

Efficient and Incentive-Compatible Liver Exchange

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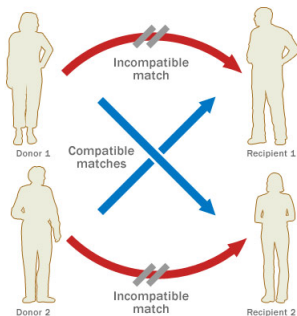
Boston College

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- **Kidney Exchange** became a wide-spread modality of transplantation within the last decade.
- About 600 patients a year receive kidney transplant in the US along through exchange, about 12% of all live-donor transplants.
- In theory **live donor organ exchange** can be utilized for any organ for which live donation is feasible.
- **Liver** is the second most transplanted organ after kidneys; moreover, live-donor lobar liver donation is feasible.



- Human organs cannot be received or given in exchange for "valuable consideration" (US, NOTA 1984, WHO)
- However, **live donor kidney exchange** is not considered as "valuable consideration" (US NOTA amendment, 2007)



- **Kidney Exchange Literature:** Plenty...



- **Kidney Exchange Literature:** Plenty...
- **Liver Exchange Literature:**
 - Hwang et al. [10] proposed the idea and documented the practice in Korea since 03
 - Chen et al. [10] documented the program in Hong Kong
 - Dickerson & Sandholm [14] asymptotic gains from liver+kidney exchange over isolated liver exchange and kidney exchange
 - Ergin, Sönmez, & Ünver [17] proposed and modeled exchange for transplants each of that needs two live donors: lung, simultaneous liver+kidney, dual-graft liver



- We model liver exchange as a matching problem – different from kidney exchange due to **size-compatibility** requirement.
- We find the structure of feasible two-way exchanges and a sequential algorithm to find an efficient matching for two patient/donor sizes.
- The requirement of **size compatibility** induces an incentive problem for the pair/donor to donate
 - the larger/riskier/easier to match **right lobe** or
 - the smaller/safer/more difficult to match **left lobe**
- For a continuum of patient/donor sizes, we propose a **Pareto-efficient** and **incentive-compatible** mechanism that elicits willingness to donate **right lobe** truthfully.
 - A new class of bilateral exchange mechanisms for **vector-partial-order-induced** weak preferences.



- Live donor liver donation is more common in Asian countries where deceased donation is at a minimum due to cultural reasons and legal non-recognition of "brain death".

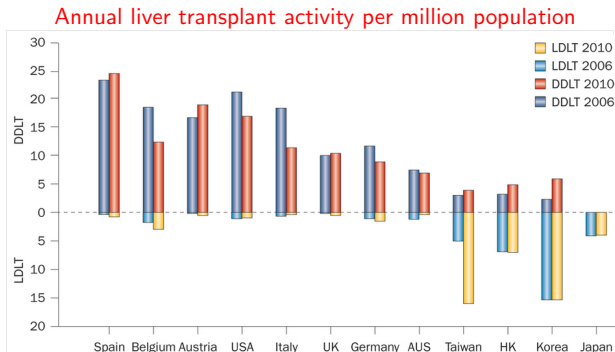
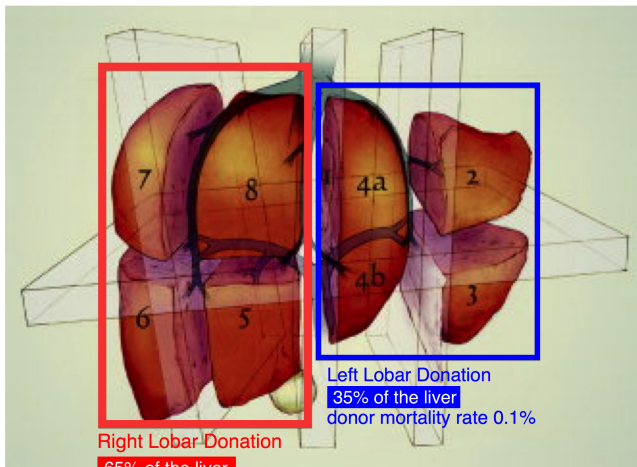
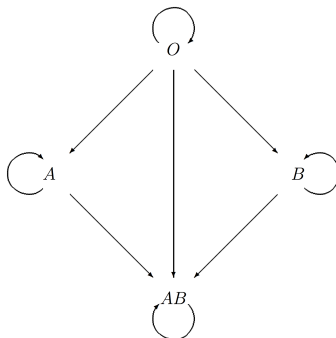


Figure from Chen et al Nature Reviews Gastroenterology & Hepatology 2013





- Blood-type compatibility is required.



