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Patient characteristics of deceased organ donors in Switzerland 1998-2008

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Summary

BACKGROUND: Transplantation today faces two major issues: organ shortage, and the increasing age of deceased donors. The former leads to growing waiting lists and, as a consequence, to a higher mortality of patients in need of an organ. The latter results in an augmentation of the proportion of organs from so called extended criteria donors, which, in the past, have been associated with inferior outcomes.

METHODS:Retrospective analysis of the patient characteristics of all deceased organ donors in Switzerland from 1 January 1998 to 31 December 2008. A total of 1004 patient records of brain-dead organ donors (59.4% male, 40.6% female) were analysed for demographic data and causes of death.

RESULTS:The average donor age increased from 43.8 ± 16.9 years in 1998 to 51.7 ± 18.0 years in 2008 (+27.6%; p <0.05). When comparing 2008 with 1998, cerebral haemorrhage accounts for 52.8% of the causes of death in 2008 (+16.7 percentage points), whereas the proportion of cranio-cerebral trauma decreased by 22 percentage points to 16.9%. The largest group of donors are the 16–55 yearolds with a 50% quota in 2008 (–23.6% compared with 1998). During the study period, the \geq 65 year-old group had a 216.2% growth rate, and it accounts 26.6% of the donors in 2008 (p <0.05).

CONCLUSIONS: A shift in the causes of death significantly increased the average age of donors, and transplantation medicine is confronted with a growing number of extended criteria donors. Nevertheless, 10-year survival of transplant recipients is better than ever before.

Key words: organ transplantation; deceased organ donors; extended criteria donors; organ procurement; donor characteristics; cause of death; age; Switzerland

Introduction

Solid organ transplantation nowadays is confronted with two major issues: first, organ shortage, and second, the increasing age of deceased donors. Switzerland has one of the lowest organ donation rates in Europe (15.4 per million of population [pmp] in 1998, 11.8 pmp in 2008). Since

2003, the gap between donated organs and patients on the waiting list has widened dramatically. By the end of 2008, 942 patients were listed for a transplantation (+47% compared with 2003), while the quota of organ donors and the total number of transplanted organs declined by 5.3% and 3%, respectively [1]. As a consequence, the waiting list mortality has risen by 12.7% between 2003 and 2008 [1, 2]. Various reasons can be cited for the mounting average age of the donors. Generally, the demographic trend in "first world" countries shows an increase in life expectancy, so it should come as no surprise that the donors are getting older as well. A more specific reason might be found in the fact that in Switzerland, during the study period, several measures and campaigns to improve the security for leisure activities and road safety were implemented, most prominently the reduction of the legal blood-alcohol limit for driving to 0.5% on 1 January 2005.

Up to now, the transplantation of organs from the so-called marginal or extended criteria donors (ECDs) has been associated with inferior outcomes. There is no generally recognised definition of the criteria which have to be fulfilled by a donor to be considered an ECD, but the following parameters are usually taken account of: donor age above 50–65 years (depending on the organ), prolonged cold and/or warm ischaemia time, and prolonged donor ICU stay. Given that ECDs will possibly become more prevalent (due to organ shortage and the growing average age of the donors), a careful donor and recipient profiling and selection may gain relevance in the not so distant future – but for now, let the results of our analysis of the deceased organ donors' patient characteristics speak for themselves.

Patients and methods

Data were obtained from Swisstransplant, Switzerland's national organ procurement organisation. In a retrospective study, patient records of all deceased (brain-dead) organ donors in Switzerland between 1 January 1998 and 31 December 2008 were reviewed. There were 1004 brain-dead donors in total (59.4% male, 40.6% female) aged on average 46.1 ± 17.6 years. Data were collected and analysed for heart, lung, liver, kidney, and pancreas retrieval.

Statistical methods

Frequency and distribution of donor criteria, overall and number of organs procured per donor were evaluated using Excel 2007 (Microsoft Inc., CA, USA) and GraphPad Prism 5.00 for Windows (San Diego, CA, USA). Significance of differences was analysed in Excel using a two sided Student t-test and a p-value <0.05 was considered significant.

