

# An analysis of heart donation after circulatory determination of death

Anne Laure Dalle Ave,<sup>1,2</sup> David Shaw,<sup>3</sup> James L Bernat<sup>4</sup>

<sup>1</sup>Ethics Unit, University Hospital of Lausanne, Switzerland

<sup>2</sup>Institute for Biomedical Ethics, University Medical Center, Lausanne, Switzerland

<sup>3</sup>Institute for Biomedical Ethics, Universität Basel, Basel, Switzerland

<sup>4</sup>Department of Neurology, Dartmouth-Hitchcock Medical Center, Lebanon, New Hampshire, USA

## Correspondence to

Anne Laure Dalle Ave, Ethics Unit, 21 rue du Bugnon, Lausanne 1010, Switzerland; Anne.Dalle-Ave@chuv.ch

Received 4 November 2015

Revised 14 December 2015

Accepted 4 January 2016

## ABSTRACT

**Background** Heart donation after circulatory determination of death (DCDD) has provoked ethical debate focused primarily on whether heart DCDD donors are dead when death is declared and when organs are procured.

**Objective and design** We rigorously analyse whether four heart DCDD programmes (Cape Town, Denver, Australia, Cambridge) respect the dead donor rule (DDR), according to six criteria of death: irreversible cessation of all bodily cells function (or organs), irreversible cessation of heart function, irreversible cessation of circulation, permanent cessation of circulation, irreversible cessation of brain function and permanent cessation of brain function.

**Conclusions** Only death criteria based on permanency are compatible with the DDR under two conditions: (1) a minimum stand-off period of 5 min to ensure that autoresuscitation is impossible and that all brain functions have been lost and (2) no medical intervention is undertaken that might resume bodily or brain circulation. By our analysis, only the Australia heart DCDD programme using a stand-off period of 5 min respects the DDR when the criteria of death are based on permanency.

## INTRODUCTION

Organ donation after circulatory determination of death (DCDD), also termed donation after circulatory death, donation after cardiac death and non-heart beating donation, has become increasingly common in the USA and in Europe. DCDD programmes most frequently provide kidneys for transplantation and, less frequently, lungs and livers. To date, only four centres have been reported doing heart DCDD: Cape Town in 1967,<sup>1</sup> Denver in 2008,<sup>2</sup> Sydney in 2015<sup>3</sup> and Cambridge in 2015.<sup>4</sup>

The 2008 report by Boucek *et al*<sup>2</sup> of heart DCDD in three newborns provoked a firestorm of ethical controversy,<sup>5 6</sup> principally around the question of how donors who were declared dead by cardiocirculatory criteria could actually be dead if their hearts regained normal function in recipients. Because of this unresolved ethical issue and the technical problem of heart susceptibility to warm ischaemia time (WIT), heart DCDD was not reported again until recently.<sup>3 4</sup> Although technical difficulties apparently have been resolved by centres in the UK and Australia, the ethical debate over heart DCDD continues. In this article, we provide a rigorous analysis of the acceptability of heart DCDD to address the question of whether heart DCDD donors are dead at the moment that death is declared and when organs are procured.

## MEDICAL CONSIDERATIONS—DESCRIPTION OF HEART DCDD PROTOCOLS

In [table 1](#), we compare the heart DCDD protocols of Cape Town,<sup>1</sup> Denver,<sup>2</sup> Cambridge<sup>4</sup> and Sydney.<sup>3</sup> Because there have been no publications of the cases of heart DCDD in Cambridge, we based our analysis on a 2009 article by Ali *et al*,<sup>7</sup> in which they described the use of a heart DCDD protocol until heart explantation, but without heart transplantation, which appears similar to the description found in the press.<sup>4</sup>

Because the heart is particularly susceptible to WIT, heart DCDD protocols aim to minimise it. Studying published heart DCDD protocols,<sup>1-3 7</sup> we identified three principal means to decrease the ischaemic insult to the heart:

1. The use of organ preservation techniques after the declaration of death:
  - A. Cardiopulmonary bypass,<sup>1</sup> also termed in situ normothermic regional perfusion,<sup>7</sup> extracorporeal membrane oxygenation (ECMO) or extracorporeal support.<sup>9</sup>
  - B. Cold fluid infusion through a femoral cannula.<sup>2</sup>
2. A reduced stand-off period, defined as the time between circulatory arrest and the declaration of death, to 75 s,<sup>2</sup> 2 min<sup>3</sup> or 3 min.<sup>2</sup>
3. The use of ex vivo perfusion.<sup>3 4</sup>

