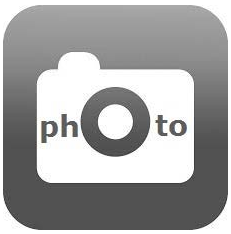
Local hospital headed paper to be used



PHOTOdynamic versus white light-guided treatment of non-muscle invasive bladder cancer: randomised trial of clinical and cost-effectiveness

Patient Information Sheet

We are inviting you to take part in a research study called PHOTO.

Before you decide whether to take part it is important that you understand why the research is being done and what it will involve. One of your doctors or nurses will go through this information sheet with you and answer any questions you may have. Please take time to read the information carefully and to discuss it with relatives, friends and your GP if you wish.

Please ask if anything is unclear or you need any further information.

Thank you for reading this and considering taking part in our research.

What is the purpose of the study?

For some people, early (non-muscle invasive) bladder cancer comes back after it is first treated. Initial treatment for early bladder cancer involves surgery to remove the cancer (transurethral resection of a bladder tumour (TURBT)). This surgery is done using a telescope put into the bladder through the urethra (cystoscopy).

Surgery is normally done using a white light to allow your surgeon to see the cancer and remove it.

There is an option to use blue light during surgery instead. If this is used after a liquid has been applied to the inside of the bladder, it may help your surgeon see more of the cancer and to remove it more effectively. It is not known if this blue light approach is better than the standard white light surgery and this is the focus of the study.

The purpose of PHOTO is to find out whether using blue light during surgery means that the cancer is less likely to come back within 3 years after treatment.

Why have I been invited?

We have invited you to be a part of this study because you have recently been diagnosed with an early bladder tumour that will require surgery. Your doctor feels that surgery using either white or blue light would be a suitable option for you. Hospitals all over the UK are taking part and 533 people like you will be included in this study.

Do I have to take part?

No, it is up to you to decide whether to take part. If you do decide to take part, you will be given this information sheet to keep and be asked to sign a consent form. You will be free to withdraw from the study at any time and do not have to give a reason. This will not affect the standard of care you receive. If you choose not to participate, you will have surgery according to the usual practice within the department.

What will happen to me if I take part?

All participants in this study will have surgery to remove their bladder cancer. Everyone who agrees to take part in this research study will be allocated at random to one of two groups. Half of the people taking part will receive white light surgery and half will have blue light surgery.

The only way to make sure that the two groups are as similar as possible is to have the treatment decided upon by chance: a process called randomisation. This process ensures that the treatments are compared fully and fairly.

If you agree to take part, your doctor or nurse will contact the research centre. The centre will record your details and tell your specialist your treatment, which will be selected by chance - you will have an equal chance of having either of the treatments. Please ask your doctor which group you are in on the day of surgery, if you would like to know. Whichever group you are in, you will be treated with the best possible care and will be monitored closely.

The two groups are:

White light surgery: You will have surgery to remove the cancer, called a transurethral resection of bladder tumour (TURBT). This surgery will be done using a standard white light cystoscopy. It is likely that you will have a temporary catheter placed after this surgery.

Blue light surgery: You will have a TURBT using a blue light. A liquid called HAL (hexaminolevulinate) will be placed in your bladder using a catheter 1 hour before your surgery. You will be asked not to pass urine for 1 hour, to give the HAL time to work before surgery. HAL is absorbed by cancer cells and glows red when seen under blue light. It is likely that you will have a temporary catheter placed after this surgery.

What are the side effects of the treatments?

You may experience side effects associated with the surgery to remove the tumour. Your doctor will discuss specific risks of the operation with you.

You may also experience side effects associated with the anaesthetic or HAL liquid given before surgery. Around 1 in 50 people may experience the following:

- Headache
- Nausea, vomiting, constipation, diarrhoea

You may see a small amount of blood in your urine for up to 2 weeks after treatment, whichever group you are in. You should avoid any heavy lifting, sports, sexual activity or driving a car during this time.

There is no reliable way to predict whether you will have any of these side effects, or how serious they might be. Your doctor will give you treatment for any problems you may experience.

What do I do if I have side effects?

It is important that you tell your hospital doctor or nurse about any problems you have at each visit so that they can give you treatment if needed. Their telephone numbers are at the end of this information sheet. There is also 24 hour support available from your hospital for patients who experience any side effects, to provide access to immediate medical care in the event of any serious problems.

Pregnancy, contraception and the PHOTO study
Two forms of contraception, including one barrier method (eg intra-uterine device (IUD) and condom, diaphragm with spermicide and condom) should be used if you or your partner are likely to become pregnant, while receiving treatment in the PHOTO study. If you have any questions about what type of contraception to use, please speak to your hospital doctor.