- Size compatibility is required unlike kidneys: A patient requires a graft relatively large to survive.
- Tissue-type compatibility is **not** required unlike kidneys.



- Right-lobe transplant has been utilized for size compatibility despite its heightened donor mortality risk.
 - Patient needs roughly at least 40% of his own liver size to survive.
 - Donor needs at least 30% remnant liver volume to survive.
 - Usually right lobe is $\sim 65\%$, left lobe is $\sim 35\%$ of liver.
 - In many occasions, size compatibility is only satisfied through right-lobe donation.



11	2003	Japan	A mother in her late 40s donated a right lobe and died 9 months later from complications of hepatic failure.
12	2002	USA	A 57-year-old brother donated a right lobe and developed gastric gas gangrene and <i>Clostridium perfringens</i> infection 3 days after surgery and died.
13	2005	Brazil	A 31-year-old female right lobe donor of unknown relationship to the recipient died 7 days after surgery from a subarachnoid hemorrhage.
14	2003	India	A donor of unknown age and unknown relationship to the recipient donated an unknown lobe and died 10 days after surgery of unknown causes.
15	2003	India	A 52-year-old wife donated an unknown lobe and became comatose 48 hours after surgery from unknown causes and remains in chronic vegetative state.
16-18	1993	Germany	A 29-year-old mother donated a left lateral lobe and died of a pulmonary embolus 48 hours after surgery.
18, 19	2000	Germany	A 38-year-old father donated a right lobe , and 32 days after developing progressive hepatic failure, died during transplantation of acute cardiac failure. The cause of the donor's death was attributed to Bernardinelli-Seip syndrome, a lipodystrophy syndrome characterized by loss of body fat, diabetes, hepatomegaly, and acanthosis nigricans.
18, 20	2000	France	A 32-year-old brother donated a right lobe and developed sepsis and multiple organ system failure 11 days after surgery and died of septic shock 3 days later.
18	2000	Europe	A 57-year-old wife donated a right lobe and died of sepsis and multiple organ system failure 21 days after surgery.
21, 22	1999	USA	A 41-year-old half-brother donated a right lobe and died of pancreatitis and sepsis 1 month later.
22, 23	1997	USA	A mother of unknown age donated an unknown lobe to a pediatric recipient and died 3 days after surgery of unknown causes.
24	2005	Asia	A 50-year-old mother donated a right hepatic lobe . She had no history of peptic ulcer disease and received a 2-week course of H2 antagonist. She died 10 weeks after surgery from an autopsy-proven duodenal ulcer with a duodenocaval fistula causing air embolism.
25	2006	Asia	A 39-year-old male "close relative" who donated an unknown lobe died of a myocardial infarction 4 days after donation. The patient reportedly had a preoperative electrocardiogram and treadmill test.
26	2005	Egypt	A brother of unknown age who donated a right lobe died of complications of sepsis from a bile leak 1 month after donation.
Donor deaths "possibly" related to donor hepatectomy			
27	2005	USA	A 35-year-old brother donated a right lobe and died of a self-induced drug overdose 23 months later.
27	2005	USA	A 50-year-old uncle donated a right lobe and died of a self-inflicted gunshot wound to the head 22 months after donation.

- **Right-lobe** donation is reported to be 5 times riskier in mortality rates than **left-lobe** donation (0.5% to 0.1%).
- A high profile death of a live **right-lobe** donor in the US in 2002 decreased live donation steadily not only for livers, but for other organs including kidneys in the US.
- Trotter et al. [06]: Documented living liver donor deaths due to donation.
- About half of live donations **right lobe**.



- **Liver exchange** first done in Korea, followed by Hong Kong and Turkey.
- Liver exchange can have two benefits:
 - It can **increase** the number of transplants.
 - It can **decrease** the share of **right-lobe transplants** (and increase donor safety) through matching with respect to size.

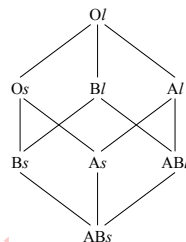


- Liver exchange differs from kidney exchange in two key ways:
 - The lack of **tissue-type incompatibility**, and
 - the presence of **size incompatibility**.
- In the absence of **size incompatibility** the scope for liver exchange would be very limited: The only viable exchange would be between
 - a blood type A patient with blood type B donor and
 - a blood type B patient with blood type A donor.



