Post-Donation Regret and Anxiety Among Family Living Liver Donors: The Mediating Roles of Family Relationships and Sense of Mastery

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Background: Post-donation regret in family living liver donors can impact their mental well-being. This study examined whether the relationship between post-donation regret and anxiety is mediated by family relationships and a sense of mastery.

Material/Methods: We conducted a secondary analysis of de-identified cross-sectional data from a prior study that included 124 living liver donors. These donors underwent partial hepatectomy between January 2011 and March 2021 at a tertiary hospital in Seoul, South Korea. The data included demographic and clinical characteristics, along with the results from administering the following measures: the Post-Donation Regret Scale, Family Relationships Index, Pearlin Mastery Scale, and the Generalized Anxiety Disorder-2 scale.

Results: Among family living liver donors, 5.6% had anxiety after donation. The total effect of post-donation regret on anxiety was significant (B=0.41, p<0.05). However, the direct effect of regret on anxiety was not significant (B=-0.05, p=0.733). Post-donation regret had indirect effects on anxiety, solely through family relationships (B=0.329, 95% CI=0.130, 0.563) and sequentially through family relationships and mastery (B=0.088, 95% CI=0.008, 0.232), even after controlling for sex, age, postoperative complications, years since donation, and recipient’s death. In addition, postoperative complication was a predictor of anxiety (B=0.64, p<0.05).

Conclusions: Providing family-centered and mastery-enhancing interventions may help alleviate the anxiety of family living liver donors.

Keywords: Anxiety • Emotions • Family Relations • Liver Transplantation • Living Donors

Full-text PDF: https://www.annalsoftransplantation.com/abstract/index/idArt/944176
Introduction
Liver transplants are the second most frequently performed solid organ transplantation [1]. Over the past decade, living donor liver transplants (LLDTs) have become increasingly popular, especially in Asia, with South Korea emerging as the leading country in terms of LLDT activity [2]. A total of 1158 LLDTs were performed in South Korea in 2021, representing 76% of all liver transplants [3]. Characteristically, more than 90% of donor-recipient relationships of LLDTs in Asia, including South Korea, were within a second degree of familial relation [3-5].

Liver living donors (LLDs) reported moderate anxiety during the postoperative phase, with no discernible decline in anxiety levels over time [6]. Prevailing research indicates a higher prevalence of postoperative anxiety compared to postoperative depression among organ donors [7,8]. Qualitative investigations have shed light on the LLDs’ postoperative psychosocial experiences, emphasizing the importance of valuing the donation decision as pivotal for effective adjustment after donation [9,10]. Additionally, quantitative studies have highlighted the impact of negative emotions such as regret on the mental quality of life of LLDs, with post-donation regret correlating with anxiety [11,12].

Family donors have acknowledged their vulnerability to ongoing uncertainties after donation, endeavoring to restore normalcy to familial routines, and maintain control over their well-being [13]. The adjustment of the donor has been shown to exert a substantial influence on the overall familial well-being, with strained relationships with the recipient exacerbating anxiety among LLDs [13,14]. Nevertheless, despite familial relationships being central to the majority of donation cases, their assessment and impact within the transplant process have remained relatively underexplored [15]. Moreover, many family LLDs embark on the donation process willingly, perceiving it as a means to contribute to the recipient’s and their own family’s future well-being [16]. Such donors may approach the donation process with a sense of mastery, believing in their ability to shape the trajectory of their lives. Conversely, individuals who experience coercion or exhibit hesitation until the eve of surgery tend to manifest greater anxiety regarding their future after donation [13]. Therefore, there exists a need to explore the roles of family relationships and the sense of mastery in understanding the association between post-donation regret and anxiety.

The stress process model (SPM) posits that individuals’ resources serve as crucial mediators between stressors and mental health outcomes, delineating the process through which stressors manifest into psychological distress [17]. The SPM framework, previously applied to elucidate the psychological dynamics of caregivers and general adult populations [18,19], holds relevance for understanding the experiences of family organ donors. Given their comparable health status and the caregiving strain experienced by donors in supporting recipients [20,21], the SPM framework can be effectively utilized to identify stressors and resources influencing post-donation anxiety symptoms among family LLDs. According to this theory, primary stressors include subjective indicators (eg, negative reaction to an event), which predict mental health outcomes (eg, anxiety). In these relationships, role strains (eg, family conflicts) and personal resources (eg, mastery, self-esteem) can mediate stressors’ effects on mental health [22].