## The use of ECMO in DCDD

Although ECMO has been used in some DCDD centres to decrease WIT,<sup>7 9 10</sup> it is not the norm. Because cardiac and brain circulation resumes with the use of ECMO, DCDD protocols employing this technique use either an inflated thoracic aortic balloon<sup>9</sup> or a clamp on the ascending aorta<sup>10</sup> to exclude simultaneous cardiac and brain perfusion, and to avoid brain resuscitation. The two heart DCDD protocols reporting the use of ECMO were those of Cape Town<sup>1</sup> and of Cambridge.<sup>4 7</sup> Brain and cardiac perfusion were not prevented in the first heart DCDD reported by Barnard.<sup>1</sup> In the Cambridge heart DCDD protocol, a cross-clamp was positioned on the aortic arch vessels to permit bodily and cardiac recirculation, while still excluding brain circulation.<sup>4 7</sup> This is an unusual protocol because it 'revives' the heart inside the donor's body by using ECMO. As the heart recovered normal function, ECMO was weaned. The team waited 50 min before the heart was procured, in order to confirm good heart function for transplantation.

## Cold fluid infusion

Cold fluid infusion is widely used in DCDD protocols.<sup>11</sup> Because blood is replaced by cold fluid

**To cite:** Dalle Ave AL, Shaw D, Bernat JL. *J Med Ethics* Published Online First: [please include Day Month Year] doi:10.1136/medethics-2015-103224

**Table 1** Comparison between the four heart DCDD protocols

	Barnard, 1967, Cap Town <sup>1</sup>	Boucek <i>et al</i> , <sup>2</sup> Denver	Press 2015, Cambridge <sup>4</sup> Ali <i>et al</i> , 2009 <sup>7</sup>	Dhital, 2015, Australia <sup>3</sup>
Organ donors	1 p	3 newborns, severe neurological injury or birth asphyxia	1 p <sup>4</sup> 1 p, 57 y.o., intracranial bleeding <sup>7</sup>	3 p, ≤40 y.o., 2 trauma, 1 hypoxia
Premortem intervention	ECMO cannula	In situ cooling cannula	No	No
Organ preservation after death	ECMO	Cold fluid infusion	ECMO+ex vivo cardiac perfusion device	Ex vivo cardiac perfusion device
Brain circulation prevented	No	No	Cross-clamp via aortic arch vessels	No
W-LST:			NA for <sup>4</sup> for <sup>7</sup>	
▶ Where	▶ In the OR	▶ In the OR	▶ In the ICU	▶ In the OR or ICU
▶ By whom	▶ NA	▶ By the ICU physician	▶ By the ICU team	▶ By the ICU team
▶ How	▶ NA	▶ Extubation	▶ Extubation	▶ NA
Stand-off period	5'	1p: 3', 2 p: 75 s	5' <sup>4</sup>	1p: 5'; 2 p: 2'
Criteria used to diagnose death	No activity on ECG No spontaneous respiration No brain stem reflexes	No cardiocirculatory functions by auscultation and arterial pulsation	NA	Cessation of circulation
Restart of heart function	In recipient	In recipients	In the donor for 50' <sup>7</sup> +ex vivo	Ex vivo

DCDD, donation after circulatory determination of death; ECMO, extracorporeal membrane oxygenation; ICU, intensive care unit; NA, not available; OR, operating room; p, patient; W-LST, withdrawal of life-sustaining treatment; y.o, years old.

infusion, there is no possibility of restoring cardiac or cerebral function with this method. This method was chosen by the team in Denver.<sup>2</sup>

### A reduced stand-off period

In a majority of DCDD protocols, a stand-off period of 5 min is used, but the stand-off period varies between 2 and 20 min depending on the DCDD protocol.<sup>12</sup> Because the heart is sensitive to WIT, a reduced stand-off period was used in the heart DCDD protocols of Denver (75 s and 3 min)<sup>2</sup> and Sydney (2 min for two donors).<sup>3</sup>