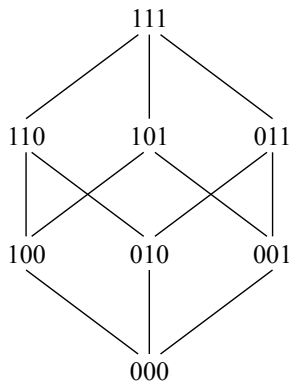
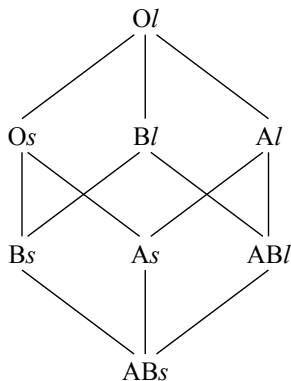
- $\underbrace{\{O, A, B, AB\}}_{\mathcal{B}} \times \underbrace{\{I, s\}}_{\mathcal{S}}$: Set of individual types
- Initially, we focus on live-donor **left-lobe** liver transplants.
- **Left-Lobe Compatibility**: A donor can donate to a patient if and only if
 - (1) the patient is **blood type compatible** with the donor, and
 - (2) the donor is **not smaller** than the patient.

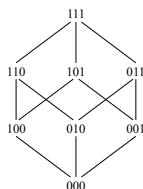
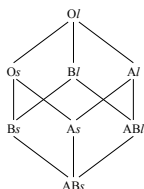
Liver Donation Partial Order \trianglelefteq on $\mathcal{B} \times \mathcal{S}$





- Consider the following two partially ordered sets:
 - The **liver donation partial order** \succeq on $\mathcal{B} \times \mathcal{S}$, and
 - the **standard partial order** \geq over the corners of the three-dimensional cube $\{0, 1\}^3$.





- Note that $(\mathcal{B} \times \mathcal{S}, \supseteq)$ and $(\{0, 1\}^3, \geq)$ are order isomorphic, where the order isomorphism associates each individual type $\tau \in \mathcal{B} \times \mathcal{S}$ with the following vector $X \in \{0, 1\}^3$:

$$X_1 = 0 \iff \tau \text{ has the } A \text{ antigen}$$

$$X_2 = 0 \iff \tau \text{ has the } B \text{ antigen}$$

$$X_3 = 0 \iff \tau \text{ is small}$$

- For notational transparency, we will work with the equivalent representation $(\{0, 1\}^3, \geq)$.



- The **type** of a patient-donor **pair** is represented through the individual types of its patient and donor, respectively, as $X - Y \in (\{0, 1\}^3)^2$.

Definition

A **liver exchange problem** is a list $\mathcal{E} = \{\mathcal{I}, \tau\}$ where $\mathcal{I} = \{1, 2, \dots, I\}$ is a set of pairs, and for each $i \in \mathcal{I}$, $\tau(i) = X - Y$ is the type of pair i .



- A (left-lobe) direct transplant consists of a single pair i of type $X - Y$ such that $Y \geq X$. Such pairs are called (left-lobe) compatible pairs.
We represent it as $\{i\}$.
- A (left-lobe-only two-way) liver exchange consists of two pairs i of type $V - W$ and j of type $X - Y$ such that $Y \geq V$ and $W \geq X$.
We represent it as $\{i, j\}$.
- There is embedded endowment bias.
- A matching is a collection of mutually exclusive exchanges and direct transplants such that if a pair is compatible, then it participates in a direct transplant.
(individual rationality in embedded for now in definition).



Observation

In any liver exchange problem, the only types that could be part of a two-way exchange are

$$X - Y \in (\{0, 1\}^3)^2 \text{ such that } X \not\preceq Y \text{ and } Y \not\preceq X.$$



- **Value** of a pair type $\underbrace{X_1 X_2 X_3}_X - \underbrace{Y_1 Y_2 Y_3}_Y$ is defined as

$$v(X - Y) = \sum_{k=1}^3 Y_k - \sum_{k=1}^3 X_k.$$

- **Waste** of an exchange between pair types $V - W$ and $X - Y$ is defined as $v(V - W) + v(X - Y)$.
- All feasible exchanges have non-negative waste.
- **Observation:** All feasible exchanges are either Waste 0, 1, or 2.



- A left-lobe-only two-way matching is (Pareto) efficient if and only if it maximizes the number of transplants.

(follows from *matching matroid* - e.g. see Lawler [1976]).

