* midnight the day prior to the 18th birthday

This form is consistent with and should be used in conjunction with, the AoMRC (2008) A Code of Practice for the Diagnosis and Confirmation of Death¹ and has been endorsed for use by the following institutions: Faculty of Intensive Care Medicine, Intensive Care Society, Paediatric Intensive Care Society, Royal College of Paediatrics and Child Health and National Organ Donation Committee: Paediatric Subgroup. Date for

Number HOSPITAL ADDRESSOGRAPH or

Surname First Name Date of Birth NHS/CHI number

review: 1/5/2023

Objective of Care

• To diagnose and confirm the death of a mechanically ventilated, severely brain injured patient in coma, using neurological criteria.

Academy of the Medical Royal Colleges Definition of Human Death (2008)1

"Death entails the irreversible loss of those essential characteristics which are necessary to the existence of a living human person and, thus, the definition of death should be regarded as the irreversible loss of the capacity for consciousness, combined with irreversible loss of the capacity to breathe. The irreversible cessation of brain-stem function whether induced by intra-cranial events or the result of extra-cranial phenomena, such as hypoxia, will produce this clinical state and therefore irreversible cessation of the integrative function of the brain-stem equates with the death of the individual and allows the medical practitioner to diagnose death."

Context

- National professional guidance advocates the confirmation of death by neurological criteria wherever this seems a likely diagnosis and regardless of the likelihood of organ donation.^{3,4,7}
- UK General Medical Council (GMC) guidance on end of life care (2010) states that national procedures for identifying potential organ donors should be followed and, in appropriate cases, the specialist nurse for organ donation (SN-OD) should be notified. NICE guidance and UKPICS Standards recommend that the specialist nurse for organ donation (SN-OD) should be notified at the point when the clinical team declare the intention to perform brain-stem death tests.^{2,3,4}

Date and time of referral to SN-OD:

- Whilst most patients will already be in an Intensive Care Unit (ICU) when the diagnosis is suspected, some patients may be in other areas, e.g. the Emergency Department. On such occasions, it is legitimate, if considered necessary, to transfer a patient to the ICU for the diagnosis to be made.
- For many clinicians, the diagnosis and confirmation of death using neurological criteria, will be a relatively infrequent task and may be complicated by uncertainties regarding the nature of the primary diagnosis, irreversibility and the availability of suitably experienced personnel. Updated guidance on the diagnosis and confirmation of death by neurological criteria was published by the Academy of the Medical Royal Colleges (AoMRC) in 2008. ¹ For infants from 37 weeks to < 2 months guidance was published by the Royal College of Paediatric and Child Health in 2015. 6</p>

The person with parental responsibility for the child and other close family members should be made aware that the purpose of testing is to confirm the child's death. If given an opportunity to witness the neurological examination, they should be appropriately supported and prepared for the possibility of spinal reflexes and their lack of relevance, as far as the diagnosis of death by neurological criteria is concerned. Whether the child's close family witness the clinical examination or not, the child's need for dignity and privacy, should remain paramount.

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Preparation

1. Evidence for Irreversible Brain Damage of known Aetiology

Case records, past medical history including possibly contacting the GP, relevant imaging.

2. Exclusion of Reversible Causes of Coma and Apnoea

Standard ICU cardio-respiratory monitoring (to ensure haemodynamic stability), medication chart and history, blood and urine drug assay results (where relevant), drug antagonists (e.g. flumazenil, naloxone), peripheral nerve stimulator, recent serum glucose and biochemistry, thermometer, patient warming device.

3. Tests for Absence of Brain-Stem Function

Brain-stem reflexes: Bright light source; small sterile gauze swabs, otoscope with disposable ear pieces, 20 or 50 ml luer lock syringe and disposable quill, ice-cold water; a spatula, Yankauer sucker or laryngoscope, endotracheal suction catheters.

Apnoea test: Haemodynamic monitoring (continuous ECG, invasive arterial pressure), arterial blood gas analysis including blood gas syringes x 4, pulse oximetry and end-tidal CO_2 monitoring, means of delivering oxygen to the trachea by bulk flow (i.e. using a circuit which maintains oxygenation and CPAP e.g. Mapleson C , Ayres T piece, or endotracheal suction catheter and oxygen tubing).