### Ex vivo perfusion

To decrease the cold ischaemia time, that is, the time from heart explantation to heart transplantation, the Australian team has used the 'normothermic ex vivo cardiac perfusion device',<sup>3</sup> also used by the Cambridge team.<sup>4</sup>

### ANALYSIS OF THE DEAD DONOR RULE IN HEART DCDD

The dead donor rule (DDR) states that vital organ procurement must not kill the donor and therefore the donor must be dead before organ procurement begins. Several scholars have raised concerns that the DDR is violated in DCDD programmes because the donor is not dead at the moment of donation.<sup>13 14</sup>

This issue continues to stir controversy. Obviously, whether heart DCDD protocols respect the DDR depends on the choice of the criterion of death. In our analysis, we analyse each heart DCDD protocol according to six plausible criteria of death (see table 2):

1. Irreversible cessation of all bodily cells function (or organs)
2. Irreversible cessation of heart function
3. Irreversible cessation of circulation
4. Permanent cessation of circulation
5. Irreversible cessation of brain function
6. Permanent cessation of brain function.

This analysis is necessary because, as Veatch stated, 'virtually all observers have assumed that DCDD could, in principle, provide any vital organs except hearts'<sup>5</sup> and because several scholars questioned the validity of the diagnosis of death in heart DCDD.

### Irreversible cessation of all bodily cells function (or organs)

Death can be defined as an irreversible event, or as a gradual process<sup>8</sup> 'that occurs over a continuum of time',<sup>15</sup> beginning with organs sensitive to ischaemia, such as the brain and the heart, and finishing with to the very last cells of the body, such as the skin.

If death were determined using the criterion of irreversible cessation of all bodily cells function (or organs), neither donation after the brain determination of death (DBDD) nor DCDD would respect the DDR. In DBDD although the functions of the whole brain have been irreversibly lost,<sup>16</sup> some brain cells remain alive.<sup>17</sup> Furthermore, in the context of DBDD, all other organs are functional because circulatory and respiratory functions are sustained by life-sustaining therapy. In DCDD, the functions of organs have not been lost irreversibly because organs regain function once transplanted.

In practice, death is not determined by the criterion of irreversible cessation of all bodily cells function (or organs). Typically in the intensive care unit (ICU) setting, ICU physicians declare death after 2–10 min of cessation of cardiocirculatory functions,<sup>12</sup> and thus long before the point at which the function of all bodily cells has ceased. This criterion of death is unnecessarily conservative, is incompatible with usual medical practice and eliminates all organ transplantation.<sup>13</sup>

### Irreversible cessation of heart function

Death is often declared using the criterion of irreversible cessation of heart function. Because the heart is the organ that produces circulation, once the heart stops and circulation ceases (assuming no medical intervention), the function of all the other organs ceases progressively, commencing with the brain.

The term *irreversible* implies that no technology (such as defibrillation or the use of ECMO) or action (such as cardiopulmonary resuscitation (CPR)) can restore heart function, which is thus irretrievably lost. Using this criterion, a deceased organ donor who has been previously determined dead by irreversible cessation of heart function cannot donate her/his heart for transplantation because the heart will have lost any possibility of functional restoration, and thus cannot be used for successful transplantation. If heart transplantation were to succeed in such

