

Prostate Cancer Donor Program

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INTRODUCTION

- Prostate cancer is the most commonly diagnosed cancer in Australia.
- Effective, preclinical research relies on robust and accurate models of prostate cancer.
- Patient derived xenografts (PDXs) are human tumour samples grown in mice and allow preclinical studies on prostate cancers similar to those seen in clinical practice.
- Patients die from prostate cancer that has developed resistance to all available treatments.
- PDXs from resistant tumours need to be created to accurately test new therapeutic options to treat prostate cancer.
- Patients often express a desire to be organ donors after death but are unable to due to their cancer diagnosis.
- An alternative is to donate tumour tissue to research.

Aims

- To assess the feasibility of collecting cancer samples with a biopsy needle at the bedside from prostate cancer patients immediately after death; and
- Use these samples to create serially-transplantable PDX models.

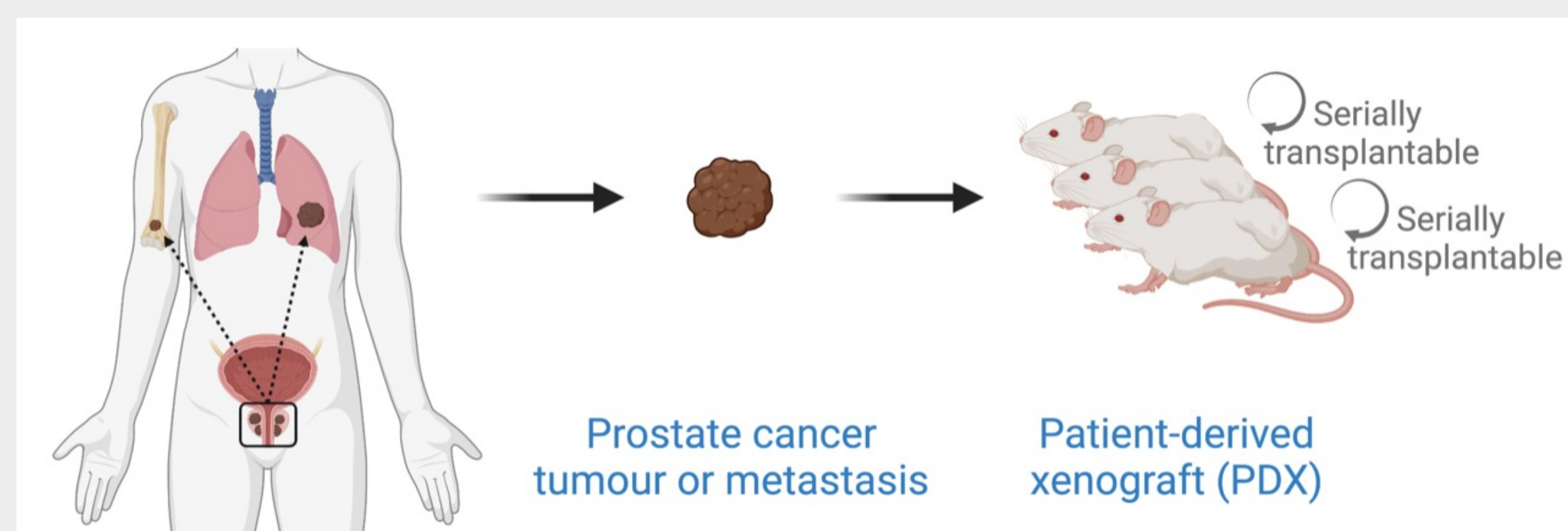
METHODS

SAMPLE COLLECTION

- This pilot study is enrolling prostate cancer patients likely to die in palliative care.
- Eligibility Criteria: 1) Over 18 years of age, 2) confirmed metastatic prostate cancer, 3) lesions amenable to core needle biopsy or bone marrow trephine and 4) not serologically positive for HIV and Hepatitis B and C.
- Patients were recruited by one of their treating clinicians.
- The patient and their next of kin signed the consent form and were made aware that collection may not be possible.
- Sites of disease were identified on previous imaging.
- The research team were notified when the patient died and arrived at the bedside within six hours of death.
- Bone lesions were biopsied using a Jamshidi 11G Bone Marrow Biopsy/Aspiration needle.
- Soft tissue lesions were biopsied using a Bard Mission 14G Disposable Core Biopsy Instrument Kit.
- Samples were placed into MACS Tissue Storage Solution and kept on ice while transported to the laboratory at Monash University.
- A follow-up phone call was made to the next of kin two weeks after collection to obtain feedback from the family.
- Clinical information was collected from medical records.