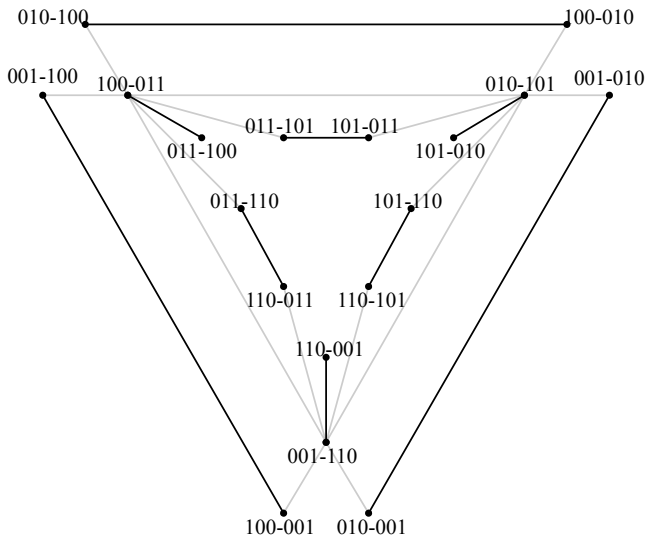
- We propose a greedy (or sequential matching) algorithm to achieve an efficient matching.

In general graphs, efficient two-way matchings cannot be found greedily (e.g. see Edmonds [1965])

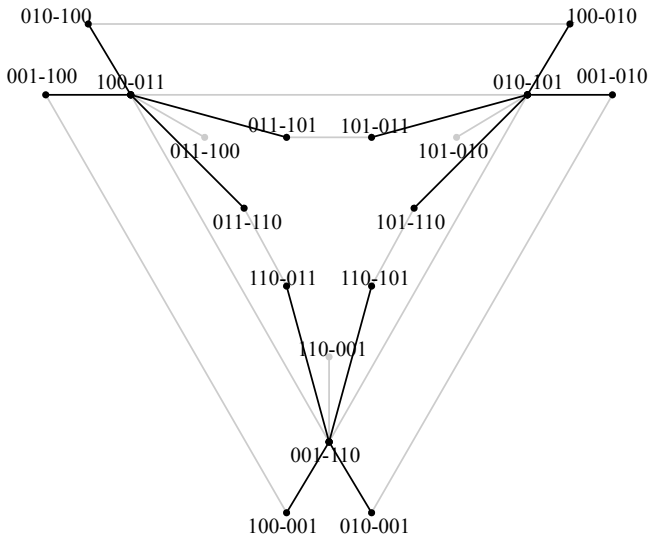


- Step 0. Clear all feasible direct transplants.
- Step 1. Clear **Waste 0** Exchanges: Match the maximum number of $X - Y$ and $Y - X$ types for all $X, Y \in \{0, 1\}^3$.
- Step 2. Clear **Waste 1** Exchanges: Match the maximum number of 100 – 011, 010 – 101, and 001 – 110 types, *without matching them to each other*.
- Step 3. Clear **Waste 2** Exchanges: Match the maximum number of 100 – 011, 010 – 101, and 001 – 110 types *among each other*.

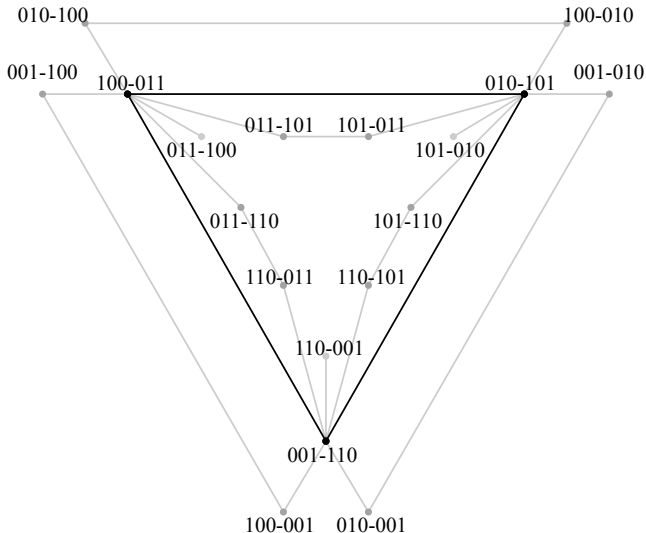
Algorithm Step 1: Waste 0 Exchanges



Algorithm Step 2: Waste 1 Exchanges



Algorithm Step 3: Waste 2 Exchanges





Theorem

Given a liver exchange problem, the *two-size left-lobe-only sequential exchange algorithm* maximizes the number of *left-lobe-only* two-way exchanges.



- Donation Possibilities:
 - Left-lobe donation: Less risky for the donor.
Blood-type compatible donor should be at least as large as the patient.
 - Right-lobe donation: More risky for the donor.
Blood-type compatible donor can donate to a larger patient.
- Pair Preferences:
 - Over donation
 - Donating left lobe is always preferable to donating right lobe or not donating at all
 - The pair may prefer donating right lobe to not donating at all: Type willing (w).
 - The pair may prefer not donating at all to donating right lobe: Type unwilling (u).
 - Over reception fixing the lobe donated:
A pair prefers direct transplant to exchange.



