に角膜を移植に使用することができる.

- b. アイバンクは、眼球あっせん業を厚生労働大臣から許可を得て いる
- c. 医師は、移植術に使用されるための臓器を、死体から摘出するこ とができる("臓器の移植に関する法律"第6条).
- d. 移植に使用されなかった臓器の処理は、焼却して行わなければな らない. ("臓器の移植に関する法律"第9条. "同施行規則"第4条).
- e. 脳死下、心臓死ともに、本人の提供に関する書面同意がなくて も、遺族の承諾があれば提供は可能である。

模範解答 a, b, e

カコモン読解 第20回 一般問題20

正しいのはどれか.

- a 医師は臓器の斡旋を行える.
- b 臓器移植法に眼球は含まれない.
- c アイバンクは知事の設立認可が必要である.
- d 眼球提供に本人の意思表示は不可欠である.
- e 心臓死の場合で遺族の承諾があれば眼球摘出できる.
- **解説** a. 臓器のあっせんはアイバンクが行う. 臓器のあっせんを 行う場合, 臓器の別ごとに、厚生労働大臣の許可を受けなければな らない("臓器の移植に関する法律"第12条).
- b. "臓器"とは、人の心臓、肺、肝臓、腎臓、その他、厚生労働省令で 定める内臓および眼球をいう("臓器の移植に関する法律"第5条).
- c. アイバンクの設立には, 厚生労働大臣の許可が必要であり, 知事 の許可は必要としない.
- d. 本人の意思表示は必ずしも必要とせず、本人の拒否の意思表示 がない限り、遺族の承諾で提供が可能である.
- e. 脳死下、心臓死ともに、本人の提供に関する拒否の意思表示がな い限り、遺族の承諾があれば摘出は可能である.

模範解答 e

(井原正裕, 青木 大)

特集「拡大臓器提供推進委員会 2012 年度」

厚生科研における「移植医療の 社会的基盤に関する研究」の概要

篠崎尚史,青木 大

厚生労働省科学研究費補助金免疫アレルギー疾患等予防・治療研究事業「移植医療の 社会的基盤に関する研究」研究代表者,東京歯科大学市川総合病院角膜センター



| はじめに

わが国の臓器移植は、改正臓器移植法の施行に伴い、脳死下臓器提供数は明らかに増加した。このことは改正法施行後に発生した事例のほとんどが家族承諾により提供に至っていることから、法改正の一つの効果の表れといえる。しかし、改正点は他にもあり、運転免許証・健康保険証に意思表示できるような施策も進んでいる。こちらの効果が表れるにはある程度の時間を要するが、こうした状況を鑑みると、今後わが国での臓器提供数は、徐々に増加することが予想される。同時にこのことは、移植コーディネーター不足という状況になることを示唆するものであり、移植コーディネーター、特に院内コーディネーターの教育機関の設立は急務といえる。

このような現状の中,厚生労働省科学研究費補助金 免疫アレルギー疾患等予防・治療研究事業「移植医療 の社会的基盤に関する研究」において,わが国におけ る移植医療の適正な発展に必要な社会基盤について検 討することを目的として研究がなされている。



研究目的

本研究では、コーディネーター教育機関の設立に向けた基盤整備として、教育プログラムの設計、教育ツールの開発、指導指針の作成、実践的教育プログラムの作成を行う。また、将来的な事業化に向けての運用制度設計を実施する。また、現場に必要な教育、処遇待遇および、システムと、教育体制について研究すると

ともに、提供施設への負担軽減のための直接的、間接 的な介入研究を行う。また、関連する分担研究との連 携の中で活動を進め、臓器提供者の増加と同時に提供 家族への配慮がなされる提供施設構築を図ることを目 的としている。



| 研究概要

現在,当研究班では前項に示す目的に向け,さまざまな角度からの研究を行っている。以下,研究の概要を示す。

1. Donor Action Program (DAP)

DAPとは、臓器提供病院にフォーカスして臓器提供に至るプロセスを分析し、臓器提供に至らなかった原因を明確にし、問題点の改善策を盛り込んだ、アクションプランを作成、実行することにより、臓器提供の増加を図る実践的なプログラムである。日本では2002年度より厚生労働科学研究「病院開発モデル作成」として取り組んでいる。

具体的には、施設合意が得られ、院内に精通した担当者・委員会が設置された段階で、2つの調査を実施する。1つが、Hospital Attitude Survey(HAS:医療従事者意識調査)で、もう1つが Medical Record Review(MRR:カルテレビュー)である。HAS は、その施設のスタッフ個人の臓器移植に対する考え方、知識、どんなことにストレスを感じているか、提供システム・教育プログラムについての要望などを分析するツールである。MRR は、その症例がどういった理由により臓器提供に至らなかったのかを分析するツールであ

る。以上の調査結果を、国際データベースに入力・分 析することで,他施設,他国と比較し特徴を把握する ことができる。具体的な各プロセスは以下のとおりで ある。

- ①ターゲットを絞る
- ②DAP を導入することについて施設合意を得る
- ③施設内に DAP 委員会を設置する
- ④運用方法をデザインする
- ⑤HAS を実施し、職員の移植医療に対する意識を 解析する
- ⑥MRR を実施し、過去の死亡症例から臓器提供適 応患者の解析と、その結果を職員と共有する
- ⑦これらの情報を基に、臓器提供者の増加に向けた アクションプランを検討, 実行する
- ⑧アクションプランを評価する

臓器提供数が伸びるためには、標準的な臓器提供プ ロセスを前提に, 医療機関においてポテンシャルド ナーがどのような現象で臓器提供に至らないかを解析 し, 科学的に、そのボトルネックを解消する必要があ る。

HASでは、ニーズを把握し、ニーズに応じたセミ ナー等を実施した後に再度、HAS を実施して改善の 検証をすることができる。また、MRRでは、分析か ら浮かび上がる提供に至らない理由の中に, 改善点を 見出すことができる。これらから、その施設特有のボ トルネックを解消していく作業こそが、臓器提供増加 への道となる。DAPをどう活かすかで、終末期にお ける臓器提供の意思確認を推進することが可能とな り、ゆえに臓器提供数増加に結び付けることができる システムである。

現在, DAP 導入病院にて実施している DAP, MRR の調査データに関して、DAP財団の運営する WEB サ イトにおいてデータベース管理を行い、その分析を進 めるとともに、その結果これまでの実績から有効性の 高い施設に限定して実施し,病院全体として取り組む ための具体的手法のさらなる確立, 各都道府県コーデ ィネーターを通じて把握し,移植医療推進に対する地 方行政の行政支援の在り方について検討し整備を進め ている。その結果、HAS(33,791人/76施設)のデー タから, 医師, 看護師など医療職種においても, ①一 般に移植には賛成であり、半数近くが、死後自分の臓 器提供を希望していること,②脳死を死の妥当な判定 方法であると考えるものは、医師の約6割に比較し て、看護師、事務職では4割程度に過ぎないこと、③

ドナー候補の特定、臓器提供の同意を得ることに必要 な能力・知識を有すると考えるものは、医師で約2 割,看護師ではごく少数であること,が分かる。その ような中で、MRR(6,662人/40施設)の結果から、 全参加病院において, 家族へのオプション提示の割合 が年々着実に増加していることが分かった。一方、12 道県(新潟県,福岡県,神奈川県,沖縄県,千葉県, 富山県, 熊本県, 静岡県, 兵庫県, 福井県, 愛知県, 北海道)で61施設がDAP導入を図っている。また、 2012年4月1日~11月30日の期間において, DAP 導入地域の臓器提供は27例、うち脳死下臓器提供14 例である。全国の臓器提供数と DAP 導入都道府県の 提供の割合は、腎提供で全国95例のうち27例 (28.4%), 脳死下臓器提供の全国 33 例のうち 14 例 (42.4%) であり、心停止下、脳死下いずれの場合も 全体の3~4割がDAP導入地域で提供されていること となる。

2. 教育機関の基盤整備

現在日本で活動している臓器提供(ドナー)側の調 整を行う移植コーディネーターは、社団法人日本臓器 移植ネットワーク(以下, JOT)所属のコーディネー ター, JOT からあっせん業の委託を受けた都道府県 コーディネーター, そして提供施設に所属する院内 コーディネーターに分けることができる。加えて、組 織提供にかかわる組織移植コーディネーター,献眼に かかわるアイバンクコーディネーターも存在する。院 内コーディネーター以外は, 事例発生時に提供施設に 派遣され、事例ごとにチームが組まれる。実際の提供 事例では、「臓器・組織にかかわらず可能なものをす べて」提供したいというケースも多く、各種のコーデ ィネーターがかかわることとなる。しかし現状では、 各種コーディネーターに対する標準化されたドナー コーディネーターの教育体制は存在しない。現場では 同じ立場でかかわりながら,標準化された教育体制が ないことは望ましいことではない。