Diagnostic caution is advised in the following 'Red Flag' patient groups. (Based on the literature and unpublished case reports.) For advice in difficult circumstances contact the local or regional Clinical Lead for Organ Donation or the regional paediatric intensive care units.

- 1. Testing **< 6 hours** of the loss of the last brain-stem reflex
- 2. Testing <24 hours from the loss of last brain stem reflex where aetiology primarily anoxic damage
- 4. Patients with **any neuro-** 6. Prolonged **fentanyl** infusions **muscular disorders**
- 5. **Steroids** given in space occupying lesions such as abscesses
- 7. Aetiology **primarily** located to the **brain-stem or posterior fossa**
- 3. **Hypothermia** (24-hour observation period following re-warming to normothermia recommended)

Examining Doctors				
Date and time:	Patient Location			
Doctor One, Name and Designation	Doctor Two, Name and Designation			
Name:	Name:			
Grade:	Grade:			

Guidance

- 1. The diagnosis of death by neurological criteria should be made by at least two medical practitioners. Both medical practitioners should have been registered with the General Medical Council (or equivalent Professional Body) for more than five years and be competent in the assessment of a patient who may be deceased following the irreversible cessation of brain-stem function and competent in the conduct and interpretation of the brain-stem examination. In infants, children and young adolescents it is recommended that one of the doctors should be a consultant, one of them should normally be a paediatrician or should have experience with children and one of them should not be primarily involved in the child's care. Those carrying out the tests must not have, or be perceived to have, any clinical conflict of interest and neither doctor should be a member of the transplant team. Clinical Leads for Organ Donation can carry out testing and are likely to have significant expertise.
- 2. Testing should be undertaken by the nominated doctors acting together and must always be performed on two occasions. A complete set of tests should be performed on each occasion, i.e., a total of two sets of tests will be performed. Doctor One may perform the tests while Doctor

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Two observes; this would constitute the first set. Roles may be reversed for the second set. The tests, in particular the apnoea test, are therefore performed only twice in total.

Evidence for Irreversible Brain Damage of known Aetiology

Diagnosis:

Evidence for Irreversible Brain Damage of known Aetiology:

Guidance

- 1. The patient must have a Glasgow Coma Scale score of 3 and be mechanically ventilated with apnoea.
- 2. There should be no doubt that the patient's condition is due to irreversible brain damage of known aetiology.
- 3. It remains the duty of the two doctors carrying out the testing to be satisfied with the aetiology, the exclusion of all potentially reversible causes, the clinical tests of brain-stem function and of any ancillary investigations (see page 8); so that each doctor may independently confirm death following irreversible cessation of brain-stem function.
- 4. Occasionally it may take a period of continued clinical observation and investigation to be confident of the irreversible nature of the brain injury. The timing of the first test and the timing between the two tests should be adequate for the reassurance of all those directly concerned.
- 5. It is recommended that there is a minimum of twenty-four hours, of continued clinical observation, in patients where anoxic damage, following cardiorespiratory arrest, is the aetiology of the brain injury. If prior treatment of the patient has included induced hypothermia, it is recommended that there is a minimum of twenty-four hours, of continued clinical observation, following re-warming to normothermia. See above for 'Red Flag' patient groups.
- 6. Stabilisation of the patient prior to testing, especially support of the cardiovascular system, is a prerequisite to testing. Mean blood pressure should be consistently maintained at age appropriate levels (see appendix 1)⁵ and appropriate fluid resuscitation administered. This almost invariably requires the use of inotropes / vasopressors via central venous access.
- 7. Diabetes insipidus can develop rapidly and should be suspected in patients with a high urine output (typically > 4 ml/kg/hr) and rising serum sodium concentration. Matched urinary and plasma electrolytes and osmolality may assist in the diagnosis. Treatment with desmopressin; 2 month 12 years 400 nanograms, 12 18 years 1 4 micrograms IV boluses, is usually sufficient for treatment but repeated doses or vasopressin infusion may be required. Plasma sodium should ideally be maintained within normal limits.