**Table 2** Choices for a criterion of death

Terminology	Consequences
Irreversible cessation of all bodily cells functions (or organs)	<ul style="list-style-type: none"> <li>▶ All bodily cells have irreversibly ceased to function</li> <li>▶ Incompatible with transplantation in general, including cornea, DBDD, DCDD and heart DCDD</li> </ul>
Irreversible cessation of heart function	<ul style="list-style-type: none"> <li>▶ Diagnosis based on the cessation of heart function (flat ECG, no heart beat on auscultation, no heart contraction on echocardiography) and on irreversibility (the heart cannot regain function by any means or actions)</li> <li>▶ Incompatible with DBDD, DCDD and heart DCDD</li> </ul>
Irreversible cessation of circulation	<ul style="list-style-type: none"> <li>▶ Diagnosis based on the cessation of circulation (flat arterial line, no central pulse, no opening of the aortic valve on echocardiography) and on irreversibility</li> <li>▶ Incompatible with DCDD and heart DCDD</li> </ul>
Permanent cessation of circulation	<ul style="list-style-type: none"> <li>▶ Diagnosis based on the cessation of circulation (flat arterial line, no central pulse, no opening of the aortic valve on echocardiography)</li> <li>▶ Permanency prohibits the use of any techniques that would restore circulation after death (ECMO, CPR) and requires a minimal stand-off period of 5 min to ensure that the possibility of autoresuscitation has passed</li> <li>▶ Compatible with heart DCDD if the stand-off period is <math>\geq 5</math> min and if ECMO is not used</li> </ul>
Irreversible cessation of brain functions	<ul style="list-style-type: none"> <li>▶ Diagnosis based on the cessation of brain functions (brain death tests, including the brain stem tests) and irreversibility</li> <li>▶ Incompatible with DCDD and heart DCDD</li> </ul>
Permanent cessation of brain functions	<ul style="list-style-type: none"> <li>▶ Diagnosis based on the cessation of brain functions (brain death tests, including the brain stem tests)</li> <li>▶ Permanency prohibits the use of any techniques that would restore circulation after death (ECMO, CPR) and requires a minimal stand-off period of 5 min to ensure that the possibilities of autoresuscitation and awareness have passed</li> <li>▶ Compatible with heart DCDD if the stand-off period is <math>\geq 5</math> min and if ECMO is not used</li> </ul>

CPR, cardiopulmonary resuscitation; DBDD, donation after the brain determination of death; DCDD, donation after circulatory determination of death; ECMO, extracorporeal membrane oxygenation.

a case, the donor ‘cannot have been previously declared dead’.<sup>5, 14</sup> The criterion of irreversible cessation of heart function is thus incompatible with heart DCDD programmes. Furthermore, heart electrical and contractile function may be restored hours after circulatory cessation, as proven by the experience of successful heart transplantation.<sup>3</sup> Thus, the time interval necessary to reach an irreversible cessation of heart function is incompatible with heart DCDD programmes, as well as with DCDD programmes for other organs.

### Irreversible cessation of circulation

The irreversible cessation of heart function is a traditional criterion of death, stated most explicitly in our technological era as the irreversible cessation of cardiocirculatory function. When heart function ceases, circulation ceases rapidly followed by the cessation of respiratory and brain functions. In the context of cardiocirculatory cessation, health professionals (depending on the clinical situation and their clinical habits) determine death by the cessation of heart function (using a stethoscope or waiting for an ECG flat line), by the cessation of circulatory function (detecting an absent central pulse or waiting for an arterial flat line), by the cessation of respiratory function and/or by the cessation of brain functions (confirming for instance the absence of consciousness or of the absence of pupillary reflex to light or dark).

Let us consider heart transplantation in the context of the irreversible cessation of cardiocirculatory function.<sup>18</sup> When circulation is sustained by extracorporeal support, there is a moment at which there is no heart in the recipient’s body. Because circulation is maintained, the recipient cannot be considered dead during the surgical procedure. Thus, what is essential to determine death is not the absence of heart function but rather the absence of circulation.<sup>18</sup>

According to the criterion of cessation of circulation, heart DCDD respects the DDR because even if the donor’s heart resumes function in the recipient, the donor’s circulation has been lost.<sup>18</sup> However, if death is determined by the *irreversible* cessation of circulation, this criterion of death does not sustain scrutiny because irreversibility means that no action or

technology can reverse the lost functions. However, extracorporeal support can restore circulation after it has stopped, even after a long period of circulatory arrest. Further, the heart function, and thus the circulatory function in consequence, can be restored by the use of appropriate resuscitation technology (CPR or extracorporeal–cardiopulmonary resuscitation) after a long period of cardiac arrest.<sup>19</sup>

If the criterion of irreversible cessation of circulation were used, then DCDD donors (including heart DCDD donors) would not be dead<sup>14</sup> and the DDR would be violated. Consequently, some scholars advocate abandoning DCDD programmes or abandoning the DDR.<sup>6</sup> One of us (JLB) proposed that death determination should be based on the *permanent* cessation of circulation rather than on its *irreversible* cessation.<sup>18</sup>