そのため、質の担保という意味でも、移植コーディ ネーターの教育に必要な,教育プログラム,教育ツー ル,教育機関運用制度等の社会基盤が必要である。

特に救急現場において, 臓器提供施設への負担の軽 減と, 臓器提供の有無にかかわらず医療機関で継続的 にご家族との連携や院内連携を行う「院内移植コーデ ィネーター」の存在が重要である。従来のポテンシャ ルドナーが発生した後に介入し,臓器提供がある場合

のみ家族ケアを行うコーディネーターではなく,医療 機関での重症患者への対応を組織的に行い,終末期医療全体の管理を行う体制整備に必要な人材育成に関しての検討を行っている。つまり,医療の中で臓器提供だけに特化することなく,救急医療現場での一連の医療行為の中で終末期に至った場合,選択肢の一つとして臓器提供が位置付けられ,質の高い家族ケアやスタッフサポート,院内教育などが実施できる人材育成が必要である。そのためのクオリティマネージャー養成を目的とした教育プログラムを作成,試行し,その内容について検証している。なお,研究を進めるにあたっては,日本救急医学会,日本臨床救急医学会の協力を得て実施している。

一方、現在、スペインでは人口 100 万人あたりの臓器提供者数が世界一であるが、その背景には家族への意思確認・家族ケアから、脳死判定・ドナー管理にまで及ぶ知識を有したコーディネーターが臓器提供施設に配置されている体制がある。そして、そのコーディネーター教育を実施しているのが Transplant Procurement Management (TPM) という教育機関であり、現在では多くの国の移植コーディネーターが TPM の教育プログラムに参加している。わが国においても、日本版 TPM のような教育機関の設置が望まれる。

3. 提供施設支援ツールの開発

移植医療の社会的基盤として, 提供施設の負担軽減 は必要不可欠なものである。提供件数の増加が見込ま れる状況では、今後わが国での移植医療が発展するか どうかの鍵は、まさしく提供施設の負担軽減といえ る。そのためのツールの開発は重要である。研究では, 提供施設内の情報共有システムの構築により、必要な タイミングで受取るべき人だけに情報を配信するシス テム, さらに, 臓器提供の各段階においての作業手順, マニュアル、法令・ガイドラインの解説が自由に参照 できるツールを開発し、提供施設スタッフの負担,不 安を軽減する。また、これまでのドナー家族が訴えた 精神的負担などを症例として蓄積し, 今後の精神的ケ アに寄与するためのデータベースの構築, そして, 脳 死下臓器提供が進むことで、これまで以上に1人のド ナーから多数のレシピエントに提供される症例が増え ることから, 双方の安全性確保のためにドナー・レシ ピエント双方の情報が登録できる統一システムを目指 した調整を行っている。これらのツールを整えること により, 提供施設の負担軽減に寄与できれば, 終末期 における医療現場での臓器提供の意思確認も進むこと が期待できる。

4. ドナー評価・管理および摘出手術の呼吸循環管理の体制整備

脳死臓器提供におけるドナー評価・管理および摘出 手術の呼吸循環管理と移植後の成績を検討し、わが国 にあったドナー評価・管理法を開発するとともに、そ こで得られた結果を基にマニュアルを作成し、メディ カルコンサルタント,提供病院のスタッフ,ドナーコー ディネーターの研修を行うとともに、そのメンバーを 基本にした体制整備を行っている。わが国では、欧米 と異なり、2002年以降メディカルコンサルタント制 度を導入しており、その結果、2012年11月20日現 在(200件の脳死臓器提供)のわが国の臓器提供率は、 心臟 74.0%, 肺 62.4%, 腎臟 95.6%, 肝臟 80.2%, 膵 臓 70.9% と高い水準を示している。臓器提供率を米 国と比較すると、腎臓は同程度で、肝臓は少なかった (脂肪肝,ショック肝が多いため)が,心臓,膵臓, 肺は3~4倍の臓器提供率である。それぞれの移植後 の成績も欧米の成績と遜色はない。呼吸器外科医を中 心とするメディカルコンサルタントがドナーの呼吸管 理に参画することで、さらに肺の提供が増加するとと もに、肺移植後の成績が向上している。スペイン、韓 国, 米国の Organ Procurement Organization (OPO) と 連携して、ドナー評価・管理システムを検討し、臓器 提供にかかわる医療者の教育・研修システムを構築中 である。

□□□ まとめ □□□

DAPの導入病院は増加傾向にあり、データ数は増加しつつある。HAS、MRR は DAPでの主要なツールであるが、全体の集計によりおおよその動向を知ることが可能であるとともに、個別病院における問題把握、介入効果判定のツールとして利用が可能である。また、症例の考察から、救命救急治療とともに家族ケアが充実しており、家族と医療者の信頼関係が十分であることが DAPを推進するうえで最も重要な要素であることが推察される。また、地域独自の取り組み、例えば、官民一体の活動の強化や MRR を多用した詳細な医療機関診断からのアプローチ、さらに家族ケアの観点から"救急における看取り医療の充実"などにより、家族にとっても、医療スタッフにとっても、満

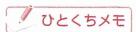
足度の高い医療が展開されていることが DAP の大き な特徴である。加えて、臓器提供者数は、わが国の人 口比 0.88 人/pmp (2010 年度) のうち 0.2 人/pmp を DAP 導入地域がカバーしている。DAP 導入地域における 活動のさらなる発展と、また非導入地域への DAP の 導入・展開によって臓器提供数は飛躍するものと考え る。また、継続的に運用しているドナー家族精神的ケ ア窓口であるが、ドナー家族が抱える精神的問題点を 詳細に発掘する為にはその運用にさらなる工夫が必要 である。一方、提供施設のスタッフの負担や不安を軽 減するための、情報を参照できるツールの開発やコー ディネーターの教育プログラムの必要性が重要な位置 付けと考えられる。

おわりに

臓器移植法の改正後,脳死下臓器提供が増加する 中,移植医療の社会的基盤としての研究により、提供 施設の負担の軽減およびコーディネーター教育が重要 な課題であることが明らかにされたが、提供現場への 具体的教育, 処遇待遇がシステムとして機能するの か、また教育体制の確立が重要である。このことが、 提供者家族が一貫した医療行為の延長上にある臓器提 供のステップを踏んで行けるための重要な鍵となる。

移植医療に関する院内システムを構築する際の介入

ポイントの設定の見直しが必要である。従来. 介入ポ イントは、患者の予後不良診断後からの動きにフォー カスされていた。しかし、救急搬入患者家族の多くは 突然の発症、すなわち非日常の出来事を受け止めなく てはならない。一方, 医療機関側からすれば救命救急 治療の限界点で移植医療が突然介在してくるのには違 和感があることは否めない。院内システム構築の際に は、むしろ患者搬入時からの取り組みこそが重要であ り、そのプロセスのなかからポテンシャルドナーを見 出し、患者に対する可能な限りの救命救急治療を提供 するのと並行して、刻々と変わる病状を受け止めなけ ればならない家族に対するケア, 救命できなかった場 合の看取りの医療から臓器提供へとつながる連続的な 流れを構築してゆくように医療機関啓発活動の内容を 見直すことが重要である。DAPの手法はある程度集 約されてきた。そのことを十分に行うことのできる人 材育成. とりわけ院内コーディネーターについてはそ の発展系の、すなわち多角的視野で組織展開ができる "クオリティマネージャー"の育成が重要である。各 分担研究ともリンクしながら院内システムとそれを遂 行するプロフェッショナルの育成が最重要課題であ り、同時に、提供施設の負担軽減のためのデータベー スシステムの構築を含め本研究を進めていくことが必 要である。



移植医療と再生医療

篠崎尚史

iPS 細胞研究が 2012 年ノーベル生理学・医学賞を受賞し、再生医療が再燃している。再生医療は夢の医療のように報道されているが、わが国の再生に関する研究はすでに 70 年近い歴史をもち、世界に大きくリードしている研究分野である。1992 年に、岡田節人、江口吾朗らと日本再生研究会を立ち上げたことが思い出される。特に江口氏の網膜再生の研究は初めて脊椎動物の中枢神経系を再生させたことで世界を驚愕させた。