XENOGRAFTING

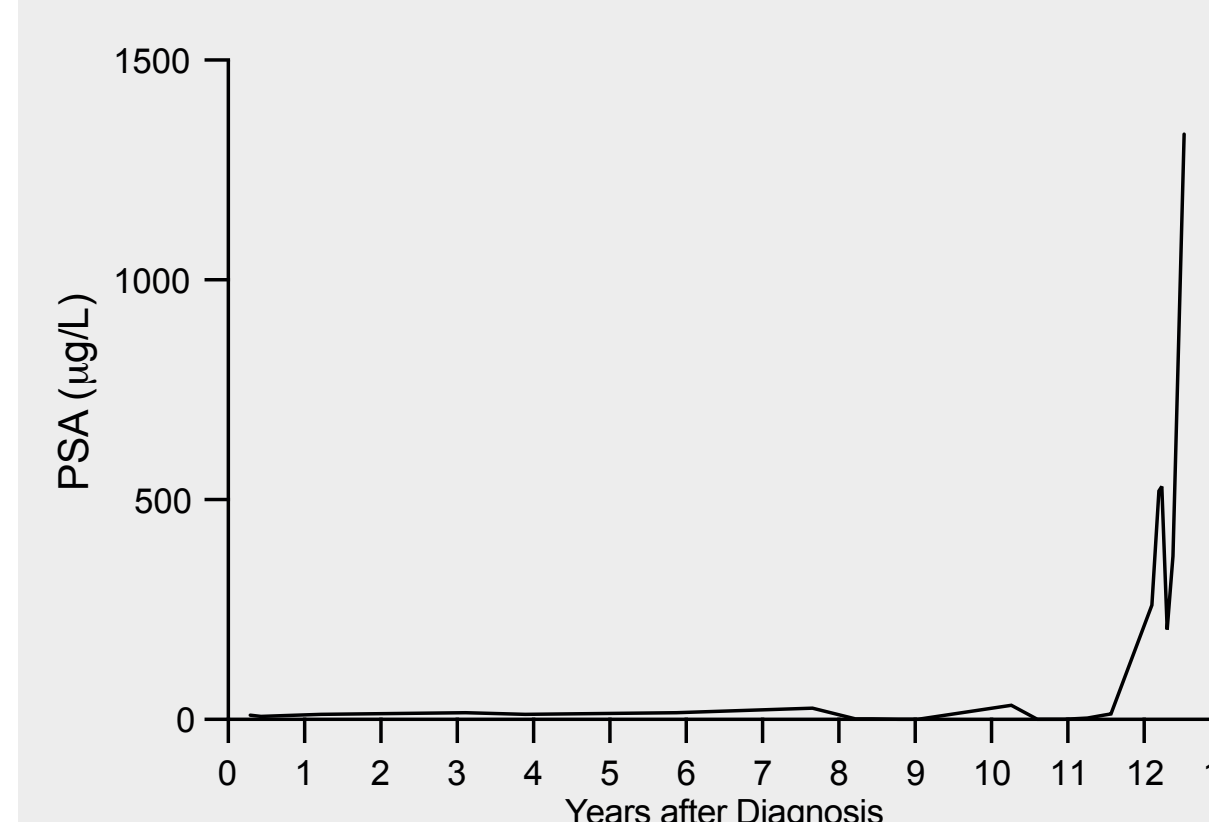
- Patient tissues were grafted into immunocompromised male NSG mice with testosterone implants.
- Samples were grafted under the renal capsule to maximise engraftment rates.
- Grafts will be monitored for up to 12 months to assess *in vivo* tumour growth.
- Actively growing tumours will be regrafted into additional mice to establish serially-transplantable PDXs and will add to the MURAL collection.