Willing preferences R_i^w :



Willing preferences R_i^w :

Left Direct

Left Exchange



Willing preferences R_i^w :

Left Direct

Left Exchange

Right Direct

Right Exchange



Willing preferences R_i^w :

Left Direct

Left Exchange

Right Direct

Right Exchange

\emptyset

\vdots



Willing preferences R_i^w :

Left Direct

Left Exchange

Right Direct

Right Exchange

\emptyset

\vdots

Unwilling preferences R_i^u :



Willing preferences R_i^w :

Left Direct

Left Exchange

Right Direct

Right Exchange

\emptyset

\vdots

Unwilling preferences R_i^u :

Left Direct

Left Exchange



Willing preferences R_i^w :

Left Direct

Left Exchange

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\emptyset

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Left Direct

Left Exchange

\emptyset

\vdots



- We focus only on **individual rational exchanges**:
 - A **left-lobe** compatible pair does not join in any exchange, but only in **direct transplant**.
 - A **right-lobe-only** compatible pair participates in an exchange only if its donor donates her **left lobe**; otherwise, it participates in **direct transplant**.



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 - A **left-lobe** compatible pair does not join in any exchange, but only in **direct transplant**.
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- **Willingness type of a pair is private information**.
- We inspect direct revelation **mechanisms** to elicit willingness types.



- Fix a willingness profile $R = (R_i)_{i \in \mathcal{I}}$ where all $R_i \in \{R_i^u, R_i^w\}$
- Type $X_1 X_2 X_3 - Y_1 Y_2 0$ w is *treated like* $X_1 X_2 X_3 - Y_1 Y_2 1$ when it donates **right lobe**. We refer to this as **transformation**.
- Hence, when type $X_1 X_2 X_3 - Y_1 Y_2 1$ covers both **native** $X_1 X_2 X_3 - Y_1 Y_2 1$ type pairs and **transformed** $X_1 X_2 X_3 - Y_1 Y_2 0$ w type pairs, we refer to it as an **auxiliary** type.



Lemma

Under left&right-lobe donation, pair types that can benefit from *right-lobe* donation are $X - Yw$ such that

- I $X \not\geq Y_1 Y_2 1$ & $X > Y$, which can only participate in exchange by donating *right lobe* ($100 - 000w$, $010 - 000w$, $110 - 000w$, $110 - 100w$, $110 - 010w$);
- II $X \not\leq Y_1 Y_2 1$ & $X \not\geq Y$, which can participate in exchange by donating *left lobe* or donating *right lobe* ($010 - 100w$, $100 - 010w$, $011 - 100w$, $101 - 010w$);
- III $X = Y_1 Y_2 1$, which can donate *right lobe* to themselves and they do not participate in exchange ($001 - 000w$, $101 - 100w$, $011 - 010w$, $111 - 110w$);
- IV $X < Y_1 Y_2 1$, which can only participate in exchange by donating *left lobe* and otherwise donate *right lobe* to themselves ($001 - 100w$, $001 - 010w$, $001 - 110w$, $101 - 110w$, $011 - 110w$).



Lemma (Observation about Efficiency and Incentive Compatibility)

Under right lobe donation, the pair types that can benefit from *right-lobe* donation are $X - Yw$ such that

- I $X \not\geq Y_1 Y_2 1$ & $X > Y$, which can only participate in exchange by donating *right lobe*
→ *Transform at the beginning*
- II $X \not\leq Y_1 Y_2 1$ & $X \not\geq Y$, which can participate in exchange by donating *left lobe* or donating *right lobe*
→ *Gradually transform as left-lobe exchanges are exhausted*
- III $X = Y_1 Y_2 1$, which can donate *right lobe* to themselves and they do not participate in exchange
→ *Direct transplant at the beginning*
- IV $X < Y_1 Y_2 1$, which can only participate in exchange by donating *left lobe* and otherwise donate *right lobe* to themselves
→ *Direct transplant at the end*



- A **mechanism** is a systematic procedure that finds a matching for each willingness type profile reported.
- A mechanism is **incentive compatible** if it is a weakly dominant strategy for each pair to reveal its willingness type truthfully.
- Any sequential algorithm using the above order of transformation for different category willing pairs and using an external fixed priority order in choosing among the same type of pairs will be incentive compatible as a mechanism.



- Sustaining Pareto efficiency is trickier than incentive compatibility.
- Pareto efficiency no longer implies transplant maximality.
- **Intuition from left-lobe-only exchanges:** Clear **waste 0**, **waste 1**, and then **waste 2** exchanges, in this order, for efficiency.
- Use the same intuition, but **transform** willing pairs with small donors as soon as their left-lobe donation possibilities are exhausted, so incentives and efficiency are aligned.
- Restart clearing from waste 0 exchanges.
Small caveat as the 2nd and 3rd/4th points can have a tradeoff.

