3%) received hormone treatment, 10 (3.1%) received both chemotherapy and hormone

Page 4 of 9

treatment, and 20 (6.1%) underwent radiation and hormone treatment (Figure 4). Seventy-eight (2.6%) women elected for no treatment. Twenty (0.68%) patients were categorized as receiving miscellaneous treatment including 8 that underwent an unspecified combination of chemotherapy, radiation, chemotherapy and hormone treatment, chemotherapy and radiation, radiation and hormone treatment, chemotherapy and radiation and hormone treatment, neo-adjuvant chemotherapy and surgery and adjuvant treatment, neo-adjuvant chemotherapy and surgery; 3 underwent an unknown surgical procedure; and 9 underwent an unspecified treatment.

Paper	Year	Median follow-up	# of pts	Surgical tx	Adjuvant tx	Overall survival	Disease free survival	Study design
Litvak et al. [13]	2006	NR	103	Mastectomy, BCS	Radiation, hormone tx, chemotherapy	NR	NR	Prospective database
Morishita et al. [32]	1997	NR	39	Mastectomy, BCS	Radiation, hormone tx, chemotherapy	10-yrs: 78%	77%	Retrospective cohort study
Strader et al. [31]	2014	NR	50	Mastectomy, BCS	Radiation, hormone tx	NR	NR	Retrospective cohort study

 Table 2: Surgical treatment excluding the axilla. NR=Not Reported, TX=Treatment, Mast=Mastectomy, BCS=Breast Conserving Surgery.

Three papers included surgical treatment strategies that excluded management of the axilla. This represented 192 patients. All three papers examined mastectomy and breast conserving surgery. Two papers treated with radiation, hormonal, and chemotherapy; one paper treated with hormonal and radiation therapy only. One paper reported both overall and disease free survival; two did not report survival at all. Follow up terms were not reported. One study was a prospectively collected database study; the other two were retrospective studies.

Paper	Year	Median follow-up	# of pts	Surgical tx	Adjuvant tx		Overall survival	Disease free survival	Breast cancer specific survival	Study design
Angarita et al. [14]	2015	14 months	217	Mastectomy, BCS, SLNB, ALND	Radiation, hormone chemotherapy	tx,	≥80: 53 mos	≥80: 17.5 mos	NR	Retrospective cohort study
Besic et al. [21]	2014	5.3 years	154	Mastectomy, BCS, SLNB, ALND	Radiation, hormone chemotherapy	tx,	NR	NR	5-yr: 83%	Retrospective cohort study
Chatzidaki et al. [25]	2011	NR	129	Mastectomy, BCS, ALND			NR	NR	NR	Retrospective cohort study
Cyr et al. [22]	2011	34 months	134	Mastectomy, SLNB, ALND	Radiation, hormone chemotherapy	tx,	65%	NR	NR	Retrospective cohort study
Evron et al. [17]	2006	70 months	135	Mastectomy, BCS ALND	Radiation, hormone chemotherapy	tx,	75%	Median: 22 mos	91%	Retrospective cohort study
Guth et al. [20]	2013	13 months	79	Mastectomy, BCS, SLNB, ALND	Radiation, hormone chemotherapy	tx,	NR	NR	NR	Prospective database
Joerger et al. [12]	2012	NR	462	Mastectomy, BCS, SLNB	Radiation, hormone chemotherapy	tx,	NR	NR	NR	Prospective database
Rosenkranz et al. [24]	2006	4 years	213	Mastectomy, BCS, SLNB, ALND	Radiation, hormone chemotherapy	tx,	Median OS: 7.28 yrs	NR	NR	Retrospective cohort study
Vetter et al. [23]	2013	14 months	151	Mastectomy, BCS, SLNB, ALND	Radiation, hormone chemotherapy	tx,	36%	NR	NR	Prospective database

 Table 3: Multi-modality treatment, surgery in all patients.