再生技術を医療に用いるとなると、臨床応用への熟慮されたシステムの確立と革新的な技術が不可欠となる、移植医療は100年以上の歴史のなかで、倫理性、安全性の担保が行えるシステム構築と、それを裏付ける法的整備が行われてきた、臓器・組織の提供者の医学的、社会医学的スクリーニングとトレーサビリティ確保、臨床成績の透明化である。

しかしながら骨髄をはじめとする角膜や多くの ヒト由来の細胞・組織・臓器が国際間でシッピン グされ. さらには生活習慣病の増加などに起因す る移植待機患者の増大により非倫理的な取り引き の報告が世界各地から寄せられ, 2003年, WHO 総会¹⁾にて、この件に関する調査 (EB112/5, EB 113/14) が決定し、2004年のWHAでは、1991 年に制定された臓器移植のガイドライン (Guiding Principles on Human Organ Transplantation) の改訂を決定した²⁾. 国際的なネットワーク によるすべてのヒト由来製剤に関して、倫理性、 安全性を担保すべきであるという結論に至り、そ の過程で国際移植学会や国際腎臓学会等の協力の 下, 臓器売買, 渡航移植を禁止するという「イス タンブール宣言」の制定に漕ぎ着けた. 2005年の マニラで開催されたアジア政府間会議には、中国 の衛生部より黄副部長が参加し、 国際的な倫理的

議論がより現実的な法整備と連動し、中国でも外国人の移植を禁止する法律「人体器官移植条例」 (2007年)が制定されるに至った.

骨髄移植では、国際間のドナー・レシピエントマッチングが有効に機能したことで、国際ルールが最も早く完備された領域である。ICCBBAが制定した ISBT(International Standard for Blood and Transplant)128 という輸血で用いられたコードを使用して、トレーサビリティを確保している。WHO のガイドライン改訂の過程で、この有用性が検討され 3 、移植医療のなかでのさまざまな incident により構築したシステムについて調査、検証がなされた。

再生医療における日本の研究・技術レベルは、世界のトップクラスである.しかし、北米や欧州でも研究は加速化しており、知的財産権の獲得数も驚くべき勢いである.このような環境下でわが国の国策として実現化するためには、安全性、倫理性に十分配慮しながらも、基礎研究からシステム化された前臨床試験、非臨床試験を支援し、治験と承認作業を急ぐ必要がある.知的財産権も世界規模での戦略で獲得する必要がある.

再生医療分野は、根本的に移植医療と同様に進める医行為の部分と、製造や管理といった薬事に係る部分が両立されなければならない新しい医療分野である。特に遺伝子改変を行った細胞の際には、その安全性、有効性試験はさらに厳格に行う必要がある。どのような医療でもリスクを完全に排除することは不可能であるが、治療不可能であった分野に新たな技術が導入され、それらを治癒させることこそが医学のイノベーションである。同時に、そのリスクをいかに最小化させ、また、万が一の副作用の際に、医学的、政策的にどのように対応するかの基本ルールを明確にして、世界的競争のなかで発展させるバランスが重要である。

- 1) WHO: Ethics, access and safety in tissue and organ transplantation: Issues of global concern. 2004.
- WHA: Human organ and tissue transplantation. WHA 57.18 WHO, Geneva, 2004.
- 3) Strong DM, et al : Cell Tissue Bank 2010 ; 11 : 305–323.

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Japanese Strategies to Maximize Heart and Lung Availabilities: Experience from 100 Consecutive Brain-Dead Donors

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ABSTRACT

Objective. Because the donor shortage is extremely severe in Japan because of a strict organ transplantation law, special strategies have been established to maximize heart and lung transplantations (HTs and LTs, respectively). We reviewed 100 consecutive braindead donors to evaluate our strategies to identify and manage heart and lung donors.

Methods. We retrospectively reviewed all 100 consecutive brain-dead donors procured since the law was issued in 1997. There were 56 mens and the overall mean donor age was 43.5 years. The causes of death were cerebrovascular disease (n=62), head trauma (n=20), and asphyxia (n=16): Since November 2002, special transplant management doctors were sent to donor hospitals to assess cardiac and lung functions, seeking to identify transplant opportunities. They stabilized donor hemodynamics and lung function by administering antidiuretic hormone intravenously and performing bronchofibroscopy for pulmonary toilet.

Results. Seventy-nine HTs, 1 heart-lung transplantations, and 78 LTs (46 single and 32 bilateral) were performed. By applying these strategies organs per donor were increased from 4.5 to 6.8. Among heart donors, 61/80 were marginal: high inotrope requirement (n = 29), cardiopulmonary resuscitation (n = 28), and/or >55 years old (n = 20). None of the 80 HT recipients died of primary graft failure (PGF). Patient survival rate at 10 years after HT was 95.4%. Among lung donors, 48/65 were marginal: pneumonia (n = 41), chest trauma (n = 4), and >55 years old (n = 9). Only 2/78 LT recipients died of PGF. Patient survival rate at 3 years after LT was 72.2%. After inducing frequent pulmonary toilet, lung procurement and patient survival rates increased significantly after LT.

Conclusions. Although the number of cases was still small, the availability of organs has been greater and the outcomes of HT/LT acceptable.

respectively) represent established procedures that show in satisfying long-term results for end-stage heart and respiratory failure patients. However, these therapies are limited by severe donor organ shortages. Therefore, optimal utilization of all suitable donor organs is mandatory to increase graft availability.

In Japan, the donor shortage has been more severe than in other developed countries because of the strict Japanese organ transplantation law issued in 1997, which requires a living person to grant written consent for organ donation after brain death. Until September 30, 2010, only 100 braindead donors have been procured in Japan since the law was issue. ^{3–6} In 2007, the cardiac donation rate per million population in Japan was only 0.08, compared with 7.3 in

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USA, 5.3 in Spain, and 0.97 in South Korea. The mean waiting times for HT and LT are extraordinary long in Japan, namely, 1,026 and 1,673 days respectively, in 2010. The organ shortage and long waiting times have forced Japanese transplant programs to consider the use of hearts and lungs from donors who would be considered to be marginal.⁶

The most troublesome issue facing transplantation is primary allograft failure (PGF). This complication is the leading cause of death in the first 30 days and in the 1 year after transplantation in the world. The use of marginal donor organs seems to increase the rate of PGF. Therefore, it is necessary to establish special donor evaluation and management systems that maximize cardiac and lung donor utilization. The purpose of the present study was to review our special strategies to identify and manage heart and lung donors among 100 consecutive brain-dead donors since issuance of the Japanese organ transplantation law.

MATERIALS AND METHODS

In this study, we reviewed retrospectively 100 brain-dead donors procured in Japan from October 17, 1997, to September 30, 2010, 56 were male, and the ages of the donors range from 18 to 72 years (mean, 43.3 ys). The cause of brain death was cerebral stroke in 61, including 52 with subarachnoid hemorrhage, 5 cerebral infarctions, and 4 cerebral bleedings, head trauma in 20 including 10 motor vehicle accidents, 16 in stavces of asphyxia, and 4 of brain injury after cardiopulmonary resuscitation.

Donor Evaluation and Management System

Organ transplantation from brain-dead donors started in 1999. Since then, each organ procurement team includes staff physicians who evaluate the condition of donor heart and lungs by echocardiography and flexible bronchofibroscopy (BFS) in the intensive care unit before the procurement operation. Since November 2002, a medical consultant (MC) has been sent to the procurement hospital, to also assess donor organ function and identify suitable organs for transplantation. They also intensively care for the donor, stabilizing hemodynamics by administering a bolus infusion of 0.01 U/kg followed by a drip infusion of 0.01 U/kg/h of antidiuretic hormone (ADH) and reducing intravenous catecholamine doses as much as possible. They seek to prevent or to treat lung infections before the procurement teams arrive.

If the ratio of arterial oxygen tension to inspired oxygen fraction $(PaO_2\text{-}FiO_2 \text{ ratio})$ was <300, one lung was transplanted if the $PaO_2\text{-}FiO_2$ ratio of the pulmonary venous blood of that side sampled at the procurement operation was >400.

After the 50th brain dead donor in December 2006, we modified the lung management. While, regular toilet and turning of the donor were performed as previously, repeated BFS was carried out when there were symptoms and/or signs of atelectasis or pneumonia on the chest X-ray or computerized tomography (CT) scan.

