#### The Lancet Commission for Breast Cancer is launched

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Breast cancer is an increasing global health, gender, socioeconomic and equity challenge. In 2020, 2.3 million women were diagnosed with breast cancer and there were 685,000 deaths worldwide<sup>1</sup>. Not only is it the highest incident cancer globally but is also the most prevalent, causing more disability-adjusted life years lost than any other malignancy<sup>1</sup>. As such, tackling breast cancer is a formidable task for healthcare systems, policy makers and many other stakeholders. The numbers of people with metastatic breast cancer who go uncounted are poignant. Cancer registries record patients initially presenting with *de novo* metastatic breast cancer, but information on those developing metastases after a diagnosis of early breast cancer is virtually non-existent<sup>2</sup>. In a world focussed on breast cancer cure, these "uncounted" people living with metastatic disease face abandonment and stigma<sup>3</sup>.

With only 0.5-1% of breast cancers occurring in men, this disease disproportionally affects women whose premature disability and deaths have immediate and devastating effects on entire families, especially children<sup>4,5</sup>. There are far-reaching consequences on society and the economy given the pivotal role of the female labour force globally, whether paid or often unpaid, in caregiving and community leadership roles<sup>6</sup>. Given the medical costs, care-giving burden and tragic loss of life, breast cancer is financially damaging to families. Financial toxicity is intersectional and intricately linked to empowerment: women often rely on partners for financial support to access care and can face abandonment at diagnosis or during treatment and rejection by their families due to high out of pocket care costs. This is compounded by cultural barriers including cancer myths, stigma, and fatalism, causing workplace discrimination, community neglect and psychological trauma.

Perhaps the most difficult issue to tackle is inequity. Access to early detection, comprehensive diagnostic work-up, and timely high-quality breast cancer management depend on who you are and where you live<sup>7</sup>. There is a strongly positive and unacceptably steep relationship between 5-year survival rates and country income<sup>8</sup>. Whilst there are stark inequities comparing high, middle and lower income countries, there are also inequities *within* countries with intersectionality between race and poverty<sup>8</sup>. How can it be acceptable for the survival estimates of some black African women to be as low, or lower, than women treated in the USA nearly a century ago?<sup>9</sup> Or as a UK study of women with screen-detected breast cancer showed, why is the excess hazard ratio of death more than 50% higher for those with social deprivation after adjusting for individual, tumour, treatment factors, lead-time bias and over-diagnosis?<sup>10</sup>

Against this background, there have been dramatic improvements in breast cancer management over the last 4 decades, resulting in falling mortality rates in mostly higher income countries. Extraordinary advances in genomics are rapidly unravelling the complex breast cancer biology leading to new strategies in personalised cancer management, enabling right sizing extent of surgery, radiation and systemic treatment to optimise oncologic outcomes and minimise collateral damage, Globally, we are at pivotal point whereby either the rapid progress in breast cancer medicine/science is capitalised to ensure equitable access for all *or* the current inequalities between and within countries widen into a chasm.

The Lancet Breast Cancer Commission is timely: its over-arching aim is to present a compelling forward-thinking story to influence global policy and ultimately improve lives for those affected by breast cancer wherever they live. The Commission unites a multidisciplinary international team of leaders to present a bold key message, under-pinned with high-quality evidence-based recommendations. Proposals for transformative change will be actionable with key metrics to measure success. The Commission represents high/low/middle-income countries from 6 continents and includes patient advocate commissioners with lived experience of early and metastatic breast cancer to ensure a person-centred approach. Financial support for 1 in-person 2-day meeting in Cambridge, UK and reimbursement for patient advocates has been provided by Breast Cancer Now.

The first commission meeting identified 5 workstreams: closing the global gap in breast cancer survival; optimising prevention and early detection; effective use of personalised breast cancer management; enabling safe de-escalation of treatment; and improving lives for patients with metastatic breast cancer. Paired workstream co-chairs will co-ordinate the commissioners and collaborators in scoping and evidence synthesis to generate material with assistance from early career researchers (See Appendix 1 for list of those involved). Twelve key questions were agreed (see Panel 1) at the second meeting and a workplan developed defining specific tasks to investigate, describe and ultimately make actionable recommendations regarding these themes.

The commission has linked with other key commissions including Cancer and Health Systems and Women and Cancer, WHO's Global Breast Cancer Initiative and ABC Global Alliance<sup>6,11,12,13</sup>. This collaborative and synergistic approach has highlighted that breast cancer is not only a test case for other cancers, but also a model for other chronic diseases within health care systems<sup>11</sup>. Breast cancer illustrates the far-reaching influence of women in society<sup>6</sup>. A collaboration with commissioners from the Palliative Care and Pain Relief commission will also explore the value to patients and society of reducing the avoidable pain and suffering from breast cancer<sup>14</sup>. The final commission publication is planned for Autumn 2023.