Nine papers included surgical treatment strategies with combinations of adjuvant or neoadjuvant treatments, representing 1,674 patients. All papers included both mastectomy and breast conserving surgery, two papers included axillary node dissection as the only treatment of the axilla, one paper included sentinel node biopsy only, and the remaining six included a combination of the two axillary treatments. All papers included radiation, hormone therapy, and chemotherapy in different combinations. Three papers did not report

survival of any kind. One paper reported disease free, breast cancer specific, and overall survivals. Three reported overall survival only, one reported both disease free and overall survival, and one reported breast cancer specific survival only. The range of follow up was 13-70 months, two studies did not report follow up length. Three studies were prospective, and the other six were retrospective in nature. Overall, 2,169 surgical operations were performed (including breast-conserving surgery or mastectomy) with some women undergoing one or more

Page 5 of 9

procedures. Seven hundred sixty-nine (35.5%) lymph node dissections were done. Some of these patients (60.9%) underwent some form of adjuvant treatment, often more than one type of therapy. Ninety-two (5.1%) had adjuvant chemotherapy (1 woman in this group was given chemotherapy as neo-adjuvant treatment), in 14 (0.78%) of these women it was unclear whether this was the primary treatment or adjuvant. 463 (25.8%) underwent adjuvant radiation; 1,264 (70.1%) received adjuvant hormone treatment; 1 (0.06%) had both adjuvant chemotherapy and hormone treatment; and 384 (21.4%) underwent an

unspecified adjuvant treatment (Figure 5). In general, breast surgery and radiation were associated with low complication rates (37.1%), with few being serious complications (5.7%) [25]. Evron et al. [29] found that women who did not have axillary staging had a higher axillary failure rate, 5.3% versus 0% who underwent axillary staging. Chatzidaki et al. found it would be reasonable to recommend surgery for patients over 80 [25].

	Size			Stage				Recep	tor exp	pression						
Paper	Mean	Median	n Range	In situ	Early Stage	Late stage	Unk	ER+	ER-	ERunk	PR+	PR-	PR unk	HER2 / neu +	HER2 / neu -	HER2 / neu unk
Mery et al. [18]				4	38	17	1	34	9	1	27	16	1	2	41	1
Bouchardy et al. [15]	30 mm		4-130 mm	6	254	98	49	88	20	299						
Cortadellas et al. [16]					189	70										
Dialla et al. [19]	27 mm +/- 19.8	22 mm	2-160 mm					Positiv	e 590,	Negative 80), Unkno	wn 11*	*			
Litvak et al. [13]	2 cm +/- 2.4			5	85	13		43	14	47						
Morishita et al. [32]	2.38 cm	2.2 cm	1-7.5 cm													
Strader et al. [31]	20.1m m +/- 20.2			11	37			38			49					
Angarita et al. [14]		Invasive: 2.4 cm DCIS: 3 cm	Invasive: 1.7-3 cm DCIS: 1.7-4 cm	24	184	9		180	37		165	52		31	155	8*
Besic et al. [21]	37 mm	25 mm	5-150 mm					134	20		90	64				
Chatzidaki et al. [25]				5	100	23	12	100	16	24	78	36	26			
Cyr et al. [22]				16	103	9		94	19	33				11	101	34
Evron et al. [17]					121	11	3	Positiv	e 102,	Negative 21	, Unkno	wn 12*	*			
Guth et al. [20]					63	16		All "HF	R-positi	ve", part of i	nclusion	criteria	a	13	61	5
Joerger et al. [12]					340	178	104	438†	88† †	96				49	256	317
Rosenkranz et al. [24]				22	151	40		132	27							
Vetter et al. [23]		25 mm			108	43		Positiv	e 94, N	legative 27,	Unknow	/n 8**		15	10	1

Table 4: Tumor Characteristics. Unk = unknown. * HER2/neu equivocal. ** Did not differentiate between ER or PR. † >50% ER expression. †† 65with <10% ER expression, 23 with 10-50% ER expression.</td>

Rosenkranz et al. reported that surgical resection was safe in the older age group (0.5% major complications, 5% minor complications) but that intravenous chemotherapy was associated with a higher complication rate (30% major complications) [21].

Overall, the quality of the literature regarding treatment of elderly patients with breast cancer is low quality. Tables 5-7 present the GRADE analysis for each of our clinical questions. The risk of bias is very serious, since all papers are observational or retrospective; there are significant selection and observation biases. This causes the

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Page 6 of 9

literature to be low quality. The overall treatment recommendation that could be gathered from these studies is weak.