We defined a marginal donor heart as one from a donor with a history of cardiopulmonary resuscitation >5 minutes, with left ventricular dysfunction defined via transthoracic echocardiography demonstrating left ventricular ejection fraction <50%, with high inotrope requirement defined as a sustained need for dopamine $>10\mu g/kg/min$, or >55 years of age.

We defined as marginal donor lung was among from a donor with infections sputa or findings of pneumonia by chest-X-ray, who is hemodynamically unstable, who sustained chest trauma, or >55 years age.

Preprocurement Meeting and Management of Procurement Operation

Before starting a procurement operation, all surgeons, anesthesiologists, operating rate and nurses gathered in the meeting room. They negotiated the types of procured organs, how to procure each organ (eg, organ dissection/perfusion technique, incision lines, blood drainage technique), needed samples (eg, blood, lymph nodes, spleen), and donor management during the operation. Because most Japanese anesthesiologists have never experienced a procurement operation from a brain-dead donor. The MC also supported them to stabilize donor hemodynamics during the operation. Skillful staff surgeons, not resident surgeons, harvested the donor organs.

Because an inverse relationship between volume of intraoperative colloid and early lung allograft function has been reported, packed red blood cells and albumin were transfused to maintain the circulating blood volume and to replace proteins and fluids. To improve organ perfusion with preservation solution, additional catecholamine was not administered (as possible to dilate the vessels of organs). ADH and all catecholamine infusions were discontinued at the time of the heparin sulfate (400 U/kg) bolus.

RESULTS

Among 100 brain-dead donors, we obtained 79 HTs (80.0%), 78 LTs (32 bilateral and 46 single) from 65 donors (58.7%), and 1 heart and lung transplantations (HLT).

Heart Transplantation

Seventy-nine HTs were performed at 7 centers, 58 of them male. The overall age of the HT recipients was 8–60 years (means 36.7 ys). Their underlying diseases were dilated cardiomyopathy (DCM; n=60), dilated hypertrophic cardiomyopathy (n=6), ischemic cardiomyopathy (n=6), secondary DCM (n=5), restrictive cardiomyopathy (RCM; n=1), and in complex cardiac anomaly (n=1). All patients were transplanted under status 1 hemodynamic condition with 69 requiring left ventricular assist support for a mean duration of 821 days (range 21, 1,593). The mean waiting period for HT was 950 days (range 29, 2,595).

Among 79 HTs, 61 donors were marginal including 35 treated with high-dose catecholamines, 4 with <50% left ventricular ejection fraction, 29 with history of cardiopulmonary resuscitation, and 9 > 55 years old without coronary angiography.

Although 3 patients required mechanical support (2 for extracorporeal membrane oxygenation and 1 for intraaortic balloon pumping) and 2 required high dose inotropic support, none of the 79 HT recipients died from PGF. Two succumbed to infections at 3 months and 4 years after HT. Patient survivals at 1, 5, and 10 years after HT were 98.7%, 96.2%, and 96.2%, respectively (Fig. 1).

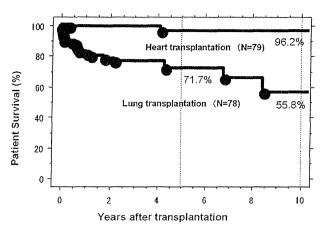


Fig 1. Patient survival after heart and lung transplantation.

Lung Transplantation

The 78 LTs included 32 bilateral (BLT) and 46 single (SLT) LTs at 7 centers, the in of 29 men. The overall age/range of the LT recipients was 19–60 years (mean, 38.3 ys). The underlying disease for LT was lymphangioleiomyomatosis (LAM; n=28) primary pulmonary hypertension (n=16), idiopathic or other interstitial pneumonia (n=14), bronchiolitis obliterans (n=4), Eisenmenger syndrome (n=3; ventricular septal defect, 1 partial pulmonary venous anomalous drainage, and 1 patent ductus arteriosus), emphysem (n=6), bronchoectasia (n=3), or another reasons (n=3). The waiting period for LT was 22–2,345 days (mean, 1,056 ds).

Among 65 lung donors, 48 were considered to be marginal including 32 due to infectes sputa or pneumonia by chest-X-ray or CT scan. There were 7 hemodynamically unstable donors, and 6 had experienced chest trauma. Nine were >55 years old.

Among the 8 recipients who, died early after LT, 4 succumbed due to PGF, 2 due to technical reasons, and 2 due to sepsis. Patient survivals at 1, 5, and 10 years after LT were 82.5%, 71.7%, and 55.8 %, respectively (Fig 1). The 1 HLT performed for Eisenmenger syndrome associated with double-outlet right ventricle, was alive after 2 years.

After the 50th brain dead donor, lungs management was modified as described above. Before modifying lung management, 29 LTs (13 SLTs and 16 BLTs) were performed from 27 donors. After modifying lung management, 49 LTs (33 SLTs and 16 BLTs) were performed from 38 donors. Lung procurement rate per donor significantly increased to 74.5% from 55.1% after the modification (Pearson, chi-square test: P = .04). However, there was no difference in the rate of early death after LT due to PGF and other causes. Patient survival at 1, 2, and 4 years after LT significantly increased after the modifications from 89.1%, 85.8%, and 85.8% before to 72.4%, 65.5%, and 62.1% respectively, after modification, (log rank test: P = .02; Fig 2).

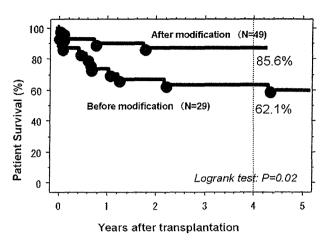


Fig 2. Patient survival after lung transplantation before and after modification of donor management.

DISCUSSION

For many years, HT and LT have represented established procedures for end-stage heart and respiratory failure patients. Over the past 2 decades, there has been a considerable increase in the numbers of patients annually listed for these procedures. Strict adherence to "standard donor criteria" resulted in a undersupply of available organs with the result of significantly extending waiting times and increasing waiting list mortality. ^{2.8}

As a consequence of this severe shortage of donor organs, hearts and lungs from marginal donors have been utilized in many countries. However, only 2,407 hearts from 7,944 deceased donors (30.2%) were transplanted in the USA in 2010. Because of the strict Japanese organ transplantation law only 100 brain-dead donors have been procured in Japan in >10 years. Only 30 HTs would have been transplanted, if the cardiac donation rate was the same as in the USA. These great pressures of the organ shortage made transplant programs consider the use of the hearts and lungs that would be considered to be marginal. Therefore, original donor evaluation and management systems have been established, such as the MC and the preprocurement meeting.⁶

High serum adrenaline concentrations have been reported to reduce myocardial β -adrenergic receptor density in brain dead nonhuman animals and human patients. These concentrations can increase the risk of PGF after HT. Therefore, the dose of intravenous catecholamine should be reduced as much as possible. The American College of Cardiology has recommended it as the initial therapy for hemodynamic support after failure of treatment of diabetes insipidus, which shows catecholamine-sparing effects. Vasopressin replacement treats diabetes insipidus as well as maintains hemodynamic stability and prevents electrolyte imbalance. A substantial number of brain-dead donors show resolution of their focal/regional wall motion abnormalities. Aggressive attempts at hemodynamic stabilization with the use of hormonal resuscitation

have dramatically improve reversed cardiac function and yield. $^{\rm [2,14]}$

The ideal lung donor shows a PaO_2 -FiO₂ ratio of >300, a positive end-expiratory pressure requirement >5 cm H_2O , a clear chest X-ray, age >55 years, a smoking history of >20 packs/year, and absence of trauma, surgery, aspiration, malignancy, and purulent secretions. Pathologic studies of lungs deemed to be unsuitable for donation have indicated that bronchopneumonia, diffuse alveolar damage, and lung consolidation are the most common reasons to reject a lung. Given these findings, it is recommended that every lung donor undergo bronchoscopy for a therapeutic toilet and to isolate potential pathogens to guide antibiotic therapy in both the donor and the recipient. 12

In the present study, after modifying lung management, lungs procured per donor significantly increased and, moreover, with significantly increased patient survival.

In conclusion, although the number of transplantations is still small, the availability of hearts and lungs has been greater in Japan than in other developed countries the outcomes of HT and LT were acceptable. These strategies may be useful to maximize HT and LT opportunities in other countries.