Women: You should inform your doctor immediately if you become pregnant within 7 days after treatment. You should not breastfeed for 7 days after treatment.
Men: If your partner becomes pregnant within 7 days after your treatment, you should inform your doctor immediately.

What will happen after surgery?

In a few cases, the tissue results from your first surgery may show you should have a second TURBT to double check that all of the cancer has been removed. This second TURBT will be done using the same type of light as your first operation.

After surgery, whichever group you are in, you will attend hospital for regular check-ups as part of normal follow up procedures. These will include a cystoscopy to check whether the cancer has returned in your bladder, with treatment as needed. The check-ups will be at the same times as those you would have if you weren't in the study and therefore the study doesn't require any additional hospital visits. As part of this study, we will perform assessments at the same time of these check-ups at around the following times:

- 3 months after surgery
- 6 months
- 9 months
- 1 year
- 18 months
- 2 years
- 3 years

You may also have extra visits not listed above as part of your hospital's normal appointment schedule.

When you join the study and before you leave hospital after your surgery you will be asked to fill in some questionnaires. Everyone in the study will also be sent questionnaires by post at approximately 3, 6, 12, 18, 24, 30 and 36 months after you join the study, up to two reminders may also be sent if we don't hear back from you. The questionnaires ask about how your treatment affects your quality of life and general wellbeing and will also collect details of how often you have visited health care professionals and the impact on any work you may do. This information will be used to investigate whether treatment with blue light would be cost-effective for the NHS. Completing the questionnaires should take about 20 minutes but you can take as long as you need. A self-addressed envelope will be provided for your convenience.

What are the possible benefits of taking part?

At the moment we don't yet know if there are any benefits from the blue light surgery, that is what we aim to find out with this study. The information we get may help us improve treatments for people with bladder cancer in the future.

What are the possible disadvantages and risks of taking part?

If you are in the blue light group, you may experience side effects from the HAL described above or discomfort from the catheter used before the operation.
If you have private medical insurance you may wish to consult your medical insurers before agreeing to take part in the study. This is to ensure that your participation will not affect your medical insurance cover.

What if something goes wrong?
Every care will be taken in the course of this study. If you are not happy with the general care and treatment you receive during the study, please speak first to your study doctor, who will try to resolve the problem. Any complaint about the way you have been dealt with during this study or any possible harm you might suffer will be addressed. You will be closely monitored both during and after therapy and any side effects will be treated as appropriate. If you are harmed by taking part in this research project, there are no special compensation arrangements. If you are harmed due to someone's negligence, then you may have grounds for legal action but you may have to pay for it.

If you wish to complain about any aspect of the way you have been approached or treated during the course of this study, the normal National Health Service complaints mechanisms are available to you. We recommend that you obtain a copy of your hospital's complaints procedure or policy if you intend to make a complaint. Concerns should be raised by speaking to a member of staff at your hospital or by talking to your local Patient Advice and Liaison Service (PALS) (or equivalent outside England).

Will my taking part in the study be kept confidential?
Your medical notes will be seen by authorised members of the research team at your hospital, so that they can collect information needed for the PHOTO trial. After you join the study, your name, date of birth, address and postcode, hospital number and NHS or Community Health Index (CHI) number will be passed to the Centre for Healthcare Randomised Trials (CHaRT) and the Clinical Trials and Statistics Unit at the Institute of Cancer Research (ICR-CTSU) who are coordinating the study. You will be given a unique registration number, which will be used to enter your data on a secure website. All information about you will be coded with the registration number and will be stored securely. It will be treated as strictly confidential and nothing that might identify you will be revealed to any third party.

Scientific and medical employees of ICR-CTSU and CHaRT, and those conducting the study with them or members of regulatory bodies, may need to examine your medical records to ensure the study is being run properly and that the information collected on the forms is correct, but your confidentiality will be protected at all times.

We will contact your hospital over the years to find out how you are getting on. We would also like to use NHS and national health and registration data to follow up your

health status. We will need enough information to identify you. This is usually your name, date of birth and NHS number (or Community Health Index and/or hospital number in Scotland). Any details we receive from these sources are confidential and will only be used for the purposes of the PHOTO study. Please initial the consent form to show that we have your permission to do this.

All the information that is sent to CHaRT and ICR-CTSU will be kept until 5 years after the PHOTO trial has ended.