### Permanent cessation of circulation

Permanent cessation of a function means that it will not resume because it will not restart spontaneously and no medical interventions will be conducted to restart it. Whereas irreversible cessation means that the function *cannot* resume, permanent cessation of a function means that it *will not* resume.<sup>20</sup> Irreversibility (the impossibility to regain function) is a quality that is independent of action or intent. Permanency (that a function will not resume), because it admits the possibility of regaining function, is dependent upon action and intent.<sup>20</sup>

In their analysis, the American Thoracic Society stated that ‘in the context of DCDD, the tension between the need for both living organs and dead donors has required the development of very explicit criteria for declaring the moment of death, despite the absence of a biological basis for this degree of precision’.<sup>15</sup> In practice, physicians often declare death before the state of irreversible death has been reached, using prevailing medical standards. In the ICU setting for instance, it is the norm for physicians to declare death once circulation has permanently ceased; physicians do not wait until the heart or circulation has irreversibly ceased.

In the USA, the stand-off period of DCDD programmes was chosen assuming a criterion of permanent cessation of circulation, that is, when cardiac autoresuscitation is no longer

possible, but while the heart remains amenable to resuscitation that will not be attempted. A systematic review showed no reported cases of autoresuscitation after withdrawal of life-sustaining treatment (when CPR was not performed), and no reported cases of autoresuscitation beyond 7 min after failed CPR.<sup>21</sup> In a review, an expert consensus panel stated that no cases of auto-resuscitation had been reported after 65 seconds<sup>18</sup> to 2 minutes<sup>8</sup> of cardiac arrest. Thus, in the context of controlled DCDD after withdrawal of life-sustaining treatment, death can confidently be determined after 5 minutes of asystole, because the cessation of circulation is permanent and will inevitably become irreversible<sup>20</sup>

Although this purported change in death criteria has been criticised,<sup>5 13 14</sup> it conforms to the usual practice of death determination using the circulatory criterion in non-donation circumstances. This criterion remains valid only if no attempts to restore circulation,<sup>22</sup> such as ECMO or CPR, are performed after the declaration of death, to avoid negating the previous declaration of death. DCDD programmes that resume circulation after death declaration using ECMO,<sup>1 4 7 9</sup> such as in the heart DCDD protocols of Cape Town<sup>1</sup> and Cambridge,<sup>4 7</sup> are incompatible with death determination using the criterion of permanent cessation of circulation.<sup>5</sup>

The Denver heart DCDD protocol<sup>2</sup> raised the concern that 'circulation was not permanently lost after 75 s before pronouncing death',<sup>5 18</sup> because autoresuscitation might have been possible after such a short stand-off period. The Australian heart DCDD protocol,<sup>3</sup> which reported a stand-off period of 2 min for two patients, raised similar concerns that autoresuscitation might have been possible. The Society of Critical Care Medicine stated that 2 min was sufficient and more than 5 min was unnecessary.<sup>23</sup> Although we agree that a 2-min stand-off period is probably sufficient to prove the impossibility of autoresuscitation in controlled DCDD,<sup>21</sup> we continue to recommend a 5-min stand-off period for heart DCDD to be absolutely certain that the cessation of circulation is permanent. This prudential recommendation acknowledges that the autoresuscitation database remains small and incomplete, and therefore lacks a high confidence limit. It also takes into account that physicians continue to report cases of autoresuscitation, including following withdrawal of life-sustaining therapy (W-LST).<sup>24</sup> Thus, if permanency is used as a criterion to determine death in DCDD programmes, there is an urgent need for studies analysing after how many minutes following W-LST the possibility of autoresuscitation vanishes. Such studies will permit the determination of a less arbitrary stand-off period for DCDD programmes, which may be particularly relevant for heart DCDD programmes that depend on a short stand-off period for organ preservation.

By this analysis, heart DCDD protocols that violate a stand-off period of at least 2–5 min, such as Denver and perhaps as Australia (for two patients), or who start ECMO after death declaration, such as Cape Town and Cambridge, may violate the DDR (but further studies are needed to confirm the stand-off period necessary to preclude the possibility of autoresuscitation). The only heart DCDD protocol that fully respects the DDR is that of Australia.

### Irreversible cessation of brain functions

Another possibility is to use brain death criteria in DCDD programmes instead of cardiac or circulatory death criteria.<sup>5</sup> When cardiocirculatory function ceases and persists long enough, all brain functions cease. But diagnosing death in DCDD using the criterion of irreversible cessation of brain functions raises two

questions: (1) which diagnostic tests should be used to determine death? and (2) how long is it necessary to wait to achieve irreversible cessation of brain function?