Exclusion of Reversible Causes of Coma and Apnoea

Guidance

Attempts should be made to maintain relatively normal cardiovascular and respiratory physiological parameters in the preceding hours prior to testing. *This may not be possible and does not necessarily preclude testing.*

The key question the two doctors must exclude is the possibility that cardiovascular and respiratory instability is the cause of the observed coma and apnoea.

The answer should be no.

Suggested cardiovascular goals for children:

• Sinus rhythm 60 - 170 beats per minute, as age appropriate (see appendix 1)⁵

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Patient Name:	DOB:	NHS/CHI Number:
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	1st 7	Гest	2nd	Test
Mean arterial pressure at time of testing? Should be consistently maintained at age		mmUa		mmlla
appropriate levels prior to testing.		mmHg		mmHg
PaCO ₂ at time of testing?				
A goal of normocarbia ($PaCO_2 < 6.0 \text{ kPa}$), if possible, is recommended in the hours prior to testing. See below for starting $PaCO_2$ in the apnoea test.		kPa		kPa
PaO₂ at time of testing? Hypoxia should be avoided (PaO ₂ >10 kPa). For infants/children with congenital cyanotic heart disease oxygen levels should be kept in their normal range		kPa		kPa
Arterial pH/[H+] at time of testing? Acidaemia and alkalaemia should be avoided, <i>if</i> possible, aiming for a relatively normal pH 7.35 - 7.45 / [H+] 45 - 35 nmol/L.	pH/[H+]=		pH/[H+]=	
Is the coma or apnoea due to ongoing cardio- respiratory instability?	Dr One	Dr Two	Dr One	Dr Two
(To diagnose death using neurological criteria, ALL answers should be NO)	Yes / No	Yes / No	Yes / No	Yes / No

Guidance

- The patient should not be receiving any drugs that might be contributing to the unconsciousness, apnoea and loss of brain-stem reflexes (narcotics, hypnotics, sedatives or tranquillisers); nor should they have any residual effect from any neuromuscular blocking agents (atracurium, vecuronium or suxamethonium).
- It remains the duty of the two doctors carrying out the testing to be satisfied that sufficient time has elapsed to ensure that any remaining drug effect is non-contributory to the unconsciousness and loss of brain-stem reflexes. This will be based on an assessment of the medications the patient has received and from knowledge of the pharmacokinetics of these agents. Renal or hepatic failure and immaturity in younger children may prolong metabolism / excretion of these drugs. See above for 'Red Flag' patient groups.
- Serum Na⁺ levels below 115 or above 160mmol/l are associated with unresponsiveness. This should be borne in mind if the primary cause of unresponsiveness is uncertain. Testing for DNC should not be carried out if serum K⁺ <2 mmol/L, or serum PO_4^{3-} and/or Mg^{2+} < 0.5 mmol/L or > 3.0 mmol/L as there may be associated severe neuromuscular weakness.
- Blood glucose should be between 3.0 20mmol/L before each brain-stem test. If there is any clinical reason to expect endocrine disturbances, then it is obligatory to ensure appropriate hormonal assays are undertaken.

	1st Test	2nd Test
Where there is any doubt, specific drug levels	Drug levels	Drug levels
should be measured (Testing for DNC should not be	(if measured):	(if measured):
carried out if midazolam level is > 10mcg/L, or		
thiopentone level is > 5mg/L)		

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Patient Name:......DOB:.....NHS/CHI Number:.....