Two-Size *Left&Right-Lobe* Sequential Exchange Mech



Fix a priority order over pairs for priority matching in each step.

Step 0. Direct transplant each left-lobe compatible and **Category III** w pair.

Step 1. *Transform* **Category I** w pairs.

Clear **Waste 0** exchanges.

Step 2. Either $010 - 100$ or $100 - 010$ (or both) type pairs are depleted.

Assume wlog $100 - 010$ types are depleted.

(Otherwise proceed symmetrically.)

Clear **Waste 1** exchanges of $100 - 010$.

Step 3. *Transform* $010 - 100$ w pairs.

Clear **Waste 0** exchanges.

Step 4. *Transform* $011 - 100$ w and $101 - 010$ w types.

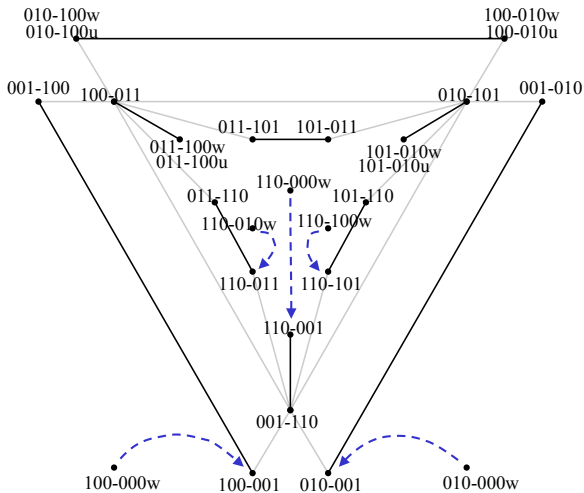
Clear **Waste 0** exchanges.

Step 5. Clear **Waste 1** exchanges.

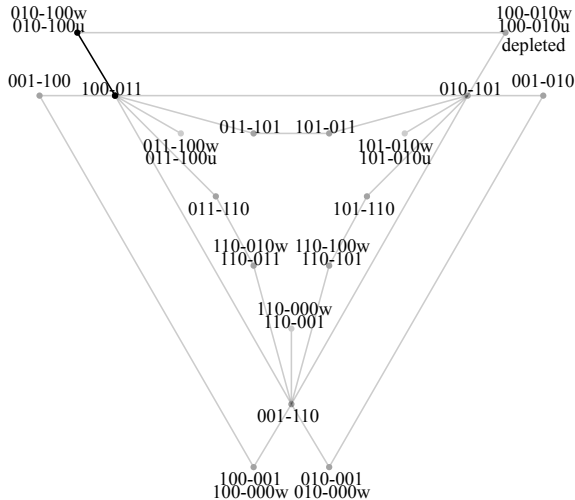
Step 6. Clear **Waste 2** exchanges.

Step 7. Direct transplant each remaining **Category IV** w pair.

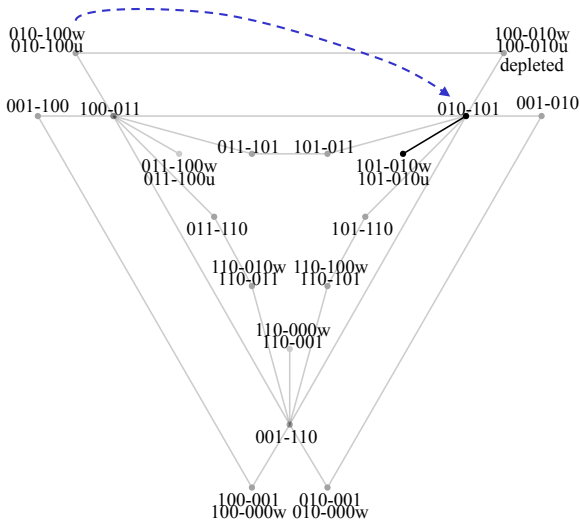
Algorithm Step 1:



Algorithm Step 2:

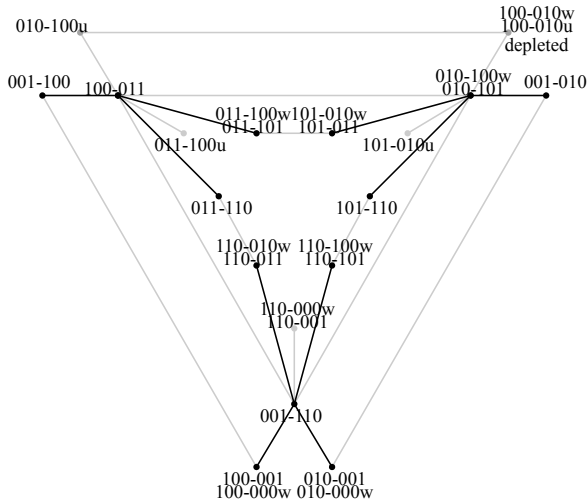


Algorithm Step 3:





Algorithm Step 5:







Theorem

The *two-size left&right-lobe sequential exchange mechanism* is individually rational, Pareto-efficient, and incentive compatible for two sizes of individuals.