Are brain death tests valid in the context of DCDD? Brain death tests were not developed for the circumstance of profound and global brain damage in which circulatory and respiratory functions are sustained medically. In Switzerland for instance, brain death tests are used to determine death in the context of DCDD.<sup>25</sup> However, we believe that, although the use of brain death tests in the context of DCDD might prove the cessation of brain function, they cannot prove that the cessation of brain function is irreversible. Before performing brain death tests, reversible metabolic or toxic disorders that might induce a transitory cessation of brain functions, and thus that might mimic brain death must be excluded but this is impossible in DCDD.

Further, the stand-off period necessary to achieve irreversible cessation of all brain functions is unknown, but stand-off periods of 5 or even 10 min are insufficient to achieve the irreversible cessation of brain function. Evidence from patients suggests that neurons can survive for up to 20 min after circulation has ceased,<sup>19</sup> and animal results suggest that brain function can return after as much as an hour without circulation.<sup>26</sup> Thus, heart DCDD protocols using a stand-off period of 5 min or less do not respect the DDR according to the criterion of irreversible cessation of brain functions.

### Permanent cessation of brain functions

As we showed in the debate over whether cessation of circulation must be irreversible or permanent, some scholars have argued that, in the setting of systemic circulatory arrest, death could be determined using the permanent cessation of brain functions. The issue would then be 'not whether the body or brain circulation and function can be resumed (because it can), but rather, whether it will be'.<sup>8</sup> Using a permanency criterion for brain death would mean that no actions (such as the use of ECMO) will be implemented to restore brain circulation, even if brain function is not irreversibly lost, because resuscitation manoeuvres might restore some brain function.

If we accept a determination of brain death based on permanent cessation, how long a stand-off period is sufficient to ensure permanency? It must be sufficiently long to ensure that the period during which autoresuscitation can occur has elapsed (as it is the case for the criterion of permanent cessation of circulation) and that all brain functions have been lost, to exclude the possibility that a donor could feel pain or have awareness. A stand-off period of 5 min would be of sufficient duration to exclude the possibility of autoresuscitation but the minimum duration is unknown. Within seconds of circulatory arrest, electroencephalographic activity becomes flat, and all brain functions cease within minutes. To ensure that the possibility of awareness or the feeling of pain has been lost, we propose a stand-off period of 5 min, despite the fact that it is an arbitrary limit. DCDD programmes need studies to demonstrate with confidence after how many minutes of brain circulatory cessation all brain functions cease. If this time is situated between 2 and 5 min, a 5 min stand-off period is a more confident limit.

Furthermore, because the brain may be responsive to restoration of its blood supply<sup>22</sup> after a 5 min stand-off period, the use of ECMO or any technology that might resume brain circulation must be prohibited.

Using the criterion of permanent cessation of brain functions, heart DCDD protocols that permit a stand-off period shorter than 2–5 min may violate the DDR (in Denver, and perhaps in

**Table 3** An analysis of four heart DCDD protocols according to six criteria of death. When is the DDR respected?

	When the criteria of death are based on irreversibility	Permanent cessation of circulation	Permanent cessation of brain functions
Barnard, 1967, Cap Town <sup>1</sup>	The 5 min stand-off period does not ensure the irreversible cessation of functions	DDR not respected because of the use of ECMO after death	DDR not respected because of the use of ECMO after death
Boucek <i>et al.</i> <sup>2</sup> , Denver	The 75 s stand-off period does not ensure the irreversible cessation of functions	DDR not respected: a 75 s stand-off period is not sufficient to ensure permanency	DDR not respected: a 75 s stand-off period is not sufficient to ensure permanency
Press, 2015, Cambridge <sup>4</sup> Description, Ali <i>et al.</i> <sup>7</sup>	The 5 min stand-off period does not ensure the irreversible cessation of functions	DDR not respected because of the use of ECMO	DDR not respected because of the use of ECMO
Dhital <i>et al.</i> , 2015, Sidney <sup>3</sup>	The 2–5 min stand-off period does not ensure the irreversible cessation of functions	DDR respected when a stand-off period of 5 min was used, but may not be respected when a stand-off period of 2 min was used	DDR respected when a stand-off period of 5 min was used, but may not be respected when a stand-off period of 2 min was used

DCDD, donation after circulatory determination of death; DDR, dead donor rule; ECMO, extracorporeal membrane oxygenation.