Outcome	Quali	ty assessment				Summary of findings							
	No of studi es	Study design	Risk of bias	Inconsist ency	Indirect ness	Impreci sion	Other considerat ions	Number of patients		Effect		Qualit y	
								Treated	Untreat ed	Relative (95% CI)	Absolute (95% CI)		
Survival	4	Retrospective cohort study, prospective database	Seriou s*	**	Not serious	Not serious	None	1018/10 37	107/10 37	***	***	Low	Critical

Table 5: Studies that included non-surgical treatment arms compared with no treatment for elderly women (80+) with breast cancer. Patients or population: Elderly women (80 yrs+) with breast cancer. Intervention: Medical treatment with or without surgery. Comparison: Elderly women (80 yrs+) with breast cancer that are not treated. *All studies are observational, which carry an inherent risk of bias. **Unable to assess from the data available. ***Unable to be calculated from the data available.

Outcome	Quali	ty assessment				Summary of findings							
	No of stud ies	Study design	Risk of bias	Inconsis tency	Indirect ness	Impreci sion	Other consider ations	Number of patients		Eff	ect	Quality	
								Treated	Untrea ted	Relative (95% CI)	Absolute (95% CI)		
Survival	3	Retrospective cohort study, prospective database	Serious*	**	Not serious	Not serious	None	192/192	NR	***	***	Low	Critical

Table 6: Surgical treatment excluding the axilla compared with no treatment for elderly women (80+) with breast cancer. Patients or population: Elderly women (80 yrs+) with breast cancer. Intervention: Surgical treatment excluding the axilla. Comparison: Elderly women (80 yrs+) with breast cancer that are not treated. NR = not reported. *All studies are observational, which carry an inherent risk of bias. **Unable to assess from the data available.

Outco me	Qualit	y assessment				Summary of findings							
	No of studi	Study design	Risk of	Inconsis tency	Indirect ness	Impreci sion	Other considerat ions	Number of patients		Effect		Qualit y	
	es		bias					Treated	Untreat ed	Relative (95% CI)	Absolute (95% CI)		
Surviva I	9	Retrospective cohort study, prospective database	Seriou s*	**	Not serious	Not serious	None	1674/16 74	NR	***	***	Low	Critical

Table 7: Multi-modality treatment, surgery in all patients compared with no treatment for elderly women (80+) with breast cancer. Patients or population: Elderly women (80 yrs+) with breast cancer. Intervention: Multi-modality treatment, surgery in all patients. Comparison: Elderly women (80 yrs+) with breast cancer that are not treated. NR = not reported. *All studies are observational, which carry an inherent risk of bias. **Unable to assess from the data available. ***Unable to be calculated from the data available.

Discussion

From this patient population of 2,947, the most common form of treatment was surgery (mastectomy or breast conserving surgery). A total of 2,169 operations were performed, and some women underwent multiple procedures. The most common form of adjuvant treatment was hormonal therapy, which was given to 1,264 (70.1%) women (pre-

or post-surgical timing of hormonal therapy is not specified in the studies).

The characteristics of primary tumors in elderly women are relatively homogeneous. More than 75% were ER positive, more than 60% were PR positive, and very few were HER2/neu positive (16.2%) [14,18,21,22,24,27,33]. In this population, less than 3% of women refused to undergo treatment of any kind, once their tumor was identified. This seemingly low rate of treatment refusal may not represent the population as a whole, though, since the percentage of octogenarians foregoing mammography is unknown.

There are several limitations of this systematic review, many of which are limitations of the body of literature as a whole. First, this lack of knowledge regarding rates of screening mammograms in octogenarians is a limitation; rates are not reported in any of the studies. The literature highlights that more than half of the tumors in octogenarians were discovered as palpable masses. Once the tumors were identified only 3% refused treatment. However, the rate of octogenarians who sought treatment (97%) may be an overrepresentation since the number of patients that continue to refuse mammograms is never captured. On the other hand, although many patients were not seeking routine screening, when a mass was identified most decided to pursue treatment. This brings to light that perhaps not enough elderly women, who would want treatment, are receiving screening mammograms.