The literature shows that negative perceptions and reactions to caregiving predicted family caregivers’ distress, which was mediated by poor family relationships [23]; anxiety in caregivers was found to be inversely correlated with mastery [24]. In general adult populations, regret influences anxious arousal [25], people with problematic family relationships and family functioning were likely to have high anxiety [26,27], greater mastery can decrease levels of anxiety symptoms [28,29], and family functioning satisfaction and sense of mastery show a positive relationship [30]. Therefore, the relationships between these 4 variables can be extended to family organ donors. Identifying roles in post-donation regret, family relationships, and mastery could be helpful in designing effective management strategies for post-donation anxiety. Therefore, for family LLDs, it is worth exploring how the resources may buffer the effect of post-donation regret on anxiety. Consequently, we propose the following hypotheses:

H1: Post-donation regret is positively associated with anxiety.
H2: The association between post-donation regret and anxiety is mediated by family relationships.
H3: The association between post-donation regret and anxiety is mediated by mastery.
H4: Family relationships and mastery sequentially mediate the association between post-donation regret and anxiety.

Material and Methods

Study Design
This cross-sectional study utilized secondary data collected from a project aimed at identifying the well-being and health adaptation of LLDs after donation [11].

Data Source and Participants
The dataset utilized in this study comprised survey responses pertaining to post-donation regret, family relationships, mastery, and anxiety, along with a review of medical records. Medical records of the participants who responded to the survey were obtained, including length of hospital stay, postoperative complications, years since donation, and recipient death. Data collection
of survey and medical record occurred between February and July 2021 at the transplant center of the Seoul National University Hospital (SNUH), in South Korea. The accessible population consisted of 535 LLDs who underwent partial heptatectomy at this transplant center within the preceding decade (from January 2011 to March 2021). Out of the 535 potential participants, 124 donors participated in the original study. The sample of the dataset comprised a total of 124 LLDs who were 19 years or older, within the second degree of familial relations with the recipients (donors were the recipients’ child, spouse, sibling, or parent), and were at least 1 month after donation [11]. For the analysis of the mediation effect, 120 LLDs were included in the mediation model after accounting for missing data.

A post hoc power analysis for linear multiple regression was performed utilizing the G*Power 3.1.9.4. With an alpha level of 0.05, a medium effect size of 0.15, 8 predictors (including post-donation regret, family relationships, mastery, and 5 covariates), and a total sample size of 120, the calculated post hoc power (1-β) was 85.1%. Moreover, considering the recommendation that the sample size for path analysis should be 15 times the number of variables [31], the sample size of this study was deemed acceptable.

Instruments

Post-Donation Regret

Post-donation regret was evaluated by a one-item questionnaire, “If you could go back to before your donation, would you still donate?” LLDs could answer based on their prevailing thoughts and sentiments from the immediate postoperative period to the time of the survey. The questionnaire employed a 4-point Likert scale, ranging from 1 (very likely), to 4 (definitely not). A higher score represents greater regret. The questionnaire was developed after a literature review [12,32], and its face validity was reviewed by 5 LLDs and 5 liver transplant professionals. They qualitatively assessed its readability, suitability, and clarity.

Family Relationships

The Family Relationships Index (FRI) [33] was utilized to evaluate the quality of familial relationships. Employing a Korean adaptation of the FRI [34], this instrument gauged LLDs’ usual perceptions of internal family dynamics, including those involving the recipient. Designed primarily for conjugal and nuclear family units, the FRI assesses familial adjustments to life transitions and crises [33]. The instrument comprises cohesion and expression-conflict subscales, consisting of a total of 20 items rated on a 5-point Likert scale (ranging from 1=“not at all true” to 5=“very true”). The cohesion subscale assesses levels of support, dedication, assistance, and open communication among family members while the expression-conflict subscale assesses the degree to which family members express anger and disagreement [34]. Total scores on the FRI range from 20 to 100. Higher scores indicate that a family has greater cohesion and expressiveness, as well as lower conflict, which reflects a supportive family environment [33]. The Korean adaptation of the FRI was validated and proved to be reliable, with Cronbach’s α values ranging between 0.74 and 0.77 [34]. In this study, the Korean version of the FRI showed an excellent Cronbach’s α coefficient of 0.93.

