

OHSL Breast Cancer Data Alliance

24.5% of all cancers.

Asia has 45.4% of these.

Asian female under 45 is 52.9%.

→ Mortality rate for this group is 51.4%.

US: 34.9% , 45.6% for under 50 years.

More stats on
ohsl.us.

{ Big Data - Large & diverse collection of data that grows over time.
Precision Medicine - Leverages big data for medical predictions.

→ For medicinal purposes, got from insurance claims, cancer registries & electronic health records

* Some ex. of big data used in breast cancer related precision:-

- MammaPrint {assesses activity of certain genes to determine risk of recurrence}
- Oncotype DX {similar to MammaPrint}
- PAM50-based Prosigna {Also calculates risk of Recurrence}
- EndoPredict {Also calc distant recurrence within 10 yrs.}

* BCDA's goal is to bring clinical & non-clinical professionals together to identify key parameters, which could explain current trends in BC.

* OHSL is trying to create a data catalog. Collab with Alation to use their tech for this.
↳ Data management platform for corpora.

"The Data Catalog provides users with a simple, searchable & unified resource that enables data discovery, no matter the source's original loc".

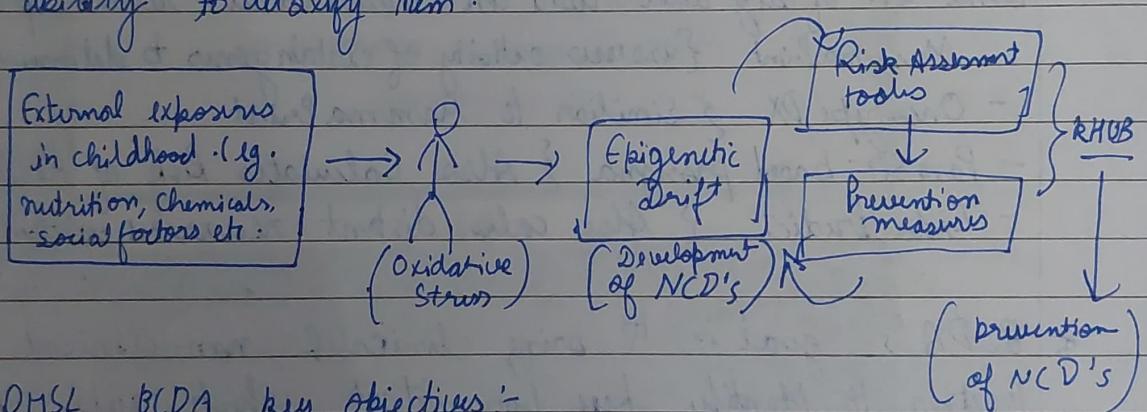
* Focusing on identifying unique features for region specific breast cancer.

OHS L has 3 main goals with BCDA:-

- implement personalized preventive interventions in pre-adolescent
- develop a multidata knowledge hub (RHUB) included with multidata catalog to identify BC risk factors & make evidence-based recommendations
- validate biomarkers of Oxidative Stress ~~mediated epigenetic impact~~ & breast cancer risk & measure the import of intervention in a preadolescent population

✓
chemical
modific.
to genes.

→ imbalance b/w productⁿ of complex oxygen species & body's ability to detoxify them.



* OHS L BCDA key objectives:-

- ↳ Data Catalog Creation
- ↳ Data Classification & Analytics
- ↳ Focus on L.M.I.C.'s. (Lower-Middle Income Countries)
- ↳ Primary Prevention

→ Normally help protect us from certain cancers.

READING REVIEW

Bunk Paper - The Social Notebook

Breast Cancer: An up-to-date review & future perspectives

- Ruaxi Hong | Binghe Xu.

- When doctors treat breast cancer, they consider:

→ specific molecules involved (like HER 2, estrogen & progesterone receptors)

→ gene mutations like PRCA1/2.

→ how immune system interacts with cancer.

- For early stage cancer, treatments like surgery & radiations are often used to remove or destroy cancer. Doctors might give medicine before surgery to shrink the tumor, which makes it easier to remove.

- For advanced cancers (like adv. BC), it is much harder to cure; but treatments like hormone therapy, targeted drugs & sometimes immunotherapy can help slow down cancer's growth.

- Doctors give tailored prescriptions to each person's specific cancer. (less or aggressive treatments)

↳ using precision medicine

↳ In BC, current treatments are based on molecular subtyping, allowing for tailored therapies. Treatments can become even more individualized, with therapy adjusted acc. to tumor biology and early predictive markers. + Further classification of BC subtypes.

