HRT IN BRCA CARRIERS following RRBSO

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• Is it effective?

• Is it safe?

- Very Little data
- Reviews > Studies





QOL AFTER RRBSO

Madalinska et al J Clin Oncol 2006

- Premenopausal women with BRCA mutation
- 286 women undergoing gynological screening (no surgery)
- 164 women post RRBSO

77 HRT -

87 NO HRT

Significantly less menopausal symptoms in HRT vs. No HRT Significantly more symptoms in HRT vs. no surgery group





QOL AFTER RRBSO Finch *et al* Gynecol Oncol 2011

- Prospective study 114 women surveyed prior to surgery and 1 year following surgery
- 66 % premenopausal , 47 % previous diagnosis of breast cancer
- Woman on HRT significantly less menopausal symptoms
- HRT didn't prevent post surgery decline in sexual satisfaction





BSO in young patients without HRT

- Bone health
- Cardiac Mortality HR 1.8
- Cognitive impairment/dementia HR 1.9





Osteoporosis risk and management in BRCA1 and BRCA2 carriers who undergo RRBSO Garcia *et al* Gynecol Oncol 2015

- 225 women median f-u 41 months
- Only 44% of BRCA carriers had a DXA scan after RRBSO
- 68% of DXA scans were abnormal.
- 4% rate of atraumatic fracture





Use of TAH and HRT in BRCA1 and BRCA2 mutation carriers undergoing RRSO

Gabriel et al Fam Cancer 2009

- Single institution study 73 carriers
- Majority of BRCA1/2 mutation carriers undergoing RRSO also underwent TAH
- A substantial number took HRT.
- TAH did not increase the likelihood of taking HRT compared to RRSO alone.





Is HRT safe?

Effect of Short-Term Hormone Replacement Therapy on Breast Cancer Risk Reduction After Bilateral Prophylactic Oophorectomy in BRCA1 and BRCA2 Mutation Carriers:

The PROSE Study Group

Rebbeck et al J Clin Oncol 2005





PROSE Study

- Prospective cohort of 462 women with BRCA1/2 mutation
- Incidence of breast cancer
 - 155 patients after RRBSO +/- HRT
 - 307 with no surgery

- Median F-U 3.6 years
- "Short term HRT"





Breast cancer risk reduction after BSO

stratified by postsurgical HRT use			
	Total sample	BSO BEFORE AGE 5	

HR

1.0

0.38

0.37

0.44

0.43

Abbreviations BSO, bilateral prophylactic, HRT, hormone replacement therapy, HR, hazard ratio, E2, estrogen, PROG, progesterone

No.

286

62

93

50

34

*adjusted for birth year, BRCA1 versus BRCA2 center of ascertainment, and parity

No HRT

No HRT

Any HRT

E2 only

PROG±E2

variable

No surgery

BSO

BSO

BSO

BSO

95%CL*

0.09 to 1.59

0.14 to 0.96

0.12 to 1.61

0.07 to 2.68

HR

1.0

0.59

0.3

0.44

0.43

No.

286

50

89

50

34

95%CL*

0.14 to 2.52

0.11 to 0.85

0.12 to 1.61

0.07 to 2.68

stratified by postsurgical HRT use			
	Total sample	BSO BEFORE AGE 50	

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stratified by postsurgical HRT use			
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Problems with PROSE study

- "Short term HRT"
 - not defined
 - -no data on duration provided
- Underpowered
- Biological plausibility



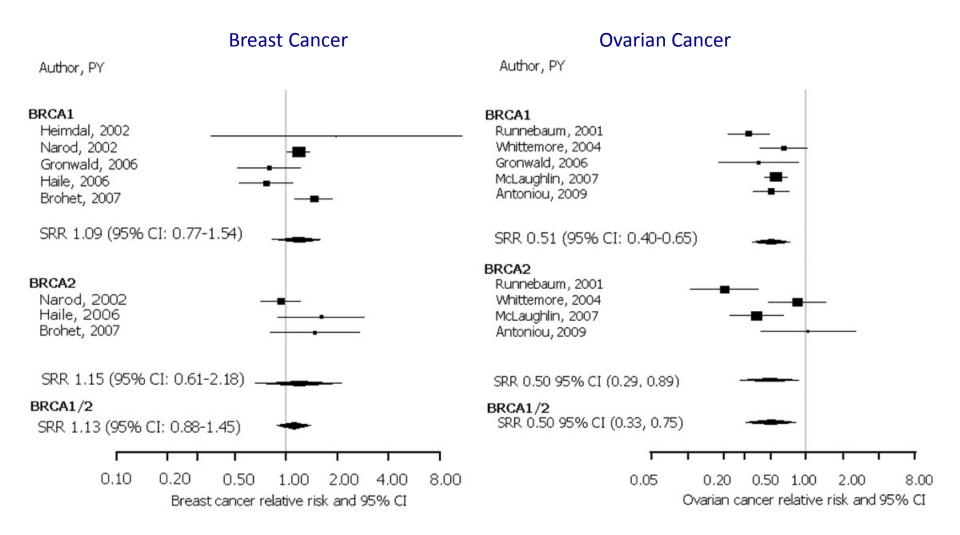


 Can we have reassurance from OC in BRCA carriers studies?





Oral contraceptive use and breast or ovarian cancer risk in IBRCA1/2 carriers: A meta-analysis lodice et al Eur J Cancer 2010



Forest plot and summary relative risk on the association between OC use and breast cancer (left) and ovarian cancer (right) in carriers. PY: publication year; SRR: Summary Relative Risk; CI confidence intervals