REFERENCES

- 1. Christie JD, Edwards LB, Kucheryavaya AY, et al. The Registry of the International Society for Heart and Lung Transplantation: Twenty-seventh Official adult lung and heart-lung transplant report-2010. *J Heart Lung Transplant*. 2010;29(10):1104-1118.
- 2. Wittwer T, Wahlers T. Marginal donor grafts in heart transplantation: lessons learned from 25 years of experience. *Transpl Int.* 2008;21(2):113–125.

- 3. Matsuda H, Fukushima N, Sawa Y, et al. First brain dead donor heart transplantation under new legislation in Japan. *Jpn J Thorac Cardiovasc Surg.* 1999;47(10):499-505.

 4. Fukushima N, Miyamoto Y, Ohtake S, et al. Early result of
- 4. Fukushima N, Miyamoto Y, Ohtake S, et al. Early result of heart transplantation in Japan: Osaka University experience. *Asian Cardiovasc Thorac Ann.* 2004;12(2):154-158.
- 5. Bando T, Gate H, Minami M, et al. First registry report: lung transplantation in Japan: the Japanese Society of Lung and Heart-Lung Transplantation. *Gen Thorac Cardiovasc Surg.* 2008;56(1):17–21.
- 6. Fukushima N, Ono M, Nakatani T, et al. Strategies for maximizing heart and lung transplantation opportunities in Japan. *Transplant Proc.* 2009;41(1):273–276.
- 7. Mcllroyl DR, Pilcher DV, Snell GI. Does anaesthetic management affect early outcomes after lung transplant? An exploratory analysis. *Br J Anaesth*. 2009;102(4):506–514.
- 8. Lima B, Rajagopal K, Petersen RP, et al. Marginal cardiac allografts do not have increased primary graft dysfunction in alternate list transplantation. *Circulation*. 2006;114(1 Suppl): I27–I32.
- 9. Sakagoshi N, Shirakura R, Nakano S, et al. Serial changes in myocardial beta-adrenergic receptor after experimental brain death in dogs. *J Heart Lung Transplant*. 1992;11(6):1054–1058.
- 10. Fukushima N, Sakagoshi N, Ohtake S, et al. Effects of exogenous adrenaline on the number of the beta-adrenergic receptors after brain death in humans. *Transplant Proc.* 2002;34(7):2571–2574.
- 11. Hunt SA, Baldwin J, Baumgartner W, et al. Cardiovascular management of a potential heart donor: a statement from the Transplantation Committee of the American College of Cardiology. *Crit Care Med.* 1996;24:1599–1601.
- 12. Novitzky D, Cooper DKC, Rosendale JD, Kauffman HM. Hormonal therapy of the brain-dead organ donor: experimental and clinical studies. *Transplantation*. 2006;82:1396–1401.
- 13. Stah VR. Aggressive management of multiorgan donor. *Transplant Proc.* 2008;40(4):1087–1090.
- 14. Yoshioka T, Sugimoto H, Uenishi M, et al. Prolonged hemodynamic maintenance by the combined administration of vasopressin and epinephrine in brain death: a clinical study. *Neurosurgery*. 1986;18(5):565–567.



Donor Evaluation and Management System (Medical Consultant System) in Japan: Experience From 200 Consecutive Brain-Dead Organ Donation

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ABSTRACT

Purpose. As the donor shortage is extremely severe in Japan because of a strict Organ Transplantation Act, special strategies must be established to maximize organ transplant opportunities. The purpose of this study was to evaluate our strategies to identify and manage 200 consecutive brain-dead organ donors.

Methods and materials. We retrospectively reviewed the 200 donors procured since the Organ Transplantation Act was issued in 1997, including 118 males, a mean overall age of 45.1 years and cause of death being cerebrovascular disease (n = 119), head trauma (n = 37), and asphyxia (n = 44).

Donor evaluation and management system. Since November in 2002, special transplant management doctors ("medical consultants") were sent to donor hospitals to assess organ function and identify transplantable organs. They also provided intensive care to stabilize hemodynamics and improve cardiac and lung functions by administering antidiuretic hormone intravenously and providing bronchofiberscopic pulmonary toilet.

Results. We obtained 146 heart, 1 heart-lung, and 154 lung (87 single and 67 bilateral), 175 liver (28 splitted liver), 142 pancreas (114 pancreas-kidney), 253 kidney and 12 small bowel grafts. Organs procured from 1 donor increased from 4.5 to 6.8 after applying these strategies.

Conclusions. Although the number of cases was still small, the availability of organs and outcomes of transplantation have been acceptable.

RGAN TRANSPLANTATION (Tx) has achieved satisfying long-term results. However, these surgical therapies are continuously limited by the severe organ donor shortage. Therefore, optimal utilization of all suitable organs is mandatory to increase graft availability. ²

The Japanese Organ Transplantation Act for brain death (BD) organ donation (the former Act) was issued in October 1997. It required written consent for BD and organ donation by a living individual and did not allow it from children younger than 15 years. From these reasons, only 81 BD organ donations were performed in Japan for 13 years. The cardiac donation rate per million populations in Japan is only 0.08, while it is 7.3 in the United States, 5.3 in Spain, and 0.97 in South Korea in 2007. The mean waiting time for heart and lung transplants of 1026 and 1673 days respectively in 2010 was extraordinary long in Japan.

© 2013 by Elsevier Inc. All rights reserved. 360 Park Avenue South, New York, NY 10010-1710 Finally the Act was revised on July 17, 2010. ¹⁻³ New organs can be donated after BD with consent from the family, if the donor did not deny organ donation premortem. Although the Act was revised in 2010 and brain-death organ donation increased from 13 to 44 cases in a year (Fig 1), the number was still extremely smaller than other developed countries. The great pressures of the organ shortage and the long waiting times made Japanese transplant programs consider the use of organs that might be otherwise considered to be marginal.⁴

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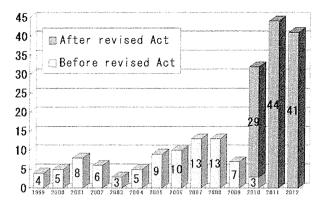


Fig 1. Brain-dead organ donation in Japan.

The most troublesome issue facing transplantation is primary graft dysfunction (PGD). This complication is the leading cause of death in the first 30 days and first year post-transplantation worldwide. The use of marginal donor organs may increase the rate of PGD. From this point of view, it is necessary to establish a special evaluation and management system to maximize cardiac and lung donor utilization. The purpose of this study was to review Japanese special strategies to identify and manage, 200 BD organ donors since issuance of the Japanese Organ Transplantation Law.

MATERIALS AND METHODS

We retrospectively reviewed all 200 brain-dead donors procured in Japan between October 17, 1997, and November 14 2010, including 118 males and 78 females with an overall mean age of 45.1 years. The cause of brain death was cerebral stroke (n = 119: subarachnoid hemorrhage, 7 cerebral infarction, and 21 intracerebral bleeding), head trauma (n = 37), asphyxia (n = 27) and brain injury after cardiopulmonary resuscitation (n = 17).

Medical Consultant System to Evaluate and Manage BD Organ Donors

Since BD organ transplantation was started on February 28, 1999, every organ procurement team has obtained a staff physician in each procurement hospital. Before the procurement operation they evaluate the condition of the donor organs by ultrasound examinations of the heart and abdomen and bronchofiberscope (BFS) candidates in the intensive care unit (ICU).⁴

Since November 2002, special transplant management doctors (medical consultants [MC]) who were usually cardiac transplant surgeons, have been sent to the procurement hospital. They assessed donor organ function to identify organs useful for transplantation. They also intensively care for the donor, stabilize the donor hemodynamics by giving antidiuretic hormone (ADH) as a bolus infusion (0.01 U/kg) followed by a drip infusion (0.01 U/kg/h) and reduce the doses of intravenous catecholamines as much as possible, and seek to improve donor cardiac and lung function by preventing and treating infection before the procurement teams arrive at the donor hospital.

Since the 50th BD donor in December 2006, we have modified lung management. Regular toilet and turning of the donor were done as previously. If there were symptoms and/or signs of atelectasis or pneumonia on chest X-ray and chest computed

tomography (CT) scan, repeated we perform BFS and frequent toileting. Since 2011, lung transplant surgeons have played a role in evaluation and management of the lungs. Currently the MCs consist of about 20 cardiac, 30 lung, and 3 liver transplant surgeons.

Current Organ Donor Evaluation System in Japan

First step: donor evaluation. Transplant procurement coordinators (PTC) of Japan Organ Tx Network (JOT) are called to a donor hospital if there is a potential BD donor. They obtain the patient's clinical course from the medical staff and examine clinical records to determine whether the patient is suitable for organ donation. If there are no absolute contraindications, such as an untreated malignancy or severe viral infection, they obtain informed consent for donation from the family. Then a legal examination for BD is performed.