In summary, this Lancet Breast Cancer Commission will investigate key biomedical topics and explore cross-cutting social, behavioural, economic, and ethical themes. As such, it is hoped that this truly person-centred approach will provide impetus to not only improve breast cancer management but for breast cancer to become an icon for wider positive culture change. This commission has a sense of urgency; the time to act is now!

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BOA is a staff member of WHO at the time of the writing of this manuscript. The authors alone are responsible for the views expressed in this article as they do not necessarily represent the decisions, policies, or views of WHO.

#### References

- 1. https://www.who.int/news-room/fact-sheets/detail/breast-cancer (accessed Dec 20, 2021).
- 2. Mailhot Vega RB, Balogun OD, Ishaq OF, Bray F, Ginsburg O, Formenti SC. Estimating child mortality associated with maternal mortality from breast and cervical cancer. Cancer. 2019 Jan 1;125(1):109-117.
- 3. Mariotto AB, Etzioni R, Hurlbert M, Penberthy L, Mayer M. Estimation of the number of women living with metastatic breast cancer in the United States. Cancer Epidemiol Biomarkers Prev. 2017 Jun;26(6):809-815.
- 4. Cardoso F, Spence D, Mertz S, Corneliussen-James D, Sabelko K, Gralow J, Cardoso MJ, Peccatori F, Paonessa D, Benares A, Sakurai N, Beishon M, Barker SJ, Mayer M. Global analysis of advanced/metastatic breast cancer: Decade report (2005-2015) Breast. 2018 Jun;39:131-138.
- 5. Galukande M, Schuz J, Anderson BO, et al. Maternally Orphaned Children and Intergenerational Concerns Associated With Breast Cancer Deaths Among Women in Sub-Saharan Africa. *JAMA Oncol* 2021; **7**(2): 285-9.
- 6. Ginsburg O, et al. The global burden of women's cancers: a grand challenge in global health. Lancet. 2017.
- 7. Farmer P, Frenk J, Knaul FM, et al. Expansion of cancer care and control in countries of low and middle income: a call to action. Lancet 2010;376: 1186–93.
- 8. American Cancer Society. Cancer Facts & Figures for African Americans 2019-2021. 2019. <a href="https://www.cancer.org/content/dam/cancer-org/research/cancer-facts-and-statistics/cancer-facts-and-figures-for-african-americans-2019-2021.pdf">https://www.cancer.org/content/dam/cancer-org/research/cancer-facts-and-statistics/cancer-facts-and-figures-for-african-americans-2019-2021.pdf</a> (accessed Dec 20, 2021).
- 9. McCormack V, McKenzie F, Foerster M, Zietsman A, Galukande M, Adisa C, Anele A, Parham G, Pinder LF, Cubasch H, Joffe M, Beaney T, Quaresma M, Togawa K, Abedi-Ardekani B, Anderson BO, Schüz J, Dos-Santos-Silva I. Breast cancer survival and survival gap apportionment in sub-Saharan Africa (ABC-DO): a prospective cohort study. Lancet Glob Health. 2020 Sep;8(9):e1203-e1212.
- 10. Morris M, Woods LM, Rachet B. What might explain deprivation-specific differences in the excess hazard of breast cancer death amongst screen-detected women? Analysis of patients diagnosed in the West Midlands region of England from 1989 to 2011. Oncotarget. 2016 Aug 2;7(31):49939-49947.
- 11. Knaul FM, Garcia PJ, Gospodarowicz M, Essue BM, Lee N, Horton R. The Lancet Commission on cancer and health systems: harnessing synergies to achieve solutions. Lancet. 2021 Sep 25;398(10306):1114-1116
- 12. Anderson BO, Ilbawi AM, Fidarova E, Weiderpass E, Stevens L, Abdel-Wahab M, Mikkelsen B. The Global Breast Cancer Initiative: a strategic collaboration to strengthen health care for non-communicable diseases. Lancet Oncol. 2021 May;22(5):578-581.
- 13. Fallowfield L, Boyle FM, Travado L, Kiely BE, Jewell P, Aubel D, Cardoso F. Gaps in Care and Support for Patients With Advanced Breast Cancer: A Report From the Advanced Breast Cancer Global Alliance. JCO Glob Oncol. 2021 Jun;7:976-984
- 14. Knaul FM, Farmer PE, Krakauer EL, De Lima L, Bhadelia A, Jiang Kwete X, Arreola-Ornelas H, Gómez-Dantés O, Rodriguez NM, Alleyne GAO, Connor SR, Hunter DJ, Lohman D, Radbruch L, Del Rocío Sáenz Madrigal M, Atun R, Foley KM, Frenk J, Jamison DT, Rajagopal MR; Lancet Commission on Palliative Care and Pain Relief Study Group. Alleviating the access abyss in palliative care and pain relief-an imperative of universal health coverage: the Lancet Commission report. Lancet. 2018 Apr 7;391(10128):1391-1454.