Results

Figure 1 shows the number of deceased organ donors per year in Switzerland from 1998 to 2008. In 1998, the total of donors was the highest (108) in the whole study period. It then declined gradually to 95 donors in 2001. In the year 2002, there was a 21% collapse of the donor quota, which then bottomed at 75. From 2003 (95 donors) to 2006 (80 donors), the rates still showed a negative trend, whereas the number of donors is increasing again since 2007 (+11.1% in 2008). Statistical analysis revealed, due to the small number of donors per year, no significant decrease of organ donors in the time period between 1998 and 2008. However the linear trend (y = 1.9818x + 103.16), displayed in figure 1, showed a tendency towards a decrease of organ donors in the observed time period.

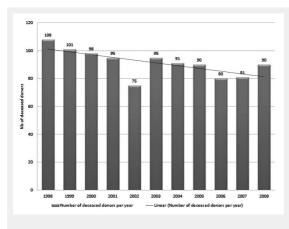


Figure 1

Number of deceased donors in Switzerland, per year

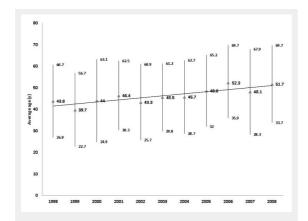


Figure 2

Average age (in years) of the deceased donors per year, displaying ±1.SD (standard deviation) and linear trend.

The mean age of the donors is displayed in figure 2. Even though there are minor variations from one year to another, it shows a clear upward trend from an average of 43.8 ± 16.9 years in 1998 to 51.7 ± 18.0 years in 2008 (+27.6%; p <0.05). This can be confirmed by the linear trend (y = 0.9527x+40.565), which clearly shows an increase in the average age of organ donors within the time period of 1998 to 2008.

The mean number of organs procured per donor increased during the study period, even though it varied quite substantially between the years, with a minimum of 3.30 ± 1.33 organs per donor in 1998, and two peaks with 3.85 ± 1.3 organs per donor in 2002 and 3.80 ± 1.28 in 2006, respectively 3.80 ± 1.35 in 2007 (fig. 3). There is a statistical significant increase in the number of procured organs per donor comparing the first time period between 01/1998 to 06/2003 and the second time period 07/1998 to 12/2008 (p <0.05).

The distribution of the causes of death is displayed in figure 4. When comparing the 2008 data with those of 1998, it is striking to note that cerebral haemorrhage (CHE) accounts for more than half of the causes in 2008 (52.8%, +16.7 percentage points). Conversely, the share of craniocerebral trauma (CTR) decreased by 22 percentage points to 16.9% of the donors' deaths. The proportion of donors who died of suicide (SUI) was relatively stable over the years (median 5.6%), except for two peaks in 1999 and 2002 (12.9% and 13.3%, respectively). In 2008, donors who suffered anoxia (ANOX) became the third largest group (13.5%, +8.9 percentage points). A quite stable share of less than 10% during the whole study period was found for cerebrovascular insults (CVI, 0-6.1%), tumours (TUMOR, 0-3.7%), others (OTH, 0-7.8%), and no available data (N/A, 0-1.2%).

Figure 5 gives an overview of the development of the donors' age distribution. When comparing the percentage shares in 1998 with those in 2008, the ≥65 years group (usually considered ECDs) had the most remarkable growth rate (+216.2%), and reaches a 26.6% share of all deceased donors in 2008. The largest group remains the 16–55 year-olds with a share of 50% of all donors in 2008 (−23.6% compared with 1998). In 2008, the joint share of the 56–64 year-olds group and the ≥65 years group

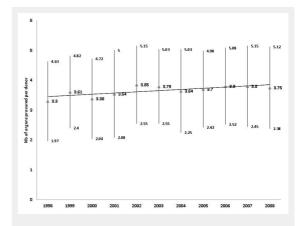


Figure 3

Average number of organs procured per donor, displaying ±1.SD (standard deviation) and linear trend.

is 46.8%, and it thus almost equals the 16–55 year-olds group. The 0–15 years group is the smallest with less than 10 donors per year.

Figure 6 shows the evolution of transplanted heart, lung, liver, kidney, and pancreas grafts per year. The average number of transplantations per year was $331.9~(\pm 23.4)$, with a minimum of 289 transplantations in 2002 and a maximum of 364 in 1999.