Data sharing

We would like to be able to combine information we collect about patients in this study with information collected for other studies, if in the future it is a useful way of advancing our knowledge of the treatment of cancer. If this happens, information about you may be passed to other researchers, but they would not be able to identify you from the information provided.

What if relevant new information becomes available?

Sometimes we get new information about the treatment being studied. If this happens, your doctor will tell you and discuss whether you should continue in the study. If you decide not to carry on, your doctor will make arrangements for your care to continue.

What will happen if I don't want to carry on with the study?

You are free to withdraw from the study or any of its components at any time. You do not have to give a reason and your future treatment and care will not be affected. If you change your mind about having the treatment in this study, we would still like to collect information about how you are getting on. The information we need is routinely recorded at your standard hospital visits and you would not need to do anything.

What will happen to the results of the research study?

Independent experts will review the progress of the research, and the results will be published in a respected journal as soon as there is enough information to be sure the results are reliable. You will not be identified in any report or publication. The results will help to decide how to treat bladder cancer in the future. Studies like these are often used in cancer research.

Your hospital will write to you when the results are known to ask if you would like to see them. The letter will explain how to get a copy. The results of this study are not likely to be available for at least 8 years.

Who is organising and funding the research?

PHOTO is organised by leading doctors across the UK. The Chief Investigator is Mr Rakesh Heer, a consultant urologist based at Newcastle University. The study is

being run by the Clinical Trials and Statistics Unit at the Institute of Cancer Research (ICR-CTSU) in collaboration with the Centre for Healthcare Randomised Trials (CHaRT) at the University of Aberdeen. PHOTO is funded by the National Institute for Health Research Health Technology Assessment (NIHR HTA) Programme.

Who has reviewed the study?
All research in the NHS is looked at by independent group of people, called a Research Ethics Committee, to protect participants' safety, rights, wellbeing and dignity. This study has been reviewed and given favourable opinion by the Newcastle & North Tyneside 2 Research Ethics Committee on behalf of all hospitals throughout the UK.

What happens now?
You will have some time to think about the study and make your decision. Your doctor or nurse will be happy to answer any questions. You may wish to discuss it with your family or friends. Once you have reached your decision please let your doctor or nurse know. You will be asked to sign a consent form and will be given a copy to keep together with this information sheet. Please keep this information sheet and copies of the signed consent form. Your GP will be told that you are taking part in the PHOTO study. If at any time you have any questions about the study you should contact your hospital doctor, whose contact details are at the end of this sheet.

If you might be interested in participating in PHOTO, we would also like to ask you to donate blood, urine and tissue samples – you can find out more information about this optional study below.

Collecting blood, urine and tissue samples

PHOTO gives us the opportunity to ask people with early bladder cancer whether we can collect samples that will aid future research into the disease.

This part of the PHOTO study is optional and you do not have to participate if you do not wish to. When you join the study and sign your study consent form, there will be a section to complete if you agree to the collection of these samples.

Your hospital may or may not be collecting all of these samples. For more information about whether you can donate samples at your hospital please talk to your doctor or nurse.

What samples are being collected?

Tissue Samples
We would like to collect tissue samples to carry out laboratory research in the future. This may enable us to better understand which treatments are best for which patients.

Bladder cancer tissue is routinely removed when you have surgery. When extra tissue is left over after your doctor has performed the necessary tests this is routinely stored in paraffin wax in the hospital. We would like to collect some of that left over tissue.

We would like to collect these tissue samples from your initial surgery, whichever group you are in and again if your cancer comes back.

Blood and Urine Samples
We would also like to collect and store blood and urine for research.

We would like to collect a blood sample just before you have surgery and then 3, 12, 24 and 36 months when you visit the hospital for check-ups after surgery. We would need to take about 15 mls of blood (about 3 teaspoons full).

We would like to collect a urine sample when you enter the study and then 3, 12, 24 and 36 months after surgery. We would also like to collect blood and urine samples if your cancer comes back in the future.

If you agree to participate your doctor or nurse will provide you with a urine sample collection kit to take home. The kit contains collection tubes, prepaid postage and instructions for collecting and posting the samples to the laboratories at University College London and Newcastle University. Around 200ml of urine will be collected each time. In the unlikely event a sample is misplaced in the post, we will kindly ask you to provide a further sample. There will only be one further request per sample collection time point.Please initial the consent form if you agree to the collection of these samples. Your participation in this part of the study is voluntary and if you decide against it, you can still take part in the main part of the PHOTO study.