→ 2 Significant Questions Remain:-

↳ whether breast surgery can be avoided in patients achieving a complete pathological response (pCR) after neoadjuvant therapy.

↳ Shrinking a tumor. { incl. chemo, radiation, hormone therapy. }

↳ If some patients can skip axillary surgery for staging & treatment.

Reserving lymph node ← from axilla area.

Mammography :- X-ray imaging method used to imagine the breast for early detection of cancer & other breast diseases.

DCIS: Ductal Carcinoma in-Situ is a condition that affects milk ducts in the breast. Bunk Page The Social Notebook

The Breast Cancer Landscape

- Department of Defense Breast Cancer Research Program.

Breast Cancer Incidence :-

↳ Likelihood of a woman being diagnosed with breast cancer has increased from 1 in 11 in 1975 to 1 in 8 today.

partly because more women are reaching older ages.

↳ But, could be due to screening practices & use of hormone replacement therapy. ↳ like mammography. ↳ changes in.

↳ Introduction of mammography has led to increase in DCIS. DCIS went from 2% (1985) → 33% now.]

(about 31% are considered overdiagnosed.) About overdiagnosis & overtreatment.

Breast Cancer Deaths :-

* Most BC Deaths occur because the cancer spreads to other parts of the body, affecting vital organs like lungs, liver & brain.

- While breast cancer incidence varies globally due to screening practices, mortality rates are consistent across regions.

Risk Factors :-

(≥ 4 times risk)	(\uparrow Risk 2x-4x)	(\uparrow Risk up to 2x)
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High-Risk Factors

Moderate R.F

Modest R.F

- Age (65+ versus <65 yrs)
- Prior diagnosis of DCIS.
- Alcohol Consumption.
- Presence of pathogenic genetic variations (BRCA1, BRCA2, PALB2, TP53)
- High endogenous hormone levels (premenopausal)
- Early Menarche (<11 yrs)
- Late age first pregnancy
- Late Menopause
- Never breastfed a child
- Diagnosis of atypical hyperplasia &/or lobular C-I-S.
- Mammographically dense breast.
- Having 2 or more relatives with BC.
- No full-term pregnancy
- Obesity & physical inactivity
- Tall height.

Receptors in or on cells that can attach to certain substances in blood.

ER - Estrogen Receptor
PR - Progesterone Receptor
HER 2 - Human epidermal growth factor receptor 2

- Still not all breast cancer cases can be attributed to these risk factors. Some factors like age, fam. history, & genetic mutations are not modifiable.
- Radiation exposure especially during childhood is a well established risk.
- ~~Risk factors~~ Risk factors may vary in their relationships to different molecular subtypes.

Heterogeneity :- Heterogeneity :-

Subtypes of BC :-

- ↳ Luminal A :- +ve for either ER or PR, but -ve for HER2
- ↳ Luminal B :- Similar to Luminal A but higher proliferation ratio & maybe ~~not~~ HER2-positive.
- ↳ HER2-expressing :- -ve for ER or PR, but +ve for HER2
- ↳ Basal Like (Triple Negative) :- Negative for ER, PR & HER2.

Distribution of these varies by factors like age, race, ethnicity etc.

- No targeted therapies are available for patients with triple negative BC.

Recurrence :-

- Around 10 to 30% of women may experience recurrence.
- Metastatic BC accounts for 90% of BC related deaths.
- Treatments are aimed at slowing growth instead of termination.

* Different BC therapies include chemo, hormonal, radiation therapy. There are less invasive surgeries like lymphectomy with radiatⁿ therapy, leading to improved QoL but no change in mortality rates. Newer treatments have increased the number of toxicities.

Metastatic BC :- { spread of cancer from places where it formed for the first time to new organs.

Histopathology: Diagnosis & study of diseases in tissue

(B.T) Benign tumor has distinct, smooth, regular borders

(M.T) Malignant tumor has irregular borders & grows faster than B.T.

Bunk Paper - The Social Notebook

Artificial Intelligence in Cancer Research

- Bhavneet Phinder et al.

Shows the transformative impact of AI on oncology.

1. Introduction to AI in Cancer Research

- AI encompasses a broad range of computational techniques that enable machines to mimic human intelligence.
- Deep learning (DL), is a subset of AI, involves neural networks with multiple layers. (hence "deep"), capable of learning from vast amounts of data.
- AI's ability of pattern recognition makes it a powerful tool in oncology.

2. Applications of AI in Oncology

2.1. Detection & Classification of Cancer

* DNN's excel in classifying images such as H&E stained multi-slide images (WSI's) of tissue samples.