Antagonists such as flumazenil, naloxone and neostigmine may be used but there is no specific pharmacological data for predicting the dose effect of these antagonists.	Drug antagonists (if used):		Drug antagonists (if used):	
Residual neuromuscular blockade can be tested for, if necessary, by peripheral nerve stimulation.	Train of Four (if measured):		Train of Four (if measured):	
Is the coma or apnoea due to depressant drugs? (To diagnose death using neurological criteria, ALL answers should be NO)	Dr One Yes / No	Dr Two Yes / No	Dr One Yes / No	Dr Two Yes / No
Body temperature at time of testing? If core temperature is ≤34°C testing cannot be carried out.		°C		°C
Serum sodium (Na+) at time of testing? Serum sodium should not be < 115 or > 160 mmol/L. Rapid rises or falls in Na+ should be avoided.	mmol/L		mmol/L	
Serum potassium (K+) at time of testing? Serum potassium should be >2 mmol/L	mmol/L		mmol/L	
Serum phosphate (PO ₄ ³⁻) at time of testing? Serum phosphate should not be profoundly elevated (>3.0mmol/L) or lowered (<0.5mmol/L) from normal.	mmol/L			mmol/L
Serum magnesium (Mg2+) at time of testing? Serum magnesium should not be profoundly elevated (>3.0mmol/L) or lowered (<0.5mmol/L) from normal.	mmol/L			mmol/L
Blood glucose at time of testing? Blood glucose should be between 3.0 - 20.0 mmol/L and should be tested prior to each test.	mmol/L			mmol/L
If there is any clinical reason to expect endocrine disturbances hormonal assays should be undertaken.	Hormone level (if measured):		Hormone (if measur	
Is the coma or apnoea due to a metabolic or endocrine disorder? (To diagnose death using neurological criteria,	Dr One Yes	Dr Two Yes	Dr One Yes	Dr Two Yes
ALL answers should be NO)	No	No	No	No

Guidance

It remains the duty of the two doctors carrying out the testing to be satisfied that the only explanation for the respiratory failure is due to the irreversible cessation of brain-stem function. A train of four examination, using a peripheral nerve stimulator, may be required. See above for 'Red Flag' patient groups.

1st Test	2nd Test

Form for the Diagnosis of Death using Neurological Criteria in Children 2 months to 18 years (full guidance version) * midnight the day prior to the 18th birthday

Patient Name:......DOB:.....NHS/CHI Number:.....

Is the apnoea due to neuromuscular blocking	Dr One	Dr Two	Dr One	Dr Two
agents, other drugs or a non brain-stem cause				
(e.g. cervical injury, any neuromuscular	Yes	Yes	Yes	Yes
weakness)?	/	/	/	/
	No	No	No	No

Tests for Absence of Brain-Stem Function

Guidance: A complete set of tests should be performed on each occasion, i.e. a total of two sets of tests will be performed. Doctor One may perform the tests while Doctor Two observes; this would constitute the first set. Roles may be reversed for the second set. The tests, in particular the apnoea test, are therefore performed only twice in total.

test, are therefore performed only twice	1st Test		2nd	l Test
To diagnose death using	Dr One	Dr Two	Dr One	Dr Two
neurological criteria, ALL answers	Examining	Observing	Observing	Examining
should be NO				
Do the pupils react to light?				
The pupils are fixed and do not respond	Yes / No	Yes / No	Yes / No	Yes / No
to sharp changes in the intensity of				
incident light - cranial nerves II, III.				
Is there any eyelid movement when				
each cornea is touched in turn?	Yes / No	Yes / No	Yes / No	Yes / No
Corneal reflex - cranial nerves V, VII.				
The use of sterile gauze is				
recommended.				
Is there any eye movement seen				
during or following the slow	Yes / No	Yes / No	Yes / No	Yes / No
injection of at least 20 - 50mls ice				
cold water over 1 minute into each				
ear with the head flexed at 30°? Each				
ear drum should be clearly visualised				
before the test. Vestibulo-ocular reflex -				
cranial nerves III, VI, VIII.				
Is the gag reflex present?				
Use a spatula or Yankauer sucker and	Yes / No	Yes / No	Yes / No	Yes / No
laryngoscope to stimulate the posterior				
pharynx - cranial nerves IX, X.				
Is the cough reflex response present				
when a suction catheter is passed				
down the trachea to the carina?	Yes / No	Yes / No	Yes / No	Yes / No
Cranial nerves IX, X				
Is there any motor response in a				
cranial nerve or somatic distribution				
when supraorbital pressure is	Yes / No	Yes / No	Yes / No	Yes / No
applied? Cranial nerves V, VII. Reflex				
limb and trunk movements (spinal				
reflexes) can be present.				

