nature portfolio

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Reporting Summary

Nature Portfolio wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Portfolio policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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For all statistical a	nalyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.				
n/a Confirmed					
☐ ☐ The exac	t sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement				
A statem	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly				
The stati	stical test(s) used AND whether they are one- or two-sided mon tests should be described solely by name; describe more complex techniques in the Methods section.				
☐ X A descrip	otion of all covariates tested				
A descrip	otion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons				
	scription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) riation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)				
For null I	hypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted fues as exact values whenever suitable.				
For Baye	sian analysis, information on the choice of priors and Markov chain Monte Carlo settings				
For hiera	archical and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
Estimate	s of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated				
,	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.				
Software ar	nd code				
Policy information	about <u>availability of computer code</u>				
Data collection	All analyses were conducted using R v3.6.1.				
Data analysis	All analyses were conducted using R v3.6.1.				
	ng custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and vencourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio guidelines for submitting code & software for further information.				

Data

Policy information about <u>availability of data</u>

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

Individual participant data that underlie the results reported in this article are not able to be shared at this time as patients remain in study follow-up.

Field-spe	ecific	reporting			
Please select the or	ne below	that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.			
X Life sciences		Behavioural & social sciences Ecological, evolutionary & environmental sciences			
For a reference copy of t	the docume	nt with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
Life scier	nces	study design			
All studies must dis	sclose on	these points even when the disclosure is negative.			
Sample size	N/A	'A			
Data exclusions	N/A				
Replication	N/A				
Randomization	N/A				
Blinding	N/A				
Poportin	a fo	conscisio materials, systems and methods			
<u> </u>		r specific materials, systems and methods			
		uthors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, rant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.			
Materials & exp	perime	ntal systems Methods			
n/a Involved in th	ne study	n/a Involved in the study			
Antibodies		ChIP-seq			
Eukaryotic cell lines		Flow cytometry			
Palaeontol					
Animals an					
Human res	•	icipants			
Dual use re		concern			
Z					
Human rese	arch _l	participants			
Policy information	about <u>st</u>	dies involving human research participants			
Population characteristics		Population Characteristics detailed in Table 1: age, sex, race, ethnicity, baseline LVEF, BMI			
Recruitment		No bias in patients recruitment was identified			
Ethics oversight The study was a		The study was approved by the institutional review board at each site.			
Note that full informa	ation on tl	e approval of the study protocol must also be provided in the manuscript.			
Clinical data					
Policy information a		nical studies with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.			
Clinical trial regis		NCT01853748			
Study protocol		Protocol previously published and attached to this submission			
Data collection	collection From May 17, 2013 to December 13, 2016, 512 patients with stage I HER2-positive breast cancer were enrolled in the ATEMPT and 497 (383 T-DM1, 114 TH) started protocol therapy and were included in this analysis				

The incidence of grade 3-4 LVSD are secondary endpoints of the ATEMPT trial and are objectives of this current analysis.

Outcomes