During the comprehensive study period, a total of 3651 heart, lung, liver, kidney, and pancreas grafts were transplanted. Kidneys accounted for 52% of the transplantations, livers for 21.1%, hearts for 11%, lungs for 9.9%, and pancreas for 5.9%. Table 1 summarises the number of organs, the quota of transplantation (which designates the respective percentage of grafts that were actually transplanted) and the average age of the donors, calculated by organ.

The quota of transplantations was 94.7% for kidneys, but only 21.6% for pancreas. As for the average age of the donors, it was highest in the liver grafts group $(49.4 \pm 13.7 \text{ years})$ and lowest in the lung grafts group $(43.7 \pm 15.7 \text{ years})$, without reaching statistical significance (p = ns).

Discussion

As mentioned before, transplantation in Switzerland is confronted not only with a low donation rate, but it is also faced with organ donors whose mean age increased by 7.9 years between 1998 and 2008 (p <0.05). When looking at the average number of organs procured per donor (fig. 3), it seems that the discrepancy between a rising number of transplant candidates and a restricted donor pool has lead to the transplantation of organs previously not allocated for various reasons. The noteworthy quota of organs procured per donor in Switzerland is similar to the

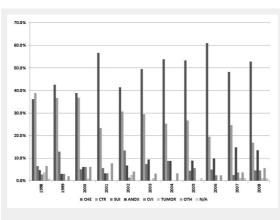


Figure 4
Causes of death (percentage) of the donors per year.

one in the UK, which has a comparable donation rate [3], and it is considerably higher than in countries with elevated donation rates [4, 5]. Unsurprisingly, the peaks in the number of organs procured per donor occurred in the same years (2002, 2006) when the number of donors was lowest (fig. 1), resulting in a relatively stable proportion of transplanted organs over the years (fig. 6). The high quota of organs procured per donor, ranging from 3.30 ± 1.33 to 3.85 ± 1.30 during the study, is even more remarkable, given the development of the age distribution of the donors, with the \geq 56 years old group having a share of over 40% since 2006 (fig. 5). These findings are astonishing considering that a large recently published analysis of the United Network for Organ Sharing (UNOS) deceased donor registry data re-

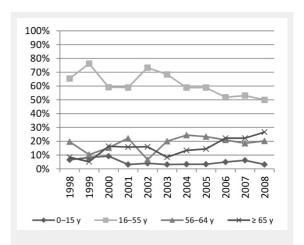


Figure 5

Age distribution of the donors per year, displaying the percentage of donors within the age range of 0 to 15 years, 16 to 55 years, 56 to 64 years and 65 years and above.

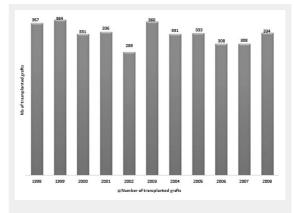


Figure 6

Number of transplanted solid organs in Switzerland. Results displayed for organs procured from deceased donors. Living is not included in this figure.

Table 1: Summary of the transplanted grafts 1998–2008 per organ (heart, lung, kidney, liver and pancreas (whole pancreas and islets); quota of transplantations = percentage of procured organs in relation to the total number of donors in this study (n = 1004)); average age of donor, displayed by procured organ. Results displayed as mean value ± 1. SD (standard deviation).

	Heart	Heart		Lung		Kidney		Liver		Pancreas	
Number	402	402		362		1900		770		217	
Quota of	40.0%	40.0%		36.1%		94.7%		76.7%		21.6%	
transplantations											
Donors' age	46.4	±16.7	43.7	±15.7	48.65	±13.85	49.4	±13.7	45.6	±9.3	
(years, mean ±1 SD)											

vealed that both, increasing age, and anoxia as cause of death significantly influence organ yield [6]. In this study, the average number of organs procured per donor was 3.13 ± 1.77 (compared with 3.65 ± 0.18 in our study), and the mean age of the donors was 40.1 ± 19 years while in our study population, it was 46.1 ± 17.6 years.

The distribution of the causes of death of the donors (fig. 4) shows growing shares for cerebral haemorrhage as well as for anoxia, while cranio-cerebral trauma, the largest group in 1998, lost more than half of its size during the study. These data are in line with a recently published overview on the causes of death in Switzerland by the Federal Statistical Office [7]. Table 2 below shows a selection of these data with the absolute numbers and the incidence per 100'000 population. When contrasting 2008 with 1998, road accidents (supposedly causing cranio-cerebral trauma) decreased 33.8% by number and 40.3% by incidence per 100'000 population.