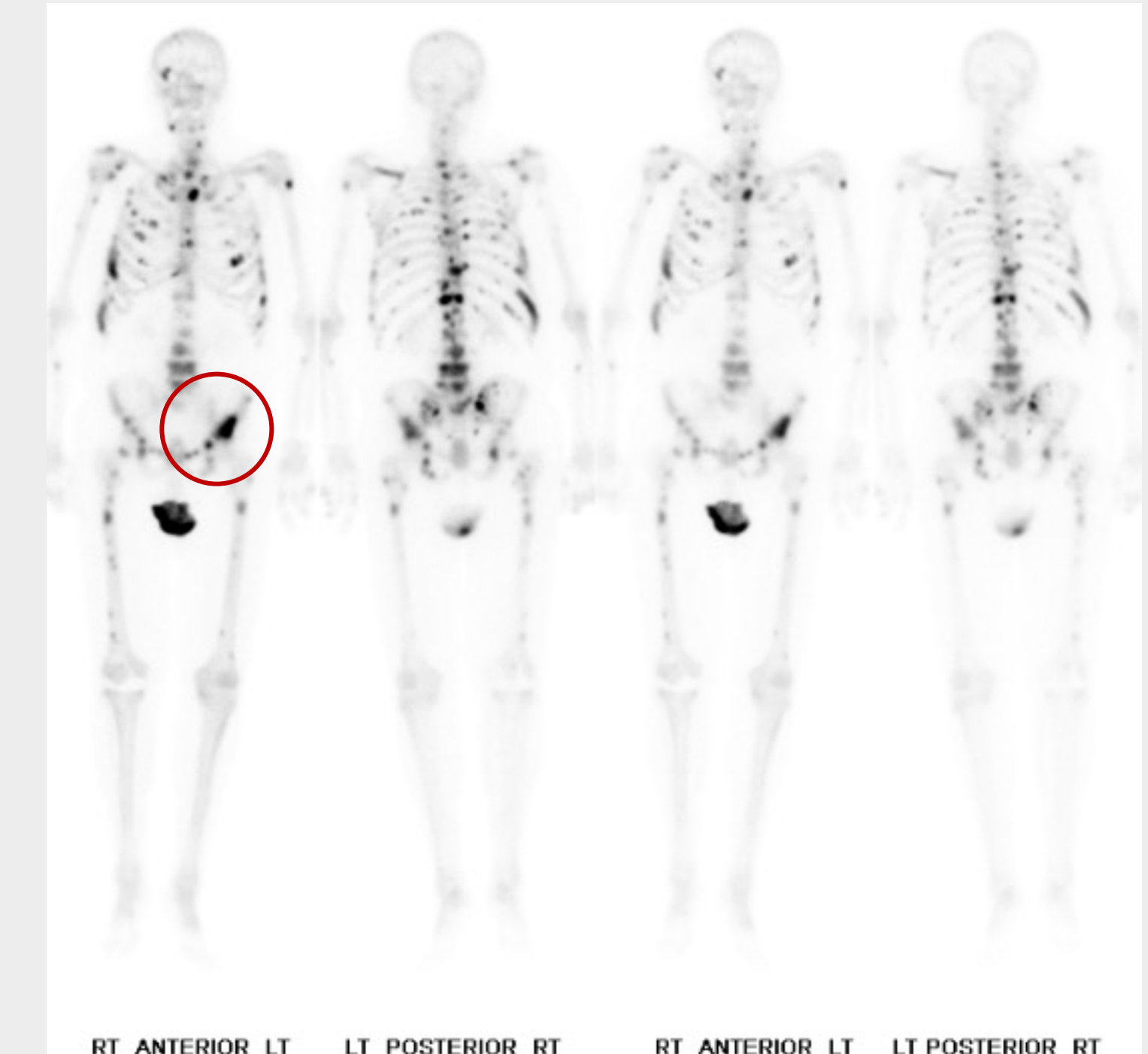
RESULTS

Patient 1 (540D)