#### Panel 1: Key questions identified by the breast cancer commission

1. How do we change the mindset that it is inevitable and therefore acceptable for 1 in 8 women to develop breast cancer during their lifetime and how do we reverse this increasing trend?

- 2. What lessons can we learn from the COVID-19 global response and how can we apply these to tackling breast cancer worldwide to move forward together in solidarity?
- 3. What can we learn from patient/public advocacy movements as a powerful & effective mechanism in other disease/health issues in order to close the global breast cancer gap?
- 4. What is the impact of stage shift to earlier breast cancer presentation on global survival rates
- 5. Financial toxicity and health-related suffering: what are the wider consequences of inaction beyond immediate healthcare costs and how can we quantify the value to patients and society of reducing the avoidable pain and suffering of breast cancer?
- 6. How can we introduce an aspirational goal of systematic risk assessment and precision breast screening and prevention for young women as part of routine broader health care?
- 7. How can we enable personalised breast cancer management to become universally applicable?
- 8. How can we transition from traditional siloed care to integrated patient-centred management, ensuring that all breast cancer patients (early and metastatic) have access to multidisciplinary specialised care?
- 9. How can patient choice in their breast cancer management be empowered through a holistic benefit-risk approach and shared decision making?
- 10. How do we design innovative clinical trials to test safe reduction in *overall* patient burden of treatment/management?
- 11. How can we quantify the hidden/forgotten global population with metastatic breast cancer?
- 12. How can we change the mindset around metastatic breast cancer from a rapidly fatal disease to potentially curable?

## Appendix 1: Lancet Breast Cancer commission commissioners, collaborators, early career researchers and other stakeholders

#### **Commissioners**

Jean Abraham, medical oncologist, UK

Ben Anderson, breast surgeon, USA (WHO Medical Officer leading Global Breast Cancer Initiative)

Fabrice André, medical oncologist, France

Carlos Barrios, medical oncologist, Brazil

Judith Bliss, statistician and methodologist, UK

Judy Boughey, breast surgeon, USA

David Cameron, medical oncologist, UK (Chair of Breast International Group: BIG against breast cancer)

Fatima Cardoso, medical oncologist, Portugal (President of the ABC Global Alliance)

Lisa Carey, medical oncologist, USA

Sanjoy Chatterjee, clinical oncologist, India

Charlotte Coles, clinical oncologist, UK (Chair of Lancet Breast Cancer Commission)

Jack Cuzick, epidemiologist, UK

Dorothy Du Plooy, nurse and midwife, South Africa

Prue Francis, medical oncologist, Australia

Julie Gralow, medical oncologist, USA

Reshma Jagsi, radiation oncologist, USA

Felicia Knaul, health economist and patient advocate, Mexico (Co-chair of Lancet Healthcare Systems and Cancer Commission)

Fei Ma, medical oncologist, China

Ritse Mann, radiologist, Netherlands

Stuart McIntosh, breast surgeon, UK

Shirley Mertz, patient advocate, USA

Miriam Mutebi, breast surgeon, Kenya (Linking Commissioner for Lancet Women and Cancer Commission)

Funmi Olopade, medical oncologist, USA

Kelly-Anne Phillips, medical oncologist, Australia

Philip Poortmans, radiation oncologist, Belgium

Tanja Spanic, patient advocate, Slovenia

Dingle Spence, palliative medicine physician, Jamaica

Hilary Stobart, patient advocate, UK

Fraser Symmans, pathologist, USA

Cynthia Villarreal-Garza, medical oncologist, Mexico

Cheng-Har Yip, breast surgeon, Malaysia

## **Early Career Researchers**

Maya Bienz, UK

Lynsey Drewett, UK

Alex Fulton, UK

Dharrnesha Inbah Rajah, USA

Farasat Kazmi, UK

Mareike Thompson, UK

Jeffrey Rubasingham, UK

Valentina Vargas, USA

#### **Collaborators**

Hector Arreola, economist, Mexico

Raj Badwe, surgical oncologist, India

Rajiv Dave, breast surgeon, UK

Ruth Etzioni, biostatistician, USA

Indraneel Mittra, surgical oncology, India

Patricia Moreno, clinical psychologist, USA

Renu Sara Nargund, research associate, USA

Cara Noble, patient advocate, South Africa

Robert Smith, epidemiologist, USA

Didier Verhoeven, medical oncologist, Belgium

Jiani Wang, research associate, China

## **Administrators**

Catherine Durance, UK

Jane Sales, UK

## **Editor**

Naomi Lee (The Lancet), UK (Editor for Lancet Breast Cancer Commission)

#### Funders

Simon Vincent, Breast Cancer Now, UK

Bunia Gorelick, Breast Cancer Now, UK

#### Other stakeholders

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