Most of the limitations of this review are due to a wide variety of reporting styles that made the review and classification of our collected data difficult. While most of the papers reported the stage of the patients at time of diagnosis, very few continued the breakdown of treatment modality by stage of disease and some did not report it at all. This presents a unique problem in that survival was reported without adjusting for stage. Specific treatment plans were also reported inconsistently and varied greatly between the papers. Some only reported whether surgery occurred, while others reported the specific surgical intervention (breast-conserving surgery, mastectomy, axillary lymph node dissection, and/or sentinel lymph node biopsy). The reporting of adjuvant treatment also varied, and many reported the specific adjuvant treatment (chemotherapy, radiation, and/or hormone treatment) but not the timing. Other papers reported if the patient was given any adjuvant treatment, without specifying modality (radiation, chemotherapy, hormonal therapy). Some included several study arms in which one cohort received medical treatment only, another cohort received surgery and adjuvant treatment, but these were not compared to each other; instead observations were reported in a retrospective fashion. In a few cases, it was unclear if the medical treatment was the primary treatment or an adjuvant treatment. In summary, the quality of the data is low in all the existing studies regarding treatment of elderly patients with breast cancer.

Despite these weaknesses, this review brings to light several important ideas. Multiple studies show elderly breast cancer patients receive the same survival benefit as younger patients with similar disease [10,11,37-39]. Yet, multiple studies have shown age to be the greatest influence on treatment choice, especially when adjuvant radiation is indicated [20,28,29,31,32,36]. Bouchardy et al. found that less than half of women aged 80 and older received standard treatment [40].

Surgery, including mastectomy, breast-conserving surgery, and axillary dissection, has been shown to have a low complication rate overall and a very low rate of major complications [23,25]. Mery et al. found that 80% of elderly patients were able to complete radiation therapy [33]. In addition, studies by Bouchardy et al., Guth et al., Litvak et al., Van Leeuwen et al., and Vetter et al. showed that both medical and surgical treatment are safe for use in elderly women and multi-modality therapy should not be withheld [20,24,30,32,36]. In the face of these studies touting the safety of multimodality treatment,

Page 7 of 9

elderly women with breast cancer continue to receive less treatment than their younger counterparts.

Undertreatment does not uniformly result from ageist beliefs of the physician. Treatment may be discontinued due to recurrence or another serious medical problem unrelated to breast cancer [30]. Undertreatment may also be the result of patient refusal. Non-compliance, especially with adjuvant treatment, is much higher in elderly women than in younger cohorts [35,36]. Vetter et al. found that 13% of elderly patients chose to forgo adjuvant hormone therapy and 49% of elderly patients refused adjuvant radiation [36].

Regardless of the reason, undertreatment is associated with a decreased breast cancer specific survival rate. Bouchardy et al. found 5-year specific breast cancer survival rate dropped from 90% in women receiving breast conserving surgery and adjuvant treatment to 46% in women receiving no treatment [40]. In comparing single modality treatment, women who underwent mastectomy had a much higher 5-year specific breast cancer survival rate at 82% compared to 51% for women receiving only hormonal treatment [24]. Cortadellas et al. have similar results - women who underwent surgery had a mean breast cancer specific survival of 108 months while women who received only medical treatment had a mean breast cancer specific survival of 50 months [26].

Despite an aging population, the relative safety of treatments and the apparent survival benefits of treatment of the elderly, it is difficult to draw firm conclusions from the current body of literature. Although most of the large, randomized-controlled multi-institutional studies exclude the elderly [12-17], CALGB 9343 examined the benefit of radiation following breast-conserving surgery with tamoxifen, in patients over 70 with smaller ER positive tumors [41]. While CALGB 9343 was not included in this systematic review because it studied a slightly younger 70-year-old cohort, it is an important study to discuss. In women with stage 1 estrogen sensitive tumors that took tamoxifen, radiation showed a modest improvement in loco-regional recurrence, but there was no difference in survival. Thus, it is reasonable given this strong level of evidence for elderly patients with favorable histology to forego radiation in this circumstance. It is difficult, though, to expand these types of conclusion to other treatments without evidence. It is necessary to perform more studies like CALGB 9343, and it is the opinion of these authors that clinical trials should begin to include an elder cohort.

Conclusions

Women aged 80 years old and older with breast cancer are most often treated surgically. The majority of these women pursue adjuvant treatment, the most common modality of which is hormone treatment. The overall quality of literature regarding treatment of elderly breast cancer is low, and only weak recommendation to provide standard treatment can be made. Future randomized clinical trials should consider including an elderly cohort to provide better evidence, and guide decision-making in this growing population.

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Conflict of Interest

None

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Page 9 of 9

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