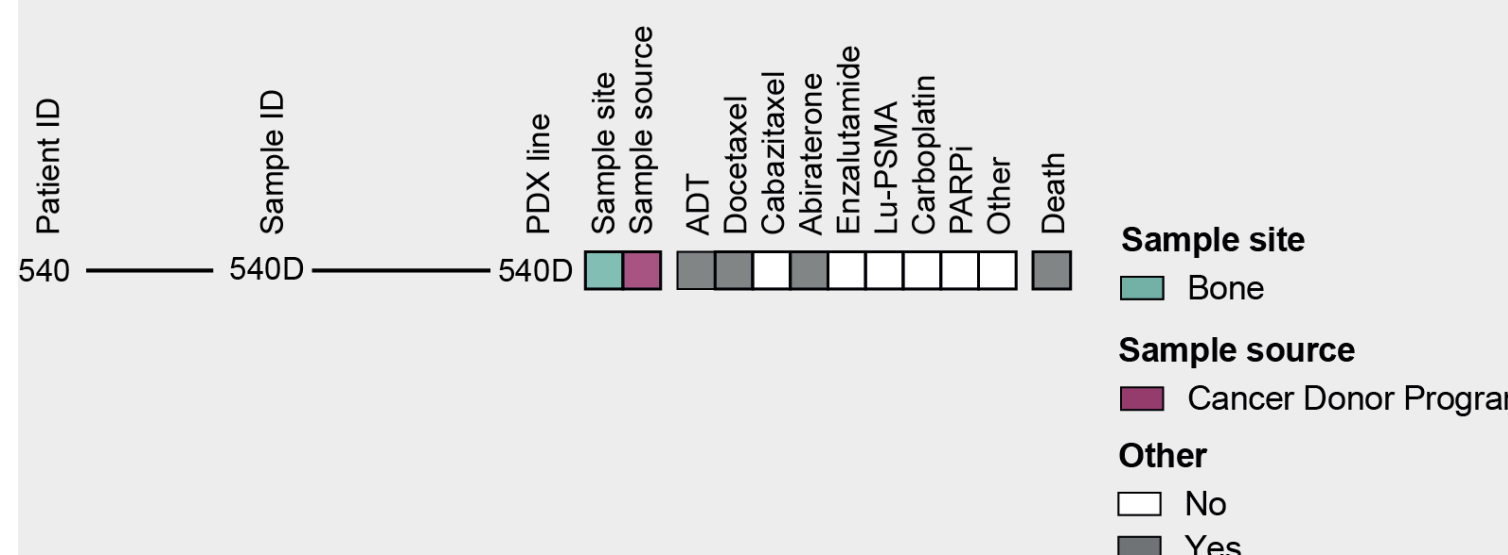
A) PSA From diagnosis to death



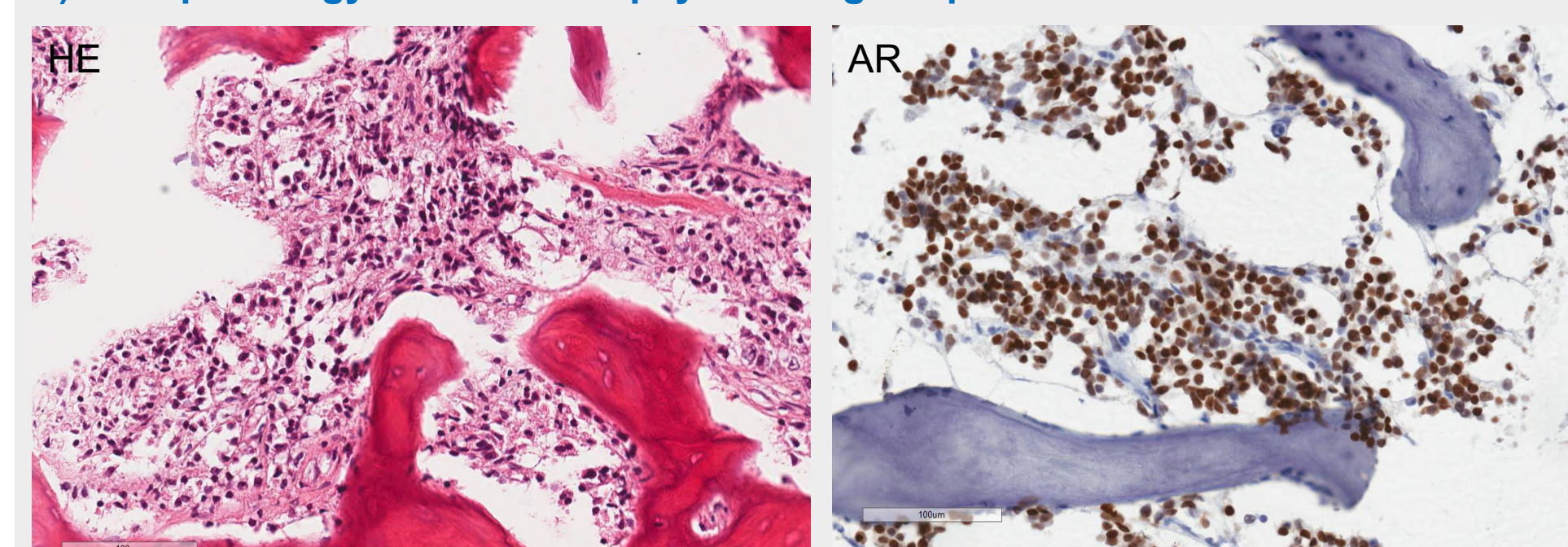
B) Whole body bone scan with biopsied site shown