Suppose $\mathcal{S} = \{0, 1, \dots, S-1\}$ is the set of possible individual sizes (instead of just $\{0, 1\}$) given a finite exchange pool $\mathcal{E} = (\mathcal{I}, \tau)$.

An individual type: $X = X_1 X_2 X_3 \in \{0, 1\} \times \{0, 1\} \times \mathcal{S}$

Pair type: $X - Y \in (\{0, 1\}^2 \times \mathcal{S})^2$

Right-lobe donation function $\rho : \mathcal{S} \rightarrow \mathcal{S}$ such that it is non-decreasing, and $\rho(s) > s$ for all $s \in \mathcal{S} \setminus \{S-1\}$:

A donor of size s size can donate **right lobe** to any patient of size $s' \leq \rho(s)$.

Let for type $Y = Y_1 Y_2 Y_3$, let $\rho(Y) := Y_1 Y_2 \rho(Y_3)$.



- 1 Greedily committing to an exchange may be no longer possible even for just **left-lobe-only** exchange.
- 2 When **right-lobe** donation is also possible, the *transformation* order of **Category II** willing pairs is non-trivial.

► Skip Diff. 1

Difficulties: 1. Left-Lobe-Only Non-Greedy Matching



A subset of pairs is **matchable** if there exists a matching that match each of them.

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Recursive **augmentation** on a graph can extend the matchable set and find an efficient matching using a **non-greedy** approach in polynomial time (Augment/Shrink-a-Blossom Algorithm of Edmonds, 1965).

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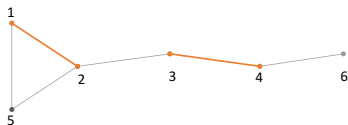
Matchable: $\{1, 2, 3, 4\}$

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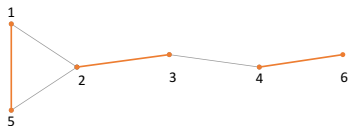


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Matchable: $\{1, 2, 3, 4\}$



Matchable: $\{1, 2, 3, 4, 5\}$

Difficulties: 2. Transformation Order of Category II Pairs

Suppose we want to use a priority approach based on matchability as the greedy approach may not work.

Transform Category II pairs after their *left-lobe* matchability option is exhausted, because of incentive compatibility. But in which priority order of pairs?

Difficulties: 2. Transformation Order of Category II Pairs

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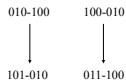
Definition

Define a directed graph on the set of *Category II pair types* ($X - Y$ such that $X \not\leq Y$ and $X \not\leq \rho(Y)$), that we will call the *precedence digraph*, where for any *Category II pair types* $X - Y$ and $U - V$:

$$X - Y \longrightarrow U - V \iff X \leq V, U \not\leq Y \text{ \& } U \leq \rho(Y).$$

If $X - Y \longrightarrow U - V$, we will also say that $X - Y$ *precedes* $U - V$.

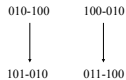
Precedence Partial Order for Two Sizes



► Two Sizes

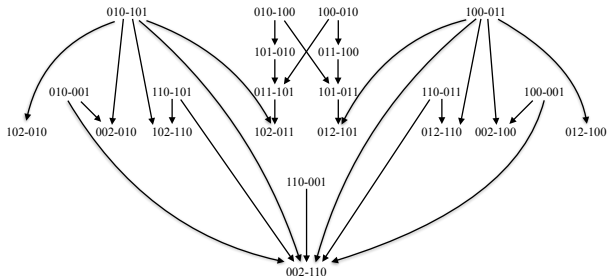
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Precedence Partial Order for Two Sizes



► Two Sizes

Precedence Partial Order for Three Sizes



Lemma (from graph theory)

*Given an acyclic digraph, there exists a linear order of all nodes, known as a **topological order**, L , that is consistent with the digraph:*

$$x \rightarrow y \implies xLy$$

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Lemma

The precedence digraph on *Category II pair types* is acyclic.

Thus, a *topological order* of the precedence digraph of *Category II pair types* exists, and we can use this as our priority order over transformation.