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Patient Name:	DOB:	NHS/CHI Number:
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Tests for Absence of Brain-Stem Function

Preparation for the Apnoea Test

- Oxygenation and cardiovascular stability should be maintained through each apnoea test. Preoxygenate FiO₂ 1.0.
- Allow PaCO₂ to rise to at least 6.0 kPa by reducing the minute ventilation prior to commencing the apnoea test. End tidal carbon dioxide can be used to guide the starting of each apnoea test but should not replace the pre and post arterial PaCO₂.
- Cardiac pulsation may be sufficient to trigger supportive breaths if the patient remains connected to the mechanical ventilator and on a spontaneous breathing mode. Performing the apnoea test whilst remaining on mechanical ventilation is not recommended.

Guidance:

It is important to maintain oxygenation above 85% and cardiovascular stability. Recommended methods:

- CPAP circuit (e.g. Mapleson C or Ayres T piece), especially if oxygenation is a problem, or
- Disconnect the patient from the ventilator and administer oxygen via a catheter in the trachea at a rate of 2 6L/minute. Ensure oxygen catheter does not occlude ETT.
- Considerable atelectasis develops in the apnoeic period. At the conclusion of the apnoea test, manual recruitment manoeuvres should be carried out before resuming mechanical ventilation.

	1st 7	Γest	2n	d Test
Arterial Blood Gas PRE-apnoea test:	1st Test		2 nd Test	
Confirm PaCO ₂ is at least 6.0 kPa but	Starting PaCo	O ₂ :	Starting PaC	O ₂ :
not substantially greater.		kPa		kPa
In patients with chronic CO ₂ retention,				
or those who have received intravenous	Should be ≥6	5.0 kPa	Should be ≥6	5.0 kPa
bicarbonate, it is recommended that				
PaCO ₂ is allowed to rise to above 6.5				
kPa.				
PRE-Arterial Blood Gas pH: Confirm			II	
pH < 7.4	pH =		pH =	
	Should be	e <7.4	Should b	e <7.4
Start time:				
Time when apnoea test was		hr : min		hr : min
commenced.	(24 hou	r clock)	(24 ho	our clock)
Arterial Blood Gas POST apnoea test:	1st Test		2 nd Test	
Ensure the PaCO ₂ has increased by	Stopping Pa(CO_2 :	Stopping Pa(CO_2 :
greater than 0.5 kPa.		kPa		kPa
	Chould have	ingressed by	Chould have	ingressed by
	Should have >0.5 kPa	ilicreased by	>0.5 kPa	increased by
Stop time:	/U.J KF a		/U.J KF a	
Time when apnoea test was ceased.		hr : min		hr : min
Time when aprioca test was ceased.	(24 hou		(24 hc	our clock)
	(211100	0.0011	(21110	
	Perform lung	recruitment	Perform lui	ng recruitment
Was there any spontaneous	Dr One	Dr Two	Dr One	Dr Two
respiration during a minimum of 5	DI OILE	DITWU	DI OILE	DITWU
(five) minutes continuous	Yes	Yes	Yes	Yes
()	100	100	100	100

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Patient Name:......DOB:.....NHS/CHI Number:.....

		•		
observation following disconnection	/	/	/	/
from the ventilator?	No	No	No	No
(To diagnose death using				
neurological criteria, ALL answers				
should be NO)				

Ancillary Investigations Used to Confirm the Diagnosis

Guidance:

Ancillary investigations are **NOT** required for the diagnosis and confirmation of death using neurological criteria.

They may be useful however, where neurological examination is not possible (e.g. extensive facio-maxillary injuries, residual sedation and some cases of paediatric hypoxic brain injury), where a primary metabolic or pharmacological derangement cannot be ruled out or in cases of high cervical cord injury, or where spontaneous or reflex movements in the patient generate uncertainty over the diagnosis. In such cases a confirmatory test may reduce any element of uncertainty and possibly foreshorten any period of observation prior to formal testing of brain-stem reflexes. Any ancillary or confirmatory investigation should be considered **ADDITIONAL** to the fullest clinical testing and examination (as outlined above) carried out to the best of the two doctors' capabilities in the given circumstances.

The utility of any additional investigation is for the two testing doctors to decide and they should seek further professional opinion from other specialities and other expert centres, where appropriate. Some possible ancillary investigations are:

- Neurophysiological demonstration of loss of bioelectrical activity in the brain (EEG, evoked potentials).
- Radiological demonstration of absent cerebral blood flow or brain tissue perfusion (CT angiography, 4 vessel angiography, transcranial doppler).