Second step: donor evaluation. After completion of the initial clinical examination, MCs are sent to the hospital. They and the JOT PTC check the clinical records of the course before and after BD, medication review, blood examination, electrocardiogram, chest X-ray, abdominal and chest CT scans, and perform ultrasound examinations of the heart, liver, pancreas, and kidneys as well as a BFS. MCs rule out malignancies from the CT scan and ultrasound examination. JOT PTCs construct donor evaluation sheets which are sent to transplant centers using a mobile system. Then the transplant centers decided whether their recipient is suitable and their procurement team is sent to the hospital.

Third step: donor evaluation. After arriving at the donor hospital, they also evaluate the condition of donor organs by ultrasound examinations for the heart and abdominal organs and BFS by themselves in the ICU before the procurement operation. They assess organ function to determine whether the organ could be transplanted to their recipient.

Final donor evaluation. After opening the chest and abdomen, the procurement team evaluates organs by watching and touching carefully. Usually a liver biopsy is performed to exclude more than moderate grade fatty liver and malignancies. They also rule out unexpected malignancies in the pleural and abdominal cavities.

Preprocurement Meeting and Management of Procurement Operation

Before starting the procurement operation, surgeons, anesthesiologists and operating room nurses gather in a meeting room. They negotiate the types of procured organs, how to procure each organ (eg, organ dissection/perfusion technique, incision lines, blood drainage technique, etc), the kinds of samples needed (eg, blood, lymphnodes and spleen), and how to manage the donor during the operation.

As most Japanese anesthesiologists have never experienced a procurement operation from a BD donor, MC also support them to stabilize donor hemodynamics during the operation. Skillful staff, not resident surgeons, harvest the donor organ.

As an inverse relationship between volume of intraoperative colloid and early lung allograft function has been reported,⁵ packed red blood cells and albumin was transfused to maintain circulating blood volume and to replace proteins and fluids during the procurement operation. To improve organ perfusion with preservation solution, catecholamines are not additionally administered seeking to dilate the vessels to organs; the ADH and catecholamine infusions are discontinued at the time of the bolus infusion of heparin sulfate (400 U/kg).

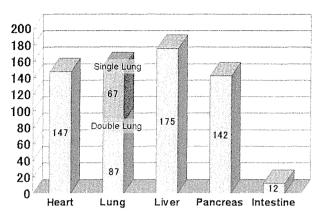


Fig 2. Organs transplanted from 200 brain-dead donors.

RESULTS

We performed 146 heart, 1 heart-lung, and 154 lung (87 single and 67 bilateral), 175 liver (28 split liver), 142 pancreas (114 pancreas-kidney), 253 kidney, and 12 small bowel transplantations (Fig 2). Although donor age has increased after the revision of the Act, the number of organs transplanted per donor and patients transplanted per donor were around 5 and 4 respectively (Fig 3). Patient survivals at 5 and 10 years are shown in Table 1.

DISCUSSION

For many years, "traditional criteria" have been used to identify an appropriate transplant donor. However, over the past 2 decades, there has been a considerable increase in the numbers of patients listed annually for organ transplantation. Strict adherence to standard donor criteria has resulted in a progredient undersupply of available organs

Table 1. Patient Survival After Organ Transplantation

1329

	Heart	Lung	Liver	Pancreas
5-y survival rate (%)	95.2	72.7	78.6	76.0
10-y survival rate (%)	95.2	54.0	70.8	62.6

with significantly extended waiting times and increased waiting list mortality. ^{2,6}

As a consequence of this severe shortage of donor organs, marginal donors have been utilized in many countries. However, only 2407 hearts of 7944 deceased donors (30.2%) were transplanted in the United States in 2010. Because of the strict Japanese Organ Transplantation Law, only 200 BD donors have been procured in Japan for 13 years and 60 heart transplantation (HTx) would have been performed if the cardiac donation rate from the deceased donors was same in Japan as in the United States. These great pressures of the organ shortage had made transplant programs consider the use of organs that would be considered to be marginal. Therefore, an original donor evaluation and management system has been established in Japan, including the MC and the preprocurement meeting.⁴

High serum adrenaline concentrations, as usually observed after its administration, reduce myocardial β-adrenergic receptor density in BD animals⁷ and patients,⁸ which may increase the risk of primary graft dysfunction after HTx. Therefore, the intravenous catecholamine dose should be reduced as much as possible. It has been recommended as initial therapy for hemodynamic support and treatment of diabetes insipidus by the American College of Cardiology,^{9–11} due to its catecholamine-sparing effects.^{6,10} Repletion of vasopressin to treat diabetes insipidus to maintain hemodynamic stability and prevent electrolyte imbalance

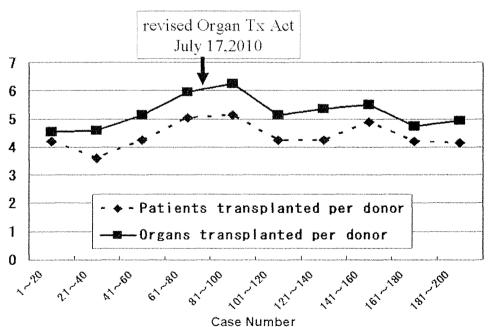


Fig 3. Patients and organs transplanted per donors.

is noncontroversial. A substantial number of BD donors resolve their focal/regional wall motion abnormalities. Aggressive attempts at hemodynamic stabilization using hormonal resuscitation have resulted in dramatic reversibility of cardiac dysfunction. 10,12

The ideal lung donor has a PaO₂/FiO₂ ratio more than 300, positive end expiratory pressure requirement greater than 5 cm H₂O, clear chest X ray, age older than 55 years, smoking history of more than 20 packs/y, and absence of trauma, surgery, aspiration, malignancy, and purulent secretions. Pathologic studies of lungs deemed unsuitable for donation have indicated that bronchopneumonia, diffuse alveolar damage, and lung consolidation are the most common reasons to reject a lung. Given these findings, it is recommended that every lung donor undergo BFS for therapeutic bronchial toilet, and to isolate potential pathogens to guide antibiotic therapy in both the donor and the recipient.¹⁰

In conclusion, although the number of transplantations is still small, the availability of organs has been greater in Japan and their outcomes acceptable. The strategies presented herein may be useful to maximize organ transplant opportunities even in other countries.

ACKNOWLEDGMENTS

We would like to acknowledge and extend our heartfelt gratitude to many heart and lung transplant surgeons who have actually worked as a medical consultant despite of daily hard work and made the completion of this paper possible.

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REFERENCES

1. Christie JD, Edwards LB, Kucheryavaya AY, et al. The Registry of the International Society for Heart and Lung Trans-

plantation: Twenty-seventh official adult lung and heart-lung transplant report—2010. *J Heart Lung Transplant*. 2010;29(10): 1104–1118.

- 2. Wittwer T, Wahlers T. Marginal donor grafts in heart transplantation: lessons learned from 25 years of experience. *Transpl Int.* 2008;21(2):113–125.
- 3. Matsuda H, Fukushima N, Sawa Y, et al. First brain dead donor heart transplantation under new legislation in Japan. *Jpn J Thorac Cardiovasc Surg.* 1999;47(10):499–505.
- 4. Fukushima N, Ono M, Nakatani T, et al. Strategies for maximizing heart and lung transplantation opportunities in Japan. *Transplant Proc.* 2009;41(1):273–276.
- 5. McIlroyl DR, Pilcher DV, Snell GI. Does anaesthetic management affect early outcomes after lung transplant? An exploratory analysis. *Br J Anaesth*. 2009;102(4):506–514.
- 6. Lima B, Rajagopal K, Petersen RP, et al. Marginal cardiac allografts do not have increased primary graft dysfunction in alternate list transplantation. *Circulation*. 2006;114(1 Suppl):I27–I32.
- 7. Sakagoshi N, Shirakura R, Nakano S, et al. Serial changes in myocardial beta-adrenergic receptor after experimental brain death in dogs. *J Heart Lung Transplant*. 1992;11(6):1054–1058.
- 8. Fukushima N, Sakagoshi N, Ohtake S, et al. Effects of exogenous adrenaline on the number of the beta-adrenergic receptors after brain death in humans. *Transplant Proc.* 2002;34(7):2571–2574.
- 9. Hunt SA, Baldwin J, Baumgartner W, et al. Cardiovascular management of a potential heart donor: a statement from the Transplantation Committee of the American College of Cardiology. *Crit Care Med.* 1996;24:1599.
- 10. Novitzky D, Cooper DKC,2 Rosendale JD, Kauffman HM. Hormonal therapy of the brain-dead organ donor: experimental and clinical studies. *Transplantation*. 2006;82:1396–1401.
- 11. Stah VR. Aggressive management of multiorgan donor. *Transplant Proc.* 2008;40(4):1087–1090.
- 12. Yoshioka T, Sugimoto H, Uenishi M, et al. Prolonged hemodynamic maintenance by the combined administration of vasopressin and epinephrine in brain death: a clinical study. *Neurosurgery*. 1986;18(5):565–567.