A similar downward trend in the share of accidents was also observed in the Eurotransplant area, where in 1998, 32% of the deceased donors belonged to this category, while in 2008, the rate had dropped to 22.9% (–28.4%) [8, 9]. As pointed out in the Introduction section, data on the incidence of road accidents in Switzerland suggest that the reduction of the legal blood-alcohol limit for driving to 0.5‰ on 1 January 2005 may have resulted in a substantial reduction of lethal road accidents, which in total decreased by 19.8% and 9.5% in the years 2005 and 2006, compared with the respective preceding year [10].

Considering the question whether the more frequent transplantation of grafts from ECDs might have a negative impact on the outcomes, recent studies in the use of ECD heart [11, 12], lung [13–16], liver [17–19], kidney [20–22], and pancreas grafts [23] show somewhat promising results. In summary, most studies emphasise the potential of ECD grafts to reduce waiting list mortality [24–26], as well as the significance of an appropriate donor and recipient profiling and selection, e.g., age matching and risk assessment [17, 21, 23, 24, 27, 28]. However, and despite this encouraging upshot, it should be kept in mind that the data presented in our study show that in Switzerland - as opposed to other countries, where ECDs are supposed to expand the donor pool, grafts from ECDs are not supplementing the availability of organs but replacing grafts no longer available from "standard" donors. This fact must be taken into account when comparing outcome data from different countries.

Limitations of the study: The authors are aware that this study gives only an overview of deceased Swiss donor characteristics in the defined study period between 1998 and 2008. However for the first time these data have been analysed in the context of a long time-period, showing significant increases in the average age of donors and in the number of organs procured per donor. The results clearly underline the importance of taking into account donor characteristics in a small country like Switzerland with a very low number of donors per million habitants, compared to the countries surrounding us, when analysing and comparing outcome data with other countries.

In conclusion, the present study shows that in Switzerland, like in other European countries, organ shortage leads to a higher number of organs procured per donor. Furthermore, a shift in the causes of death markedly increased the average age of donors, and transplantation medicine is confronted with a growing number of ECDs. The good news is that despite these adversities, outcomes do not worsen, and that nowadays, amongst others due to refined immunosuppressive regimens [29–31], the 10-year survival of transplant recipients is better than ever before.

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	1998			2008				
	Number	Percentage	Incidence per 100'000 pop. 601	Number (compared with 1998)		Percentage	Incidence p. 100'000 pop. (compared with 1998)	
Total number of deaths in Switzerland	62567			61233	(-2.1%)	100%	478	(-20.5%)
Cerebrovascular diseases	4854	8%	41	4077	(-16.0%)	7%	27.4	(-33.2%)
Cardiac diseases (total)	18852	30%	166	16956	(-10.1%)	28%	118	(-28.9%)
Ischaemic cardiac diseases	11257	18%	101	8861	(-21.3%)	14%	63	(-37.6%)
Circulation system	25443	41%	222	22321	(-12.3%)	36%	155	(-30.2%)
Pulmonary embolism	333	1%	3.1	300	(-9.9%)	0%	2.2	(-29.0%)
Accidents and violence	3561	6%	41.6	3742	(+5.1%)	6%	36.3	(-12.7%)
Accidents (total)	2036	3%	22.2	2274	(+11.7%)	4%	20.1	(-9.5%)
Road accidents	547	1%	7.2	362	(-33.8%)	1%	4.3	(-40.3%)
Suicide	1371	2%	17.5	1313	(-4.2%)	2%	14.4	(-17.7%)

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Figures (large format)

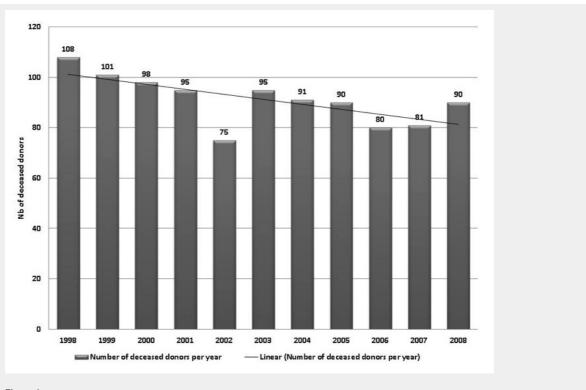


Figure 1
Number of deceased donors in Switzerland, per year.