Mastery

The Pearlin Mastery Scale (PMS) [35,36] was used to assess LLDs’ sense of mastery, reflecting their usual belief in the degree of control they exert over their life circumstances, as opposed to viewing life events fatalistically. It comprised 7 items rated on a 4-point scale: 1 (strongly disagree) to 4 (strongly agree). The total score could vary from 7 to 28, with higher values suggesting stronger mastery. The Korean version of the PMS proved to be reliable, with a Cronbach’s α value of 0.75 [36]. A Cronbach’s α in this study was 0.83.

Anxiety

Anxiety symptoms were evaluated using the Generalized Anxiety Disorder-2 (GAD-2) [37], a brief two-item measure. This instrument assesses the presence of anxiety symptoms experienced over the preceding 2 weeks. Further details and access to the GAD-2 screener are available at https://www.phgscreeners.com/select-screener. Every response is rated on a 4-point Likert scale (0=“not at all” to 3=“nearly every day”), with total scores between 0 and 6. An increased level of anxiety symptoms is indicated by a higher overall score. A total score of 3 or greater was considered positive for anxiety screening. It has already been established that the GAD-2 has suitable validity and reliability in the Korean population [38]. The Cronbach's α in this study was 0.84.

Demographic and Clinical Information

Demographic information about LLDs was obtained, including data on sex (1=female/0=male), age (continuous), relationship to the recipient (1=child/0=spouse, sibling, parent), and marital status (1=married/0=else). Clinical information was obtained, including length of hospital stay (continuous), postoperative complications (1=Grade II-III in Clavien-Dindo classification/0=no complication or Grade I in Clavien-Dindo classification), years since donation (continuous), and recipient death (1=death/0=alive). According to Clavien-Dindo classification, Grade I complications are considered natural sequelae after surgery not requiring any intervention; Grade II or III complications need medication or medical intervention [39].
Ethical Consideration

The current study was approved by the Institutional Review Board (IRB) of SNUH; since this study used secondary data, informed consent from the participants was waived by the IRB of SNUH (approval no.: 2210-085-1368). Data without identifiers were used for data analysis.

Data Analysis

Descriptive analyses were performed to determine sample characteristics. The normality of the study variables (post-donation regret, family relationships, mastery, and anxiety) was examined using the Shapiro-Wilk test, suggesting that these continuous variables were not normally distributed. Hence, Spearman’s rho correlation coefficient, a nonparametric statistical method for correlations, was used to detect the association between the 4 variables. SPSS version 25.0 was used to perform all the analyses.

To test the serial multiple mediation hypotheses illustrating psychological mechanism of LLDs after donation, Model 6 of SPSS PROCESS v.4.1 macro was utilized [40]. This approach, employing bootstrapping, does not require the assumption of normality [41]. Post-donation regret was entered as the independent variable, anxiety as the dependent variable, family relationships as the first mediator, and mastery as the second mediator. These 4 variables were treated as continuous variables. Notably, mediation analysis of the effect of post-donation regret on anxiety was conducted after controlling for sex, age, postoperative complications, time since donation, and the recipient’s death, which were identified as predictors of anxiety in previous research [14]. Using 10,000 bootstrap resamples and a 95% confidence interval (CI), the significance of the direct and indirect effects was evaluated.

Cases were excluded if the missing values constituted more than 50% of the items for each instrument, or if the information for control variables was missing (1 case for the FRI, 1 case for the GAD-2, and 2 cases for recipient death). If the missing values were less than 50% of items for each instrument, the expectation-maximization approach was used (3 cases for the FRI and 3 cases for the PMS). This imputation algorithm substitutes missing data with the best value using maximum likelihood estimation. Consequently, a total of 120 cases were analyzed in the mediation model.

Results

Sample Characteristics

Table 1 shows LLDs’ demographic and clinical characteristics. Seventy LLDs were male (56.5%). Their mean age was 37.9±11.4 years. Most of LLDs were children of the recipients (71.0%). Over half of the LLDs were single (55.6%). Among 124 LLDs, 7 (5.7%) experienced Grade II complications, and only 3 (2.4%) had Grade III complications. The average time since donation was 2.1±2.2 years. A small percentage of the recipients (8.9%) had died. The prevalence of regret and anxiety was 3.2% and 5.6%, respectively.