Australia for two patients); heart DCDD protocols that used ECMO without proving that brain circulation was excluded may have violated the DDR (Cape Town). The Cambridge protocol used an aortic arch clamp that probably excluded brain circulation. If this exclusion can be demonstrated, it probably respected the DDR. But although an aortic arch clamp blocks blood flow through both carotid and vertebral arteries, it may spare small collateral arteries from the segmental spinal arteries that arise from the thoracic aorta and anastomose with branches of the vertebral arteries. These collaterals conceivably could provide a little perfusion to the brainstem. To ensure that the aortic arch clamp completely blocks brain perfusion, brain circulation should be tested and proven absent. In particular, brain stem perfusion should be proven to have stopped because its function can continue with a smaller blood flow than the cerebral hemispheres.

Based on a criterion of permanent cessation of brain functions, only the heart DCDD protocol from Australia respected the DDR because it used a stand-off period of 5 min for one patient and did not employ ECMO. Table 3 compares the four described heart DCDD protocols in relation to six described criteria of death. Only the criteria of death based on the notion of permanency allow some heart DCDD protocols to respect the DDR.

## CONCLUSION

Our analysis suggests that some heart DCDD protocols likely violate the DDR because they use a shortened stand-off period or ECMO. In DCDD, only the criterion of death based on permanency (either the permanent cessation of circulation or the permanent cessation of brain functions) respects the DDR. If health professionals who practice heart DCDD protocols wish to comply with the DDR, they should determine death either based on the criterion of permanent cessation of circulation or permanent cessation of brain functions. But certain conditions must be respected. If the criterion of permanent cessation of circulation is used, heart DCDD protocols must follow two conditions: (1) use a minimum stand-off period of 5 min to ensure that the possibility of autoresuscitation has elapsed (unless future studies prove otherwise); (2) strictly avoid the use of any actions or technology that may resume bodily circulation.

If the criterion of permanent brain death is used, heart DCDD protocols must respect a minimum stand-off period of 5 min (unless future studies prove otherwise) to exclude the possibility of autoresuscitation, and to ensure that all brain

functions have been lost. The use of any technology that might preserve brain circulation, even partially, is forbidden. Thus, the use of ECMO would be permissible only if it confidently excluded brain circulation, including the brainstem. This condition would be achieved only if the use of ECMO were accompanied by a mechanism to prove that there was absent brain circulation. Until this is achieved, it is not defensible to merely ‘assume’ that brain circulation has ceased, and thus ECMO should be avoided.

According to our analysis, of the four heart DCDD programmes, only Australia using a stand-off period of 5 min respects the DDR when death is based on a criterion of permanent cessation of circulation. It remains unclear if heart DCDD programmes that use a stand-off period of 2 min, such as Australia’s for two patients, respect the DDR.

**Contributors** ALDA conceived the idea for the article, performed the literature search and read the papers, wrote the first draft of the article and contributed to the edits of subsequent revisions. JLB and DS worked with ALDA to develop the lines of arguments of the article, and edited and rewrote sections of the article during an iterative series of drafts. The three authors take responsibility for the arguments presented in the article.

**Disclaimer** The opinions expressed are the views of the authors and do not reflect the policy of their related institutions, the Federal Office of Public Health, or any national organisations/associations.