Current Status of In-Hospital Donation Coordinators in Japan: Nationwide Survey

S. Konaka, S. Shimizu, M. Iizawa, H. Ohkawara, O. Kato, J. Ashikari, and N. Fukushima

ABSTRACT

Objectives. When the Japanese Organ Transplantation Act was issued, the Japanese Organ Transplantation Network (JOT) was established in 1997. JOT lists recipients, assesses and manages organ donors, and educates publics and headquarters for organ donations. JOT procurement transplant coordinators (PTC) play roles in obtaining consent from relatives for organ donation, donor evaluation and management, organ recovery management, organ transport, and care of donor families during and after donation. Every prefecture has at least one PTC who is mainly working in public education and hospital development. They also help the JOT PTC at the time of organ procurement. Most prefectures commission hospital staff in the procurement hospital to be an in-hospital PTC (In-Hp PTC), who make their hospital staff aware of organ donation and support organ procurement. Although the Act was revised in 2010 with brain-dead organ donation increased from 13 to 44 cases yearly, the number was still extremely smaller than other developed countries. In these circumstances, In-Hp PTC may play greater roles to increase donation and smooth procurement procedures Our primary aim was to describe the current status of In-Hp PTC in Japan.

Materials and methods. Between December 15, 2011, and January 31, 2012, we invited 1889 In-Hp PTC to complete a letter survey using a self-designed questionnaire. In all, 56 In-Hp PTC (40%) completed and returned it.

Results. The occupation of the respondents was nurse (66%), physician (18%), or other (16%). Although 52% of respondents belonged to the hospital, which was designated for brain-death organ donation by the government, only 46% had any experience with a cadaveric donor. Only 2% were full-time In-Hp PTC. They mainly played a role in preparing their own manual for organ procurement (57%), providing in-hospital lectures (44%) or their own simulation exercise (29%), as well as coordinating donation cases. Although 77% had attended seminar about organ donation provided by JOT or the prefecture PTC, 93% wanted more professional education. However, it was difficult for them to attend these activities, to manage a rare and sudden donation case, and to find time to learn about organ donation because they had another post. The topics that they wanted to learn were donor family care (72%), overall organ/tissue donation procedures (65%), the role of In-Hp PTC (67%), simulations of donation (65%), legislation and social system of organ donation (61%), medical indications for donation (61%), current status of donation and transplantation in Japan (57%), donor management (56%), and case studies (49%). There were significant variations in the topics of interest among the occupations.

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As they had another post, they could find only a short period (1 or 2 days) to take professional education, such as lectures. Therefore, it was difficult for them to attend practical on-the-job training.

Conclusions. To establish an organ procurement system and increase organ donation, In-Hp PTC have important roles in Japan. However, none is a full-time In-Hp PTC. Most In-Hp PTC require more professional education. A systematic education program for each occupation must be established soon.

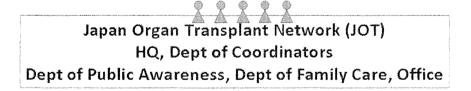
SINCE 1978, the donation of kidneys after cardiac death (DCD) has been legally accepted in Japan after family consent. Small children had been able to donate their kidneys after cardiac death. The Japanese Organ Transplantation Act for brain-death (BD) donation was issued in October 1997. The Act required a living written consent for BD declaration and organ donation; it did not allow BD donation from children younger than 15 years. For these reasons, only 81 BD organ donations were performed in Japan over 13 years since the Act was issued in October 1997.

The Japanese Organ Transplant Act issued in 1997, established the Japanese Organ Transplantation Network (JOT), which lists recipients, assesses and manages organ donors, and educates publics and headquarters of organ donations. JOT procurement transplant coordinators (PTC) play roles in obtaining consent for organ donation from relatives, donor evaluation and management, management of organ recovery, organ transport, and care of donor families during and after donation.

Every prefecture has at least one of their own 1 PTC who is mainly working on public education and hospital development. They also collaborate with the JOT PTC at the time of organ procurement. Most prefectures commission staffs in procurement hospitals to be in-hospital PTC (In-Hp PTC), who make their hospital staffs aware of organ donation and support organ procurement (Fig 1).^{1,2}

Finally the Act was revised on July 17 2010. 1-3 Renewal of the Act allowed organs to be donated after BD with family consent if not previously denied before the event. Although the Act was revised in 2010 and BD organ donation increased from 13 to 44 cases in a year, the number was still extremely smaller than that in other developed countries. The revised Act accepts organ donation from BD children younger than 15 years. However, only 158/504 (42.4%) procurement hospitals where BD organ donation is allowed by the Government have established procurement systems from children. In these circumstances, In-Hp PTC may play a great role to increase organ donation and smooth procurement procedures.

The Department of Coordinators and the JOT coordinator committee play the main roles in educating these PTC. JOT has prepared guideline manuals of standard roles and procedures of PTC during organ procurement from BD and DCD donors. Although the JOT has prepared a textbook for In-Hp PTC and held several educational programs for In-Hp PTC, they have been educated mainly by prefectural PTC or their own hospital. Therefore, educational systems for In-Hp PTC should be modified to establish an effective tool once the current status and needs of In-Hp PTC are clarified. Therefore, the primary aim of this study was to describe the current status of In-Hp PTC in Japan based upon a national survey.



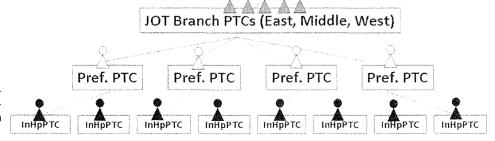


Fig 1. Organ transplant network in Japan. PTC, procurement transplant coordinator; In HpPTC, in-hospital PTC.

MATERIALS AND METHODS

We developed a 28-item self-completed questionnaire that queried: (1) occupation and status; (2) activities and issues (daily and at organ donation); (3) education and experiences of organ donation; (4) needs for learning about organ donation.

Survey letters were sent to 389 donor hospitals and 1889 In-Hp PTC. In 40 prefectures where the In-Hp PTC was delegated by the prefectural government, survey letters were directly or indirectly sent to the delegated In-Hp PTC. In 4 prefectures (Tokyo, Chiba, Saitama, and Osaka) where In-Hp PTC are not delegated, survey letters were sent to the BD donor hospitals. In all, 756 In-Hp PTC (40%) completed and returned the survey. The survey period was December 15, 2011, to March 31, 2012.

RESULTS

Among 1679 letters sent to In-Hp PTC in 40 prefectures where an In-Hp PTC was delegated by the prefectural government, there were 739 (44%) In-Hp PTC responses. But only 17 In-Hp PTC (8%) responded among 210 letters sent to In-Hp PTC in 4 prefectures that had not delegated the In-Hp PTC.

Hospital Where In-Hp PTC Were Working

Overall, 52% of respondents worked in donor hospitals accepted to undergo BD organ donation; 38% in hospitals accepted only to undergo donation after cardiac death (Fig 2a). Only 46% worked in donor hospitals where cadaveric organ donations had been performed in the past (Fig 2b). However, 63% worked in a donor hospital where committees were established for cadaveric organ donation (Fig 2c).

Occupation and Status of In-Hp PTC

The occupations of the respondents were nurse in (66%), physician (18%) or other $(16\%; \mathrm{Fig}\ 3)$. Although only 2% of individuals was a full-time In-Hp PTC; 83% had been delegated by the prefectural government and 49% were supported from an advisory committees in their hospital (Fig 4).