Correlations Between Study Variables

Table 2 outlines the ranges, mean scores, and bivariate correlations between the 4 main variables. Post-donation regret was negatively correlated with family relationships (p<0.001) and mastery (p=0.003); regret was positively correlated with anxiety (p<0.001). Worse family relationships were linked to lower levels of mastery (p<0.001) and higher degrees of anxiety (p<0.001). Furthermore, a negative association was discovered between mastery and anxiety (p<0.001).
Table 2. Descriptive statistics and correlations among post-donation regret, family relationships, mastery, and anxiety (N=124).

<table>
<thead>
<tr>
<th>Variables</th>
<th>Range</th>
<th>M±SD</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>4</th>
</tr>
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<tbody>
<tr>
<td></td>
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<td></td>
<td>(p)</td>
<td>(p)</td>
<td>(p)</td>
<td>(p)</td>
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<tr>
<td>1. Post-donation regret</td>
<td>1-4</td>
<td>1.28±0.55</td>
<td>1</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>2. Family relationships</td>
<td>34-96</td>
<td>77.02±12.42</td>
<td>-0.405***</td>
<td>1</td>
<td></td>
<td></td>
</tr>
<tr>
<td>3. Mastery</td>
<td>10-28</td>
<td>23.97±3.68</td>
<td>-0.264**</td>
<td>0.411***</td>
<td>1</td>
<td></td>
</tr>
<tr>
<td>4. Anxiety</td>
<td>0-6</td>
<td>0.46±0.99</td>
<td>0.311***</td>
<td>-0.494***</td>
<td>-0.343***</td>
<td>1</td>
</tr>
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</table>

** p<0.01, *** p<0.001.

Table 3. Results of the regression analyses testing the serial multiple mediation effect of family relationships and mastery on the relationship between post-donation regret and anxiety (N=120).

<table>
<thead>
<tr>
<th>Predictors</th>
<th>Direct effect (SE)</th>
<th>Total effect (SE)</th>
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<tbody>
<tr>
<td></td>
<td>Family relationships</td>
<td>Mastery</td>
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<tr>
<td>Post-donation regret</td>
<td>-10.60 (1.91)***</td>
<td>-0.49 (0.64)</td>
</tr>
<tr>
<td>Family relationships</td>
<td>0.10 (0.02)***</td>
<td>-0.03 (0.01)***</td>
</tr>
<tr>
<td>Mastery</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Covariates</td>
<td>Sex</td>
<td>-0.73 (0.63)</td>
</tr>
<tr>
<td></td>
<td>Age</td>
<td>0.05 (0.03)</td>
</tr>
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<td></td>
<td>Postoperative complications</td>
<td>-2.82 (1.12)*</td>
</tr>
<tr>
<td></td>
<td>Years since donation</td>
<td>-0.17 (0.14)</td>
</tr>
<tr>
<td></td>
<td>Recipient death</td>
<td>0.30 (1.08)</td>
</tr>
<tr>
<td></td>
<td>R²</td>
<td>0.24***</td>
</tr>
</tbody>
</table>

* p<0.05, ** p<0.01, *** p<0.001.

Serial Multiple Mediating Effects of Family Relationships and Mastery

Table 3 and Figure 1 show the serial multiple mediation effects of family relationships and mastery on the association between post-donation regret and anxiety after controlling all covariates. Post-donation regret had a statistically significant total effect on anxiety (B=0.41, p<0.05; R²=0.15, p<0.01). However, the direct effect of post-donation regret on anxiety was insignificant (B=-0.05, p=0.733), indicating a significant total indirect effect. In the multiple mediation model, an additional predictor of anxiety was postoperative complications of Grade II or III (B=0.64, p<0.05).