The interpretation of ancillary investigations is complex and their availability usually restricted to neurological centres.

Helpful references on ancillary testing

- 1. Wijdicks (2001) "The Diagnosis of Brain Death" NEJM 344:1215-21.
- 2. Young & Lee (2004) "A critique of Ancillary Tests for Brain Death." *Neurocritical Care*; 1:499-508.
- 3. Heran, Heran & Shemie (2008) "A review of ancillary tests in evaluating brain death." *Can J Neurol Sci*; 35:409–19.

Is there a need for any ancillary	Dr One	Dr Two
investigations?		
	Yes / No	Yes / No

If yes please outline the results of these investigations:

Form for the Diagnosis of Death using Neurological Criteria in Children 2 months to 18 years {full guidance version} * midnight the day prior to the 18th birthday

Patient Name:	DOB:	NHS/CHI Numl	oer:

Completion of Diagnosis				
	Test 1		Test 2	
Are you satisfied that death has been confirmed following the irreversible cessation of brain-stem function?	Dr One Yes / No	Dr Two Yes / No	Dr One Yes / No	Dr Two Yes / No
Legal time of death is when the 1st Test indicates death due to the absence of brain-stem reflexes. Death is confirmed following the 2nd Test.	Date: Time: Dr One signature Dr Two signature		Date: Time: Dr One signature Dr Two signature	

Appendix 1 Age- Appropriate Paediatric Physiological Parameters ⁵

Age	Heart Rate	Systolic BP	Mean Arterial BP
3 months	110 -170	65 - 105	44 - 79
6 months	110 - 160	65 - 105	44 - 79
12 months	110 - 160	70 - 105	46 - 79
18 months	100 - 155	70 - 105	46 - 79
2 years	100 - 150	70 - 110	49 - 84
3 years	90 - 140	70 - 110	52 - 87
4 years	80 - 135	70 - 110	54 - 89
5 years	80 - 135	80 - 120	60 - 94
6 years	80 - 130	80 - 120	62 - 96
7 years	80 - 130	80 - 120	63 - 97
8 years	70 - 120	80 - 120	64 - 98
9 years	70 - 120	80 - 120	64 - 99
10 years	70 - 120	80 - 120	65 - 100
11 years	70 - 120	80 - 120	66 - 100
12 years	65 - 115	90 - 140	69 - 107
14 years	60 - 110	90 - 140	70 - 108
14 -18 years	60 - 110	90 - 140	73 - 111

References

1. Academy of Medical Royal Colleges (2008) "A Code of Practice for the Diagnosis and Confirmation of Death" http://www.aomrc.org.uk

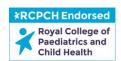
* midnight the day prior to the 18th birthday

Patient Name:......DOB:.....NHS/CHI Number:.....

- 2. GMC (2010) "Treatment and care towards the end of life." www.gmc-uk.org/guidance/ethical guidance/end of life care.asp
- 3. NICE (2011, 2016) "Organ Donation for Transplantation" http://guidance.nice.org.uk/CG135
- 4. Paediatric Intensive Care Society (2014) "PICS Organ Donation Standards" http://picsociety.uk/resources/
- 5. Advanced Paediatric Life Support A Practical Approach to Emergencies (2016)
- 6. Royal College of Paediatric and Child Health (2015) "The diagnosis of death by neurological criteria in infants less than two months old" www.rcpch.ac.uk
- 7. Report from the Organ Donation Taskforce (2008) "Organs for Transplant" http://www.dh.gov.uk/en/Publicationsandstatistics/Publications/PublicationsPolicyAndGuidance/DH 082122
- 8. A series of helpful education videos are available: https://www.odt.nhs.uk/deceased-donation-after-brainstem-death/diagnosing-death-using-neurological-criteria/

Form authorship and feedback

This form was written by Dr Dale Gardiner, Nottingham, Dr Alex Manara, Bristol and Dr Kay Hawkins, Manchester, Dr James Fraser, Bristol, Dr Margrid Shindler, Bristol and Andrea Macarthur, Manchester. Angie Scales, NHS Blood and Transplant. Comments should be directed to: reinout.mildner@nhsbt.nhs.uk





Additional NOTES