Fix a **topological order** over **Category II pairs** as i_1, \dots, i_K and a **priority order** over all pairs. Given a willingness profile R :

Step 0. Direct transplant **left-lobe** compatible and willing **Category III** pairs.

Transform willing **Category I** pairs.



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Fix a **topological order** over **Category II pairs** as i_1, \dots, i_K and a **priority order** over all pairs. Given a willingness profile R :

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Any matching in G^K that matches all pairs in \mathcal{I}^{K+N} in exchanges together with the fixed direct transplants is the outcome.



Theorem

The *precedence-order induced priority mechanism* satisfies the following:

- *for given topological and priority orders and a willingness profile, all matchings that can be chosen by the mechanism are Pareto indifferent,*
- *individual rationality,*
- *Pareto efficiency, and*
- *incentive compatibility.*

► Simulations



Intuition of the Proof.

Pareto indifference: We extend the matchable set in every step and only *transform* pairs that cannot be matched through *left lobe* in addition to the previously committed pairs. Thus those can only be matched through *right lobe*.



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Incentive compatibility: Acyclicity of the precedence digraph implies that *transformation* a willing **Category II Pair** i_k is independent of the willingness types of its lower-prioritized “graph neighbors.” Thus, they cannot affect how i_k is matched by manipulating their own willingness types.





Proposition

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Proof.

Consider an exchange pool with two sizes 0,1 such that

$$n(101 - 011) = 1, \quad n(011 - 100w) = 2, \quad n(100 - 011) = 1.$$

Any **left-lobe** donation or total transplant maximizing matching generates two exchanges with types

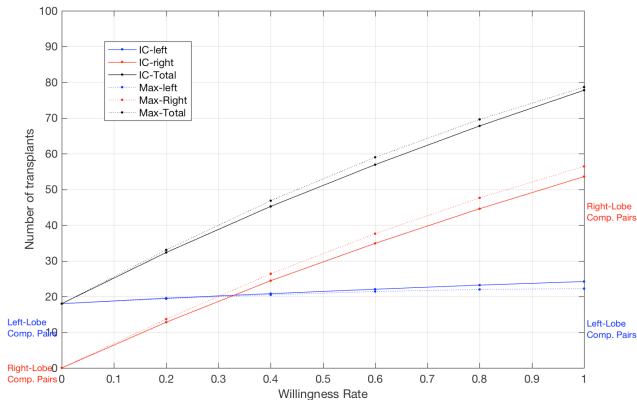
100 - 011 & 011 - 100 (type 1) 101 - 011 & 011 - 100w (type 2),
resulting with 4 transplants and 3 **left-lobe** donations.

Observe that one of the willing pairs of type 011 - 100 is matched to donate its **left lobe** (in type 1 exchange above) while the other one is matched to donate its **right lobe** (in type 2 exchange above). The one matched to donate its **right lobe** can manipulate. □

Simulations: Cost of Incentive Compatibility

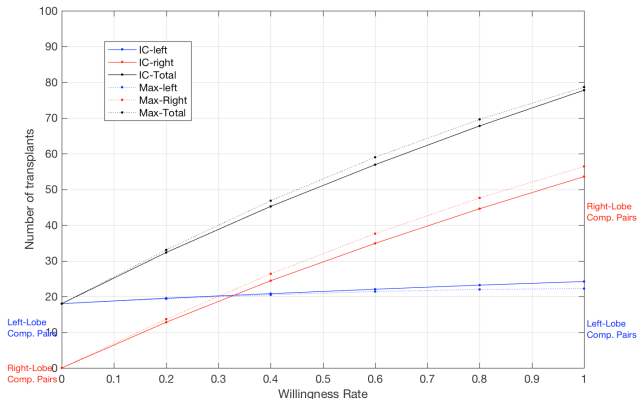


Simulated 100 pairs using South Korean patient/donor sizes and blood-type composition.





Simulated 100 pairs using South Korean patient/donor sizes and blood-type composition.



Very small cost of incentive compatibility!



- Model of **live-donor lobar liver exchange** as a market design problem. Information/incentive problems are modeled and solved through a **PE + IC mechanism**.
Simulations: Relative cost of IC is less than 2%
- **Size incompatibility** increases the benefit from exchange, more gains plausible with respect to kidney exchange
- Off-the-shelf-implementable mechanism in Middle East and East Asia: Liver transplants are more complex, two-way may be the way to start the market design
- **Implications for matching theory in general**: A new class of bilateral exchange mechanisms for **n -dimensional vector partial-order induced** weak preferences:
 - Examples: vacation house exchanges, time/favor exchanges
 - Two-size model with three dimensions is of independent interest: Induces a fully-symmetric model where **greedy mechanism design** is possible.