Activities and Issued of In-Hp PTC

Their main roles was to prepare a manual for organ procurement (57%), to provide awareness of organ dona-

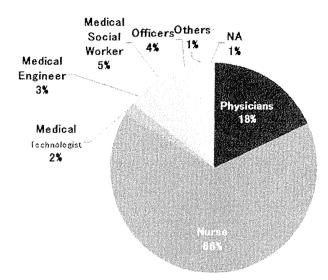
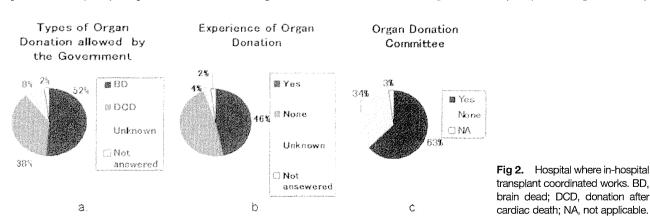


Fig 3. Occupation of in-hospital procurement transplant coordinator.

tion for patients and medical staff (56%), to arrange in-hospital seminars (44%), to consult about organ transplantation and donation (24%), to organize simulations of organ donation (29%), as well as to coordinate donation cases (Fig 5).

Among the 345 respondents who had experienced a cadaveric donor procedure, 77% had coordinated the inhospital staff and arranged organ procurement; 59% communicated with prefectural and JOT PTC; 58% had cared for the donor family and 35% had obtained informed consent for donation accompanied by a prefectural or JOT PTC (Fig 6).

However, it was difficult for them to do these activities, namely, manage a rare, sudden donation case and to learn organ donation, because they had another post with regard to daily issues, they answered "hard to work as In-Hp PTC due to part time activity" (42%) "no daily time to work as In-Hp PTC" (31%), and "no support by hospital administration and other medical staffs" (29%; Fig 7). With regard to issues at organ donation, they answered no knowledge of their activities at organ donation (77%), "cannot preferentially



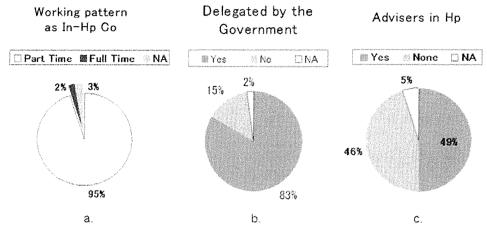


Fig 4. Status of in-hospital procurement transplant coordinator. In-Hp Co, in-hospital coordinator; Hp, hospital; NA, not available.

involve in donation" (66%), "annoying their partners because they cannot do their daily jobs" (65%), no knowledge of donor family care (49%), and no knowledge of organ donation process (36%; Fig 8).

Educational Experiences of Organ Donation

Although 77% have attended seminars about organ donation provided by JOT or the prefecture PTC (Fig 5), 93% still wanted to obtain more professional education. Nurses and medical social workers were more likely to desire more professional education than physicians or medical examiners.

The topics that they desired to learn were donor family care (72%), overall procedures of organ/tissue donation (65%), role of In-Hp PTC (67%), simulations of organ donation (65%), legislation and social system of organ donation (61%), medical indications for donation (61%), current status in Japan (57%), donor management (56%), and case studies (49%; Fig 9). There were significant variations in interested topics among the occupations.

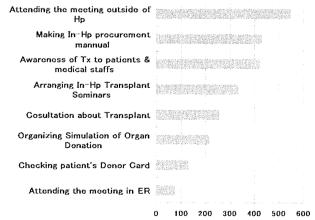


Fig 5. Daily activities as in-hospital procurement transplant coordinator; Hp, hospital; Tx, transplant; ER, emergency room.

DISCUSSION

Although the Transplantation Act was revised in 2010 and BD organ donation increased from 13 to 44 cases in a year, the number is still extremely smaller than that in other developed countries. In these circumstances, In-Hp PTC may play great roles to increase organ donation and smoothing procurement procedures.

Currently, JOT PTC and prefectural PTC conduct class-room lectures for In-Hp PTC and other medical staff in each donor hospital. The topics include organ transplantation/donation legislation in Japan, current status of DCD and BD donation in Japan, the roles of medical staffs and the donation processes: namely, initial actions, family consent, donor evaluation. Indeed, although 77% of respondents attended these seminars, 93% still wanted more professional education. Nurses and medical social workers were more likely than physicians and medical examiners to desire more professional education.

As most of In-Hp PTC had another post, they could find only a short period (1 or 2 days) to take professional education such as lectures. Even at organ donation, they could not be preferentially involved in organ donation

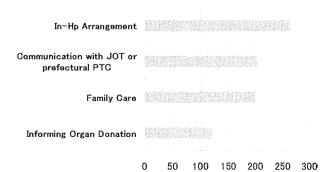


Fig 6. Activities at organ procurement. In-Hp, in-hospital; JOT, PTC, procurement transplant coordinator.

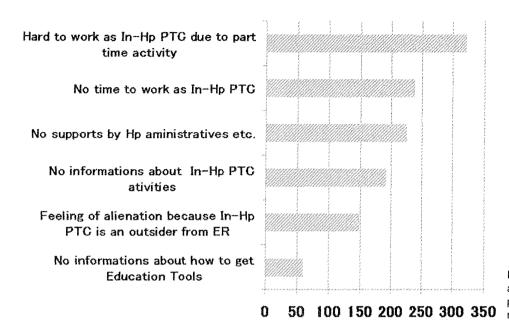


Fig 7. Issues of daily activities as In-Hp PTC. Hp, hospital; PTC, procurement transplant coordinator; ER, emergency room.

because they felt sorry for their partners if they did not do their daily jobs. Therefore, it was difficult for them to find time to obtain practical on-the-job training.

Of course, it may be effective to establish full time In-Hp PTC in every donor hospital. However, it is hard to establish such a post in every hospital, because the number of organ donations has been extremely small in Japan. Therefore, we need to establish special educational programs for In-Hp PTC.

The results of the present study led our department to hold a special educational seminar and establish a program for In-Hp PTC in Japan: namely, 2 lectures per day for 10

days from May to October 2012. Topics of this seminar were the history of transplantation, current status of organ transplantation and donation in Japan, legislation and network system of organ transplantation in Japan, outcomes of organ transplantation in Japan, nursing care of transplant recipients, pathophysiology and diagnosis of BD, the role of PTC, process of organ donation, the visit of JOT office, family care and informed consent for organ donation, survey of donor family, donor evaluation and management, role of In-Hp PTCs, case studies and simulations of organ donation, pediatric organ donation, determination of child abuse, informing organ donation, and care of the family of

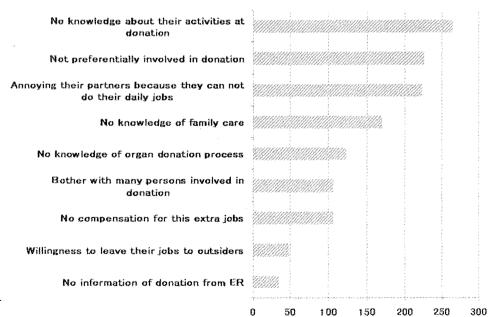
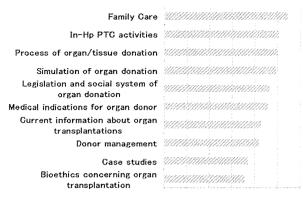


Fig 8. Issues at organ donation. ER, emergency room.



0 100 200 300 400 500 600

Fig 9. Top-10 subjects desired to learn. Hp, hospital; PTC, procurement transplant coordinator.

an end-life patient. Survey of the 16 participants showed all of them to be satisfied with the program, but they still wanted to have on-the-job training.

Although we may be able to establish an education curriculum for In-Hp PTC in the near future, national support is essential to provide these programs to all In-Hp PTC in Japan.

In conclusion, to establish the organ procurement system and increase donation, In-Hp PTCs have great roles in Japan. However, few of them have full-time positions and most of them require more professional education. Each In-Hp PTC has several problems, seeking to study the organ donation process and their role in BD the donations under the revised Transplant Act. We need to make a special national educational program for in-Hp PTCs.

REFERENCES

- 1. Fukushima N, Konaka S, Kato O, Ashikari J. Professional education and hospital development for organ donation. *Transplant Proc.* 2012;44(4);848–850.
- 2. Konaka S, Kato O, Ashikari J, Fukushima F. Modification of education system for organ procurement coordinators in Japan after the revision of the Japanese Organ Transplantation Act. *Transplant Proc.* 2012;44(4);851–854.
- 3. Fukushima N. Revised Organ Transplant Act and Transplant Surgeons. *Jpn Med Assoc J.* 2011:54(6);387–391.