**Competing interests** None declared.

**Provenance and peer review** Not commissioned; externally peer reviewed.

## REFERENCES

- Barnard CN. A human cardiac transplant: an interim report of a successful operation performed at Groote Shuur Hospital, Cape Town. *S Afr Med J* 1967;41:1271–4.
- Boucek MM, Mashburn C, Dunn SM, *et al.* Pediatric Heart transplantation after declaration of cardiocirculatory death. *N Engl J Med* 2008;359:709–14.
- Dhital KK, Iyer A, Connellan M, *et al.* Adult heart transplantation with distant procurement and ex-vivo preservation of donor hearts after circulatory death: a case series. *Lancet* 2015;385:2585–91.
- Walsh F. Europe’s first non-beating heart transplant. BBC, 26th of March 2015. <http://www.bbc.com/news/health-32056350>
- Veatch RM. Transplanting hearts after death measured by cardiac criteria: the challenge to the dead donor rule. *J Med Philos* 2010;35:313–29.
- Truog RD, Miller FG. The dead donor rule and organ transplantation. *N Engl J Med* 2008;359:674–5.
- Ali A, White P, Dhital K, *et al.* Cardiac recovery in a human non-heart-beating donor after extracorporeal perfusion: source for human heart donation? *J Heart Lung Transplant* 2009;28:290–3.
- Shemie SD. Clarifying the paradigm for the ethics of donation and transplantation: was ‘dead’ really so clear before organ donation? *Philos Ethics Humanit Med* 2007;2:18.

- 9 Rojas-Peña A, Sall LE, Gravel MT, *et al.* Donation after circulatory determination of death : the University of Michigan Experience with extracorporeal support. *Transplantation* 2014;98:328–34.
- 10 Butler AJ, Randle LV, Watson CJ. Normothermic regional perfusion for donation after circulatory death without prior heparinization. *Transplantation* 2014;97:1272–8.
- 11 Snoeijs MG, Dekkers AJ, Buurman WA, *et al.* In situ preservation of kidneys from donors after cardiac death: results and complications. *Ann Surg* 2007;246:844–52.
- 12 Dhanani S, Hornby L, Ward R, *et al.* Variability in the determination of death after cardiac arrest: a review of guidelines and statements. *J Intensive Care Med* 2012;27:238–52.
- 13 Rady MY, Verheijde JL. No-touch time in donors after cardiac death (nonheart-beating organ donation). *Curr Opin Organ Transplant* 2013;18:140–7.
- 14 Marquis D. Are DCD donors dead? *Hastings Cent Rep* 2010;40:24–31.
- 15 Gries CJ, White DB, Truog RD, *et al.* An official American thoracic society/ international society for heart and lung transplantation/society of critical care medicine/association of organ and procurement organizations/united network of organ sharing statement: ethical and policy considerations in organ donation after circulatory determination of death. *Am J Respir Crit Care Med* 2013;188:103–9.
- 16 Bernat JL. The whole brain concept of death remains optimum public policy. *J Law Med Ethics* 2006;34:35–43.
- 17 Wijidicks EF, Pfeifer EA. Neuropathology of brain death in the modern transplant era. *Neurology* 2008;70:1234–7.
- 18 Bernat JL, Capron AM, Bleck TP, *et al.* The circulatory-respiratory determination of death in organ donation. *Crit Care Med* 2010;38:963–70.
- 19 Hara M, Hayashi K, Hikoso S, *et al.* Different impacts of time from collapse to first cardiopulmonary resuscitation on outcomes after witnessed out-of-hospital cardiac arrest in adults. *Circ Cardiovas Qual Outcomes* 2015;8:277–84.
- 20 Bernat JL. How the distinction between ‘irreversible’ and ‘permanent’ illuminates circulatory-respiratory death determination. *J Med Philos* 2010;35:242–55.
- 21 Hornby K, Hornby L, Shemie SD. A systematic review of autoresuscitation after cardiac arrest. *Crit Care Med* 2010;38:1246–53.
- 22 Manara AR, Murphy PG, O’Callaghan G. Donation after circulatory death. *Br J Anaesth* 2012;108(Suppl 1):i108–121.
- 23 Kotloff RM, Blosser S, Fulda GJ, *et al.* Management of the potential organ donor in the ICU: Society of Critical Care Medicine/American College of Chest Physicians/ Association of Organ Procurement Organizations Consensus Statement. *Crit Care Med* 2015;43:1291–325.
- 24 Wind J, van Mook WN, Dhanani S, *et al.* Determination of death after circulatory arrest by intensive care physicians. A survey of current practice in the Netherlands. *J Crit Care* 2016;31:2–6.
- 25 Swiss Academy of Medical Sciences. *Diagnosis of death in the context of organ transplantation*. Ethical Guidelines. 2011.
- 26 Hossmann KA, Schmidt-Kastner R, Grosse Ophoff B. Recovery of integrative central nervous function after one hour global cerebro-circulatory arrest in normothermic cat. *J Neurol Sci* 1987;77:305–20.