* HISTOPATHOLOGY & IMAGING

- Convolution Neural Networks (CNNs), have been successful in classifying medical images, such as histopathology slides, with high accuracy.
- Deep PATH system uses CNNs to classify lung cancer subtypes with a high accuracy outperforming traditional methods.
- AI models can distinguish b/w malignant & benign tissues in breast biopsies, aiding pathologists in diagnosis.

TME is the ecosystem surrounding a tumor.

Inception - V3 architecture has been used to classify lung cancer subtypes with AUC of 0.97

- The Social Notebook

Used DNN to quantify lymphocyte infiltration.

2.2 Molecular Characterization of Tumors :-

- Tumor Microenvironment (TME) :-

> AI helps in analyzing the TME, which includes various cell types & molecular signals that interact with tumor cells. [Learning Model by Saltz et al.]

Study of genetic markers

- Genomic & Proteomic analysis:-

identification of all proteins. > AI can generate integrate data from genomics, proteomics & other omics technologies to identify biomarkers & subtypes of cancer.

characterization of normal or abnormal tissues.

> This aids in personalized medicine, where treatments are tailored based on the molecular characteristics of the individual's tumor.

2.3 Drug Discovery & Repurposing :-

- Accelerating Drug Development :-

> AI algos analyze biological data to identify potential drug targets & predict the efficacy of new compounds.

> Ex:- DrugCell model integrates cellular & drug response data to predict how cancer cells will respond to different therapies, aiding in discovery of new treatment.

- Drug Repurposing :-

> AI can identify existing drugs that might be effective against cancer by analyzing molecular effects & matching them with cancer cell vulnerabilities. This shortens time & cost consumption.

combination of histological stains:
hematoxylin & Eosin.

Immunotherapy:- Using persons own immune system
to fight cancer.

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2.4 Predicting Treatment Outcomes

- Personalized Treatment Plans :-

- > AI models predict how patients will respond to various treatments, helping to tailor therapies to individual patients.
- > Example: A study by Durie et al. used ML to analyze CT scan features & predict lung cancer patients' responses to immunotherapy.

- Survival Predictions :-

- > AI can analyze patient data, including medical images & clinical records, to predict survival outcomes & guide treatment decisions.
- > This helps in managing patient expectations & planning appropriate interventions.

3. Challenges & Future Directions :-

3.1. Data Quality & Availability }

- Large, Annotated Datasets.

- > AI models require large datasets with accurate annotations for training. Access to high-quality data is often an bottleneck.

- Data Privacy & Security.

- > Ensuring patient data privacy & security while sharing data for AI research is crucial.

This is basically what OHSL-BCDA project is aimed to contribute via a comprehensive catalog.

P.T.O

we can't trace the systems
thought - process.

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3.2 Model Interpretability:-

- Understanding AI decisions:-

> Deep learning models have a 'black-box' nature, making it difficult to interpret their decisions.

- Regulatory & Clinical Acceptance:-

> AI models must be interpretable & comply with regulatory standards.

3.3 Continuous Learning & Drift:-

enabling detailed studies of cell interaction within the TME.

- Concept Drift:

> AI models need to be updated continuously as new data becomes available & medical practices evolve.

- Maintaining Performance:-

> Ensuring models retain performance over time requires validation & retraining.

4. Technological Advancements:-

4.1 Advanced Imaging Technologies:-

MAI:-
Magnetic Resonance Imaging.

> Combining AI with advanced imaging tech like multi-modal-imaging [e.g. combining MRI & PET scans] can improve diagnostic accuracy & provide more comprehensive insight into tumor biology.

PET:- Positron Emission Tomography

P.E.T.O

> Detailed image of inner structures.

< Helps reveal metabolic or biochemical functions.

4.2 Integration Approaches:-

- > Integrating data from various sources can provide a holistic view of the patient's condition & improve decision-making.

5. Conclusion.

- Transformative Potential :-

- > AI has potential to revolutionize oncology by improving diagnostic accuracy, personalizing treatment, accelerating drug discovery & predicting patient outcomes.
- > Overcoming current challenges related to data quality, model interpretability & continuous learning is essential for fully realizing AI's benefits in cancer care.

What
BCDA
is aimed
at.

- Future Outlook :-

- > Future looks promising with ongoing research & tech. advancements helping to further enhance its applic'n & integrat'n into clinical proc.
- > Collab. b/w researchers, clinicians & regulatory bodies will be crucial in driving successful implementation of AI in oncology.