C) Summary of treatment history



D) Histopathology of tumour biopsy showing the presence of AR+ tumour cells

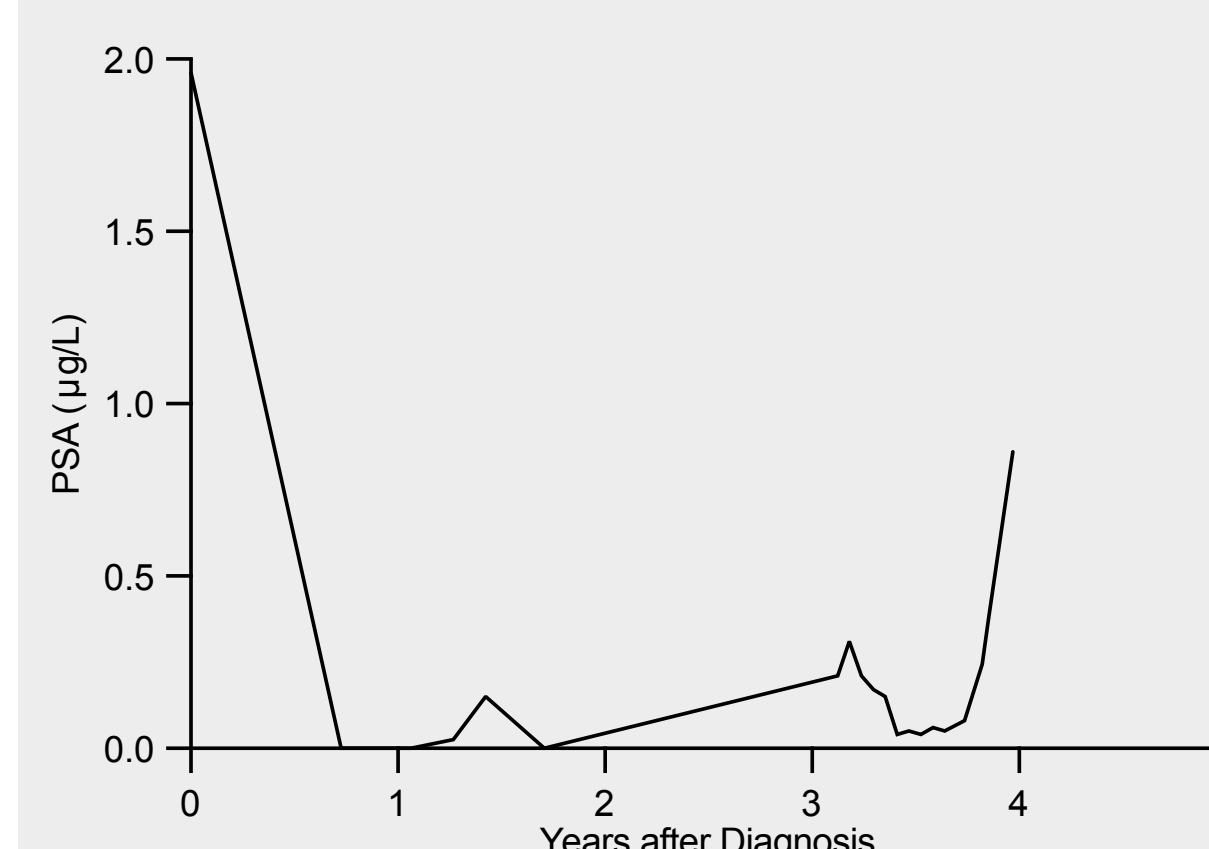


E) Next of kin response from questionnaire

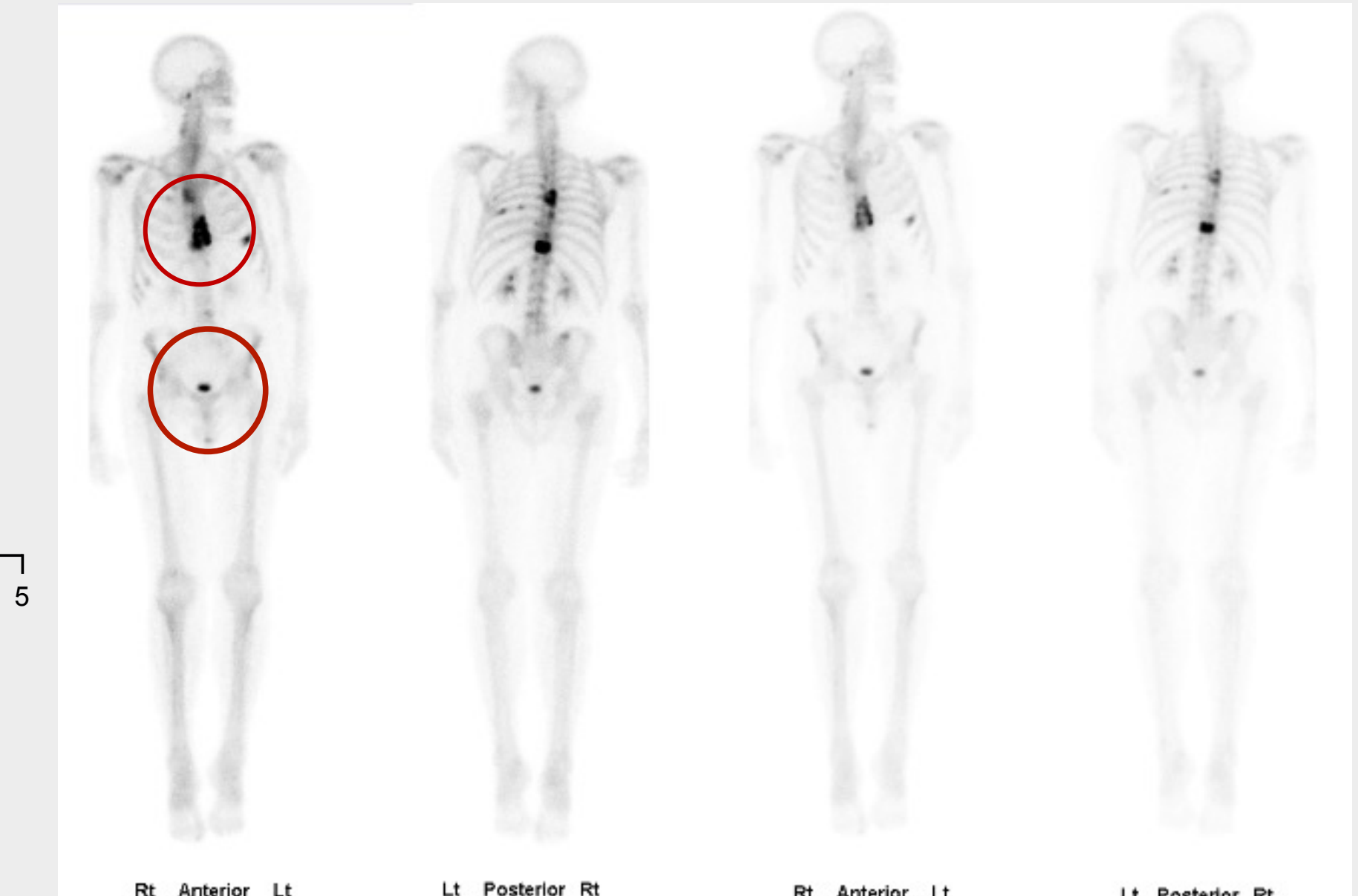
- Q1. How do you feel about your family's involvement in the cancer donor program?
R1. *Smooth process*
- Q2. Do you have any suggestions to improve the process of cancer donation?
R2. *No, easy as already in hospital.*
- Q3. Do you have any concerns that you feel the research team should be aware of?
R3. *No*
- Q4. Do you have any other feedback?
R4. *Keen to be updated on future outcomes*