What will happen to my samples?
The samples will be stored and used to test for genetic differences in the make-up of individuals that may indicate why they develop bladder cancer and how the cancer reacts to treatment. The samples will be used to extract DNA (the genetic material inside a cell) which will also be stored. If we show that genetic differences do explain why some patients develop bladder cancer or how the cancer reacts to treatment, this knowledge could help many patients in the future.

If you agree to the collection of these samples, they will be labelled with a code and any personal details will be removed. The coding will maintain your confidentiality whilst allowing biological details to be compared to treatment findings.

The group of medical professionals overseeing the main PHOTO study will also supervise the sample collection study. All samples collected in the PHOTO study will be sent to University College London and Newcastle Biomedicine Biobank for

storage. The samples collected in this study will be stored indefinitely. Samples collected will be sent to receiving laboratories by Royal Mail post.

Samples may be used in association with commercial partners, such as pharmaceutical companies, for research purposes. Any resulting benefits from these partnerships will be used to improve patient care or to enable researchers to perform further research to bring benefits to patient treatment.

Although the research will not be conducted for the purposes of profit it is possible that some of the results, once published, will be of value to commercial companies, for example in the development of new tests or treatments. If samples are used in association with commercial partners there will be no financial benefits for participants. Samples may also be transferred to other research laboratories in the UK or overseas in the future.

Your confidentiality will be fully protected at all times and your personal details will not be disclosed to laboratory researchers. Research using the samples will be conducted only after approval by a Research Ethics Committee.

Will I be told the results of research tests on my samples?
No. Nor will the result be given to your hospital doctor or general practitioner. The overall results of the research projects will be published in the scientific literature. In the future, if the research showed that there was a test which might be useful to you then you would be able to discuss the test with your doctor.

What do I have to do?
Please initial the consent form if you agree to the donation of these samples. Donation is voluntary and if you decide against it, it will not affect your treatment in any way and you can still take part in the main PHOTO study.

What if I have other concerns?
If you have any questions you would like to ask about this study please contact your local PHOTO specialist.

Your local PHOTO specialist (consultant) is: xxx
Local research nurse/clinical nurse specialist:

Address:
Telephone:
Email:
24 hour contact number:

Thank you for thinking about taking part in our research.

NHS
National Institute for Health Research

The PHOTO Trial is funded by the National Institute for Health Research's Health Technology Assessment Programme (Project: 11/142/02).

Trust headed paper to be used

The PHOTO Trial

PHOTOdynamic versus white light-guided treatment of non-muscle invasive bladder cancer: randomised trial of clinical and cost-effectiveness

Consent Form

MREC Study No: 14/NE/1062 Patient Trial ID:

Name of Clinician: ……

Please initial box

1. I confirm that I have read and understand the patient information sheet Version 2 dated 01/07/2015 for the above study and have had the opportunity to ask questions. ☐
2. I understand that my participation is voluntary and that I am free to withdraw at any time, without giving any reason, without my medical care or legal rights being affected. ☐
3. I understand that my personal contact details will be kept confidentially and securely by the study office in Aberdeen. I agree that the study co-ordinators can use my contact details to send me study questionnaires and to contact me by phone or post or email. ☐
4. I understand that sections of any of my medical notes may be looked at by responsible individuals from the research team or from regulatory authorities where it is relevant to my taking part in research. I give permission for these individuals to have access to my records. ☐
5. I agree to my GP being informed about my participation in this study. ☐
6. If I withdraw from the study, I consent to my doctor providing authorised researchers with basic clinical information that would be routinely collected and written in my medical records *(optional)*. ☐
7. I agree to my name, date of birth, postcode, hospital number and NHS or Community Health Index (CHI) number being sent to the researchers when I join PHOTO. ☐
8. I agree to the researchers using NHS and national health and registration data to follow up my health. ☐
9. Data sharing: I grant advance authorisation for the possible future sharing of information collected about me with other organisations, with the understanding that I will not be identifiable from this information. ☐
10. I agree to participate in the PHOTO study. ☐

Tissue, blood and urine samples (optional)

11. Please initial this box if you are willing to donate the samples as described in the PHOTO patient information sheet V2 01/07/2015. ☐

12. I grant advance authorisation for future research on my stored samples, with the understanding that I will not be identifiable from these samples and that prior approval of an ethics committee will be obtained. ☐

Name of Patient	Date	Signature
Name of person taking consent (if different from researcher)	Date	Signature
Researcher (PI)	Date	Signature

Original copy to be kept in the PHOTO trial site file;

1 copy to be kept in hospital notes, 1 copy to the patient, 1 copy for the Aberdeen PHOTO trial office