Table 4 presents the indirect effects in the serial multiple mediation model. The total indirect effect of post-donation regret on anxiety was significant (B=0.457, 95% CI=0.194, 0.761). The indirect effect through family relationships was found to...
findings, indicating a post-donation prevalence rate of anxiety acute pain and discomfort. A meta-analysis corroborates these previously in good health, juxtaposed with the experience of may stem from the unfamiliarity of undergoing surgery when anxiety levels. Additionally, the anxiety experienced by donors unmet expectations could plausibly contribute to heightened complications exceeding their expectations [11], the resultant impact on LLD’s anxiety levels. While a minority of LLDs reported revealed that postoperative complications exerted a discernible suits owing to organ donation [43]. Moreover, our study re exacerbation by the postponement or disruption of these life pur developments like employment and marriage, may be predisposed to heightened levels of anxiety, potentially ex accelerated by the postponement or disruption of these life purpursuits owing to organ donation [43]. Moreover, our study revealed that postoperative complications exerted a discernible impact on LLD’s anxiety levels. While a minority of LLDs reported complications exceeding their expectations [11], the resultant unmet expectations could plausibly contribute to heightened anxiety levels. Additionally, the anxiety experienced by donors may stem from the unfamiliarity of undergoing surgery when previously in good health, juxtaposed with the experience of acute pain and discomfort. A meta-analysis corroborates these findings, indicating a post-donation prevalence rate of anxiety among LLDs at 10.0%, with anxiety being most pronounced within the initial 3 months after donation [14]. Concerns over unforeseen medical expenses stemming from complications may further exacerbate anxiety levels [14]. Consequently, implementing policies that offer financial support for both donors and recipients may be a salient intervention strategy [43].

Our study underscores the notable influence of post-donation regret on anxiety symptomology. Hence, vigilantly monitoring and addressing feelings of regret among LLDs is important, given their susceptibility to heightened anxiety levels. Notably, only 3.2% of participants in this study reported experiencing feelings of regret regarding their donation decision. This finding aligns with the literature, as evidenced by a systematic review reporting a prevalence rate of post-donation regret among LLDs ranging from 0% to 11.4% [8]. However, more recent research conducted in Germany and Turkey, encompassing a follow-up period of up to 20 years after donation, revealed a regret prevalence rate of 27.5% [44]. Furthermore, a study in Egypt found that 67% of LLDs would not donate again [45]. Noteworthy distinctions between these studies and our own lie in the scope of donor–recipient relationships examined, with our study focusing on second-degree relatives, while the aforementioned studies encompassed relatives up to the fourth degree and non-relatives [44,45]. This may have contributed to the prevalence of regret. The cultural atmosphere may also have influenced the rate of regret across studies. Furthermore, short-term regret (a year after donation) has been reported to be associated with health perceptions and social functioning [46]. Donors had also expressed feeling abandoned owing to the sudden lack of tracking of their psychological issues in a clinical setting after donation [47], which might lead to long-term regret. Therefore, to prevent regret, healthcare providers should continue to care for and help donors to be aware of their health status and to reintegrate into society.

This study also demonstrated that family relationships mediate the associations between post-donation regret and anxiety. Family LLDs with greater regret had worse family relationships, ultimately leading to higher levels of anxiety, but good family relationships can buffer against anxiety. This aligns with findings from a qualitative study that showed the importance of

| Table 4. Indirect effects in the serial multiple mediation model (N=120). |
|------------------|--------|--------|--------|--------|
| B                 | Boot SE | Boot LLCI | Boot ULCI |
| Total indirect effect | 0.457 | 0.146 | 0.194 | 0.761 |
| Post-donation regret → family relationships → anxiety | 0.329 | 0.110 | 0.070 | 0.563 |
| Post-donation regret → mastery → anxiety | 0.040 | 0.069 | -0.105 | 0.180 |
| Post-donation regret → family relationships → mastery → anxiety | 0.088 | 0.058 | 0.008 | 0.232 |

LLCI – lower limit confidence interval; ULCI – upper limit confidence interval.
adequate support from family members in facilitating donors’ recovery and adapting to their future lives [9]. Additionally, Noma et al discovered that the availability of family support prior to donation is crucial in determining donors’ psychosocial states after donation [48]. Our study is meaningful in that it quantitatively explores the positive impact of supportive family relationships on family LLDs after donation. Especially in Asian societies, decision-making for an LDLT is family-centered [49]. In this context, the greater the donor’s regret about the decision to donate, the more likely it is for family conflict or discord to arise. Moreover, poorer family relationships might be correlated with higher levels of anxiety in family LLDs.

