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Reporting Summary

Nature Research 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 Research policies, see our Editorial Policies and the Editorial Policy Checklist.

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1 01	ali statisticai ali	alyses, commit that the following items are present in the figure regend, thank text, or Methods section.			
n/a	Confirmed				
	The exact	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement			
	A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly			
	The statis	tical test(s) used AND whether they are one- or two-sided on tests should be described solely by name; describe more complex techniques in the Methods section.			
	A descript	ion of all covariates tested			
	A descript	ion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons			
	A full desc	cription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) tion (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)			
	For null hy Give P valu	ypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted es as exact values whenever suitable.			
\boxtimes	For Bayes	ian analysis, information on the choice of priors and Markov chain Monte Carlo settings			
\boxtimes	For hierar	chical and complex designs, identification of the appropriate level for tests and full reporting of outcomes			
\boxtimes	Estimates	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.			
So	ftware an	d code			
Poli	Policy information about <u>availability of computer code</u>				
Da	ata collection	Standard, open-source tools used in data collection are described in the Methods section.			

For manuscripts utilizing 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 reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.

All computational tools used in data analysis are open-source tools that have been previously published, and are described in the Methods

Data

Data analysis

Policy information about availability of data

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 list of figures that have associated raw data
- A description of any restrictions on data availability

Access to patient sequence data is controlled to protect patient privacy, and thus requires the approval of the Data Access Committee. Requests to access the data should be submitted to genetics@cancerresearch.my.

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Please select the o	ne below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.					
Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences					
For a reference copy of	the document 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	The sample size was set in order to detect somatic mutation frequencies of 3% or above as described previously by Pan et al., 2020 Nature Communications. With 560 samples, the sample size also enables general and subtype-specific molecular comparisons of breast cancer to similar genomic studies of breast cancer in other published studies or datasets such as TCGA and METABRIC.					
Data exclusions	Patients were excluded from this study for the following criteria: No corresponding germline samples (n=5) and those who withdrew consent (n=12). Tumour samples were further excluded after clinicopathological review if they were found to be from rare histological subtypes and other breast diseases (n=5). Tumour samples with an average tumour content of <30% (n=50) and those with insufficient DNA (n=8) were excluded from the study. After sequencing, samples that did not reach standard sequencing quality metrics were also excluded.					
Replication	Where appropriate, all major findings were compared to previously published studies (Li et al. npj Breast Cancer 201, Lee et al. 2018). Some findings could not be compared to those studies due to lack of data or inadequate sample size.					
Randomization	Given the nature of the study, randomization was not relevant as comparator groups were based on presence/absence of germline or somatic mutation in PALB2, BRCA1 or BRCA2 and no mutations.					
Blinding	Not relevant to the study as comparisons made were made between different groups based on presence/absence of mutation. Additionally, the data analysis team used de-identified patient data and were not involved in patient recruitment or data collection.					

Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed is relevant 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 & experimental systems		Methods			
Involved in the study	n/a	Involved in the study			
Antibodies	\boxtimes	ChIP-seq			
Eukaryotic cell lines	\boxtimes	Flow cytometry			
Palaeontology and archaeology	\boxtimes	MRI-based neuroimaging			
Animals and other organisms					
Human research participants					
Clinical data					
Dual use research of concern					
	Involved in the study Antibodies Eukaryotic cell lines Palaeontology and archaeology Animals and other organisms Human research participants Clinical data	Involved in the study Antibodies Eukaryotic cell lines Palaeontology and archaeology Animals and other organisms Human research participants Clinical data			

Antibodies

Antibodies used

Antibodies used were anti-CD3 (clone 2GV6, predilute; Ventana Medical Systems), anti-CD4 (clone SD35, predilute; Ventana Medical Systems), anti-CD8 (clone SD57, predilute; Ventana Medical Systems) and anti-PD-1 (clone SP263, predilute; Ventana Medical Systems).

Validation

All antibodies used are commercially available and validated by Roche Diagnostics for in vitro diagnostic (IVD) use in sections of normal and neoplastic human tissues, as listed on the manufacturer's website.

Human research participants

Policy information about studies involving human research participants

Population characteristics

The study population was female, Malaysian breast cancer patients seeking treatment at the Subang Jaya Medical Centre, a private hospital within the Kuala Lumpur metropolitan area.

Recruitment

Patients were recruited sequentially from a hospital-based cohort. The hospital in point, Subang Jaya Medical Centre, is a private Malaysian hospital within the Kuala Lumpur metropolitan area, where patients tend to be of higher socio-economic

status relative to the general population.

Ethics oversight

The project was reviewed and approved by the Independent Ethics Committee, Ramsay Sime Darby Health Care (reference no: 201208.1), and written informed consent was given by each individual patient.

Note that full information on the approval of the study protocol must also be provided in the manuscript.