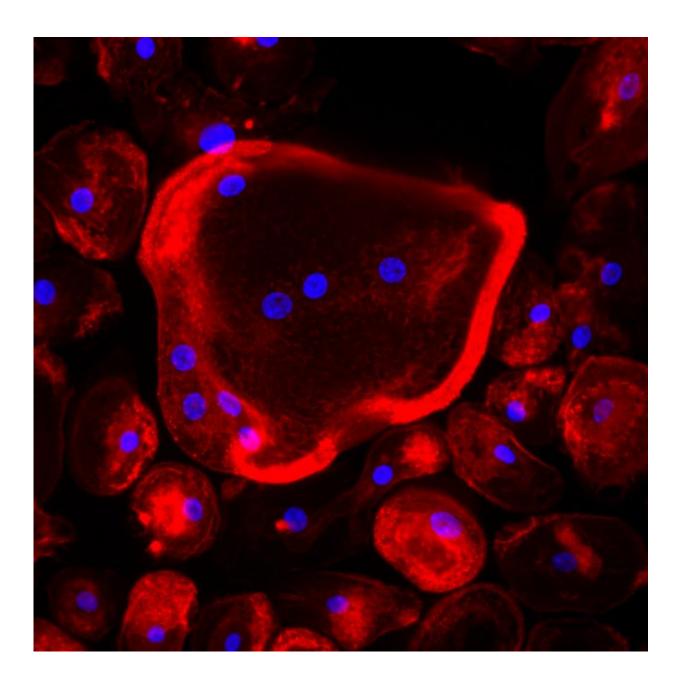


## Scientists identify key to preventing secondary cancers

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Immunofluorescence image is of multinucleate osteoclast generated by exposure of pre-osteoclast cells to lysyl oxidase (Red = cell body phalloidin, blue = DNA). Credit: Dr Alison Gartland:University of Sheffield

Leading scientists from the University of Sheffield and University of Copenhagen have identified a possible key to preventing secondary cancers in breast cancer patients, after discovering an enzyme which enhances the spread of the disease.

Secondary (metastatic) breast cancer is the main cause of the 12,000 deaths which occur from breast cancer in the UK every year.

The most common site for the disease to spread is the <u>bone</u> - occurring in around 85 per cent of <u>secondary breast cancer</u> patients.

The new research, which was conducted at The Institute of Cancer Research, London, found that the enzyme LysYl Oxidase (LOX) released from the primary tumour causes holes in bone and prepares the bone for the future arrival of cancer cells.

The findings suggest that identifying LOX in oestrogen receptor negative (ER negative) <u>breast cancer patients</u> early, could allow doctors to block the enzyme's activity, preventing <u>bone damage</u> and the spread of tumour cells to the bone (metastasis), halting the progression of the disease.

The researchers also showed that treatment with bisphosphonate, an existing class of drug which prevents the loss of <u>bone mass</u> and is already used to treat diseases such as osteoporosis, was able to prevent the changes in the bone and the spread of the disease in mice.

The pioneering research, co-led by Dr Alison Gartland at the University



of Sheffield's Department of Human Metabolism, could lead to a better prognosis for <u>cancer patients</u> in the longer term.

Dr Gartland said: "This is important progress in the fight against <u>breast</u> <u>cancer metastasis</u> and these findings could lead to new treatments to stop secondary breast tumours growing in the bone, increasing the chances of survival for thousands of patients.

"We are really excited about our results that show breast cancer tumours send out signals to destroy the bone before cancer cells get there in order to prepare the bone for the cancer cells' arrival.

"The next step is to find out exactly how the tumour secreted LOX interacts with <u>bone cells</u> to be able to develop new drugs to stop the formation of the <u>bone lesions</u> and cancer metastasis. This could also have implications for how we treat other bone diseases too."

Study co-leader Dr Janine Erler, formerly Team Leader in Cancer Biology at The Institute of Cancer Research, London, who now is Associate Professor at the Biotech Research & Innovation Centre (BRIC) at the University of Copenhagen, said: "Once cancer spreads to the bone it is very difficult to treat. Our research has shed light on the way breast <u>cancer cells</u> prime the bone so it is ready for their arrival.

"If we were able to block this process and translate our work to the clinic, we could stop breast cancer in its tracks thereby extending patients' lives."

The research, funded by Breast Cancer Campaign, Cancer Research UK, Novo nordisk foundation, Danish cancer society, lundbeck foundation, and both universities, is published today (27 May 2015) in the journal *Nature*.



Katherine Woods, Senior Research Communications Manager at Breast Cancer Campaign and Breakthrough Breast Cancer, said: "By unveiling the role that the protein LOX is playing, these results open up a whole new avenue for research and treatments that could stop breast cancer spreading to the bone. The research also adds weight to the growing body of evidence supporting the role of bisphosphonates in stopping secondary breast cancer in its tracks.

"The reality of living with secondary breast cancer in the bone is a stark one, which leaves many women with bone pain and fractures that need extensive surgery just when they need to be making the most of the time they have left with friends and family."

She added: "Secondary breast cancer kills 1,000 women each and every month in the UK alone and yet we still don't know enough about how and why breast cancer spreads to stop it.

"Our newly-formed charity is determined that by 2050, no one will lose their life to <u>breast cancer</u> and we'll do this by ramping up our research efforts, in this area in particular, doing everything possible to achieve that goal."

## More information: Nature, DOI: 10.1038/nature14492

## Provided by University of Sheffield

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