An interesting finding in our study is that family relationships and mastery sequentially mediated the association between post-donation regret and anxiety. Similarly, high levels of personal mastery alleviate anxiety symptoms among Asian family caregivers [24]. Our study adds to the evidence that personal mastery among LLDs is influenced by family relationships, which in turn affects anxiety levels. In Eastern cultures, such as South Korea, which value familial commitment, family donor candidates were reported to have more family-related motives for liver donation than self-related motives [50]. Our study suggests that this trend persists after surgery; lower levels of support, help, and encouragement, along with higher levels of conflict within a family, may undermine the sense of mastery among LLDs experiencing regret. However, family members’ expressions of gratitude for the bravery and commitment shown in donating a liver can bolster LLDs’ sense of personal mastery. Moreover, this enhanced sense of mastery might help to alleviate anxiety. Therefore, this study is meaningful as it empirically demonstrates the importance of a family-centered and mastery-based intervention in the care of family LLDs, even after donation.

To provide family-centered care, the first step is to strengthen the relationship between the donor and recipient. Although LLDs may experience both physical and psychological discomfort after donation, they often hesitate to express their struggles for fear of burdening or worrying the recipient, leading them to endure their feelings alone [43,51]. Recipients may feel infinite gratitude towards the donor but may also experience feelings of guilt and hesitation in expressing these openly [51,52]. In some cases, living donation can intensify unsolved tension, disappointments, and resentments between them [52]. Therefore, healthcare professionals should provide education and counseling to help them understand, appreciate, and navigate their shared journey together, facilitating better adaptation to changes in the dynamics of their relationships [20,52].

Additionally, in transplant families, family caregivers may feel overwhelmed with the responsibility of caring for both the recipient and the donor, potentially leading to feelings of inadequacy in attending to the donor’s needs [53]. Conversely, donors may perceive a lack of attention from the family owing to the focus on the recipient’s care [49]. To alleviate the care burden on family caregivers and ensure adequate support for donors, it is essential to implement support systems tailored to the needs of transplant families. These may include the use of care assistants for the recipient, home care services specifically designed for transplant families, and support plans for leisure activities to promote overall well-being within these families [53].

Although the effects of mastery on anxiety are more likely to be significant when preceded by positive family relationships, interventions such as cognitive behavioral therapy can be applied to promote mastery. Providing LLDs with psychoeducation regarding the mechanisms of change can help them recognize fatalistic and helpless attitudes and shift towards more flexible and constructive thoughts, thereby enhancing their sense of mastery [54]. For instance, Wirken et al [55] demonstrated improvements in postoperative psychosocial outcomes by providing cognitive behavioral therapy to living donors via a website. Incorporating the content of mastery into such interventions could further enhance their effectiveness.

This study has limitations. First, the cross-sectional design of the study may not be able to demonstrate causal relationships. However, given that the SPM was used to establish these relationships, they appear plausible. Second, the data collected from a single transplant center could hinder the generalization of our findings. Future prospective studies should include multiple transplant centers in different countries. Third, post-donation regret was measured using a single item with 4 scores, which could result in response bias. Future research should measure regret using questionnaires with more items or granular scoring. Lastly, it is pertinent to acknowledge the inherent limitations stemming from the secondary analysis used in this study. The dataset under scrutiny lacked inclusion of certain potential confounding factors, such as financial status, perceptions of overall health, and social activity, which consequently precluded their incorporation into the mediation model. This limitation necessitates a cautious interpretation of the research findings, mindful of the contextual constraints imposed by the available data.

**Conclusions**

Family LLDs with greater post-donation regret were more susceptible to anxiety symptoms. Furthermore, family relationships and mastery sequentially mediated the relationship between post-donation regret and anxiety, with family relationships being a key element in this mediation. Therefore, regular psychosocial counseling and education that includes family members are recommended. To ultimately reduce anxiety, healthcare professionals and clinical nurses should consistently monitor the psychological status of LLDs and develop...
intervention programs that foster a positive family environment and enhance mastery.

Data Availability Statement

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

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29. Declaration of figures’ Authenticity

All figures submitted have been created by the authors, who confirm that the images are original with no duplication and have not been previously published in whole or in part.


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