Patient 2 (543D)

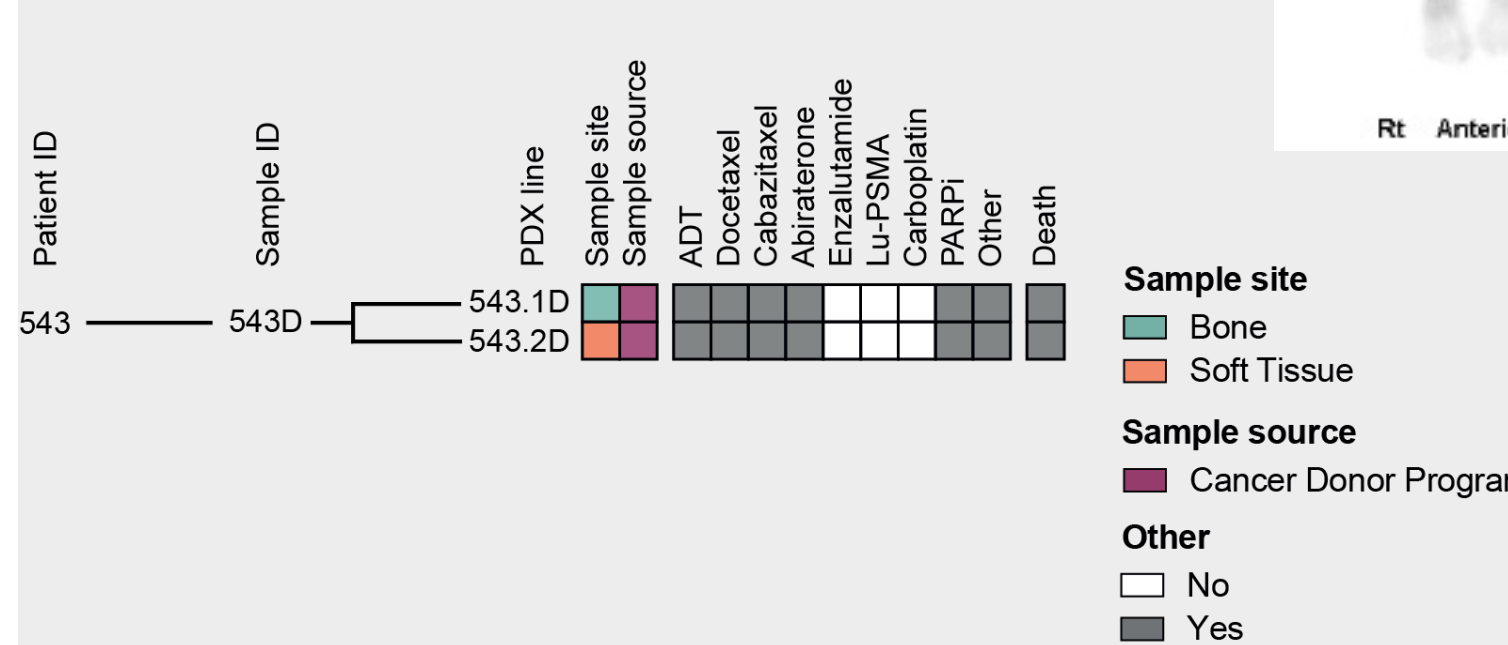
A) PSA From diagnosis to death



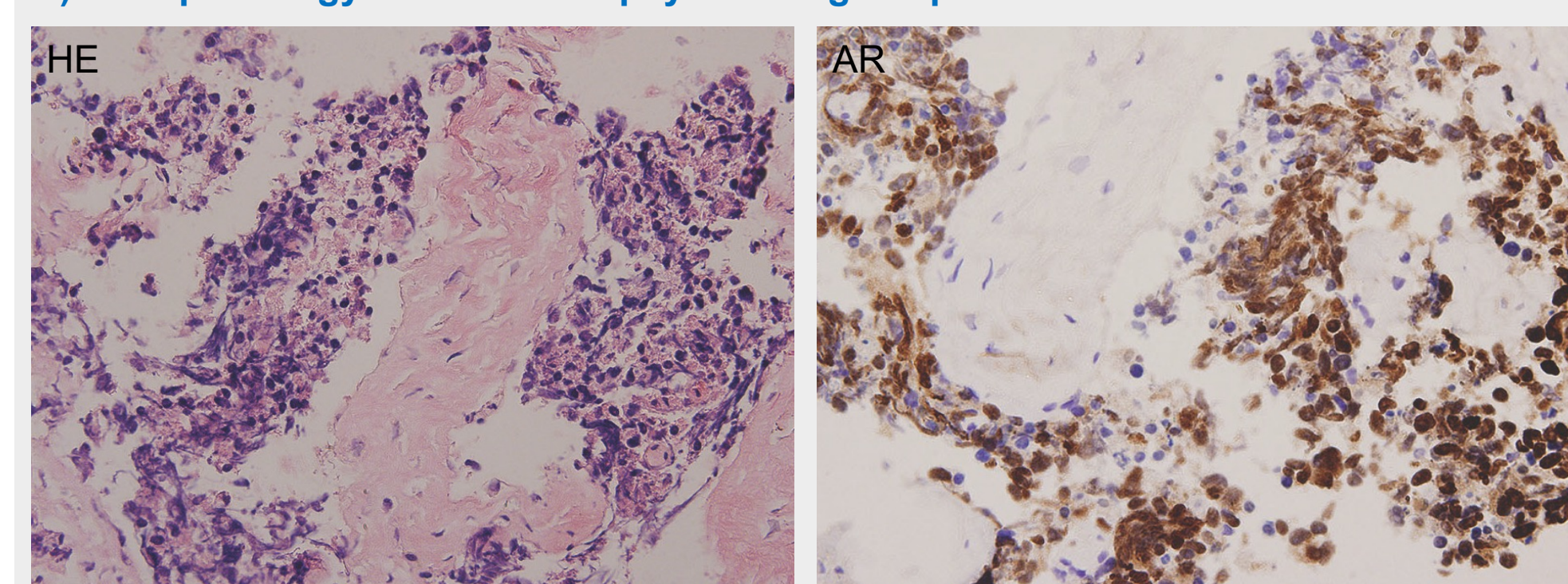
B) Whole body bone scan with biopsied site shown



C) Summary of treatment history



D) Histopathology of tumour biopsy showing the presence of AR+ tumour cells



E) Next of kin response from questionnaire

- Q1. How do you feel about your family's involvement in the cancer donor program?
R1. *Ok, it is what the patient wanted which was important to the family.*
- Q2. Do you have any suggestions to improve the process of cancer donation?
R2. *No, was an easy process.*
- Q3. Do you have any concerns that you feel the research team should be aware of?
R3. *No*
- Q4. Do you have any other feedback?
R4. *No, happy to have followed through with patient's wishes.*

CONCLUSION & FUTURE DIRECTIONS

- Prostate cancer donation performed post-mortem at the bedside appears to be a feasible approach to collect valuable cancer samples.
- Viable cancer was present in both patient biopsies and has been grafted into mice.
- Patients from some ethnic and religious backgrounds are unable to donate cancer samples if this delays funeral arrangements.
- This has been reported as a positive experience for patient families.

Future directions

- Assess PDX take rate on collected tissue
- Continue to monitor family and staff experience of donation process
- Consider expanding protocol to different tumour types

REFERENCES