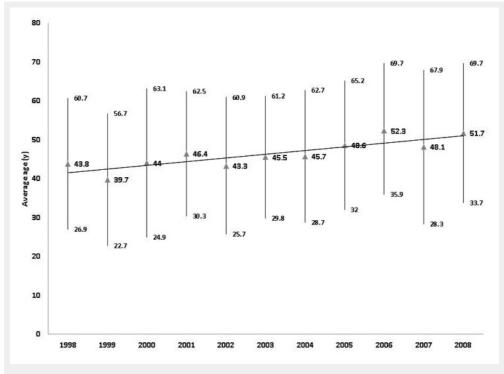


Figure 2

Average age (in years) of the deceased donors per year, displaying ±1.SD (standard deviation) and linear trend.

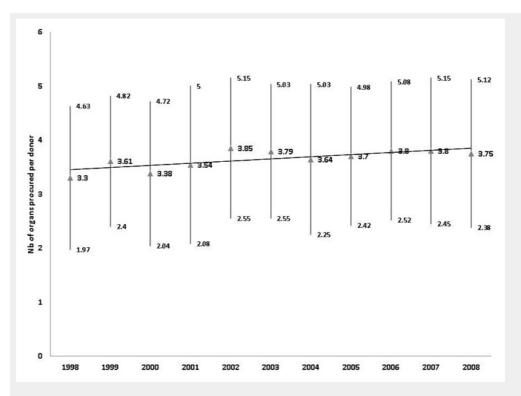


Figure 3

Average number of organs procured per donor, displaying ±1.SD (standard deviation) and linear trend.

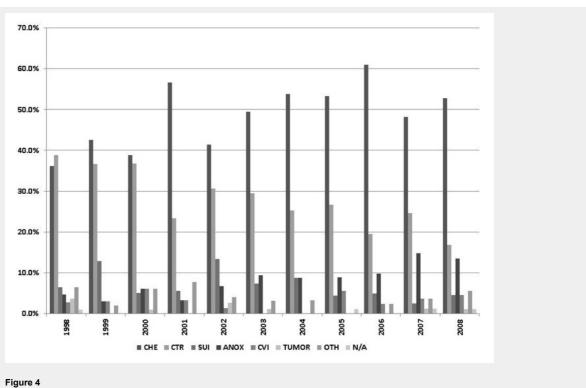


Figure 4

Causes of death (percentage) of the donors per year.

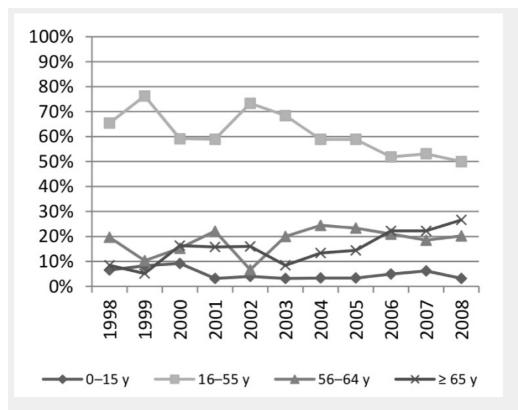


Figure 5

Age distribution of the donors per year, displaying the percentage of donors within the age range of 0 to 15 years, 16 to 55 years, 56 to 64 years and 65 years and above.

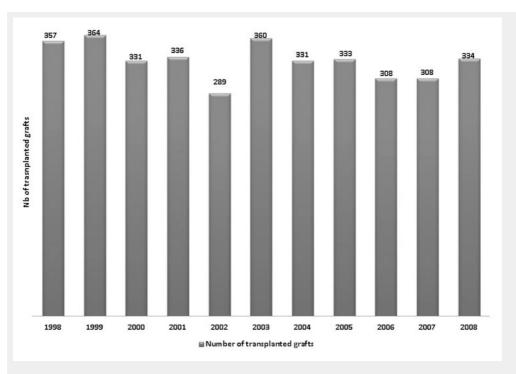


Figure 6

Number of transplanted solid organs in Switzerland. Results displayed for organs procured from deceased donors. Living is not included in this figure.