

Machine perfusion in clinical trials: "machine vs. solution effects"

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Dear Sirs,

In their letter Chatauret *et al.* [1] address an interesting issue: in hypothermic machine perfusion for kidney preservation we should have a better understanding of the machine effects versus solution effects.

During the discussions concerning the study design the scientific steering committee also discussed which preservation solutions should be used.

For more than 40 years, KPS-1 is the standard solution for clinical machine perfusion. An increasing number of animal experimental studies demonstrate that machine perfusion with either HTK [2] or UW [3] leads to good results. Nevertheless clinical data are missing so we decided to use the standard solution.

For static cold storage use of HTK or UW is clinical reality in the Eurotransplant region. Therefore both solutions were accepted for cold storage in the trial.

Vaziri [3] performed an experimental study in a porcine autotransplantation model with 60 min warm ischemia and 24 h hypothermic preservation. Four Groups were compared: Machine perfusion was with either KPS1 or UW and Cold storage with either KPS1 or UW.

Since no animal in the group with cold storage and UW survived, the authors concluded that UW is bad for kidney preservation because of its potassium content. In a similar autotransplantation model with 60 min warm ischemia but 4 h hypothermic treatment [4] we found different results: four of seven animals survived with cold storage in UW. Somehow dissociated the Chatauret *et al.* interpret the group with machine perfusion and UW. This group has the highest survival whereas the differences between MP with KPS1 and CS with KPS1 were not so "obvious." You would expect the deleterious effect of a preservation solution to be aggravated by MP that permanently exposes kidney parenchyma to the preservation solution.

Chatauret *et al.* expect "extrapolating from their results that UW solution in the CS arm of the multicenter trial pulls down the survival curve." This is pure speculation.

Of all the experimental and clinical trials up to now at least no inferior results have been shown for UW compared with HTK [5,6].

We actually analyzed the results in the ECD subgroup of the multicenter trial comparing UW and HTK preserved kidneys. There were no significant differences for DGF rates (25% UW vs. 33% HTK), PNF rates (10% UW vs. 13, 9% HTK) and 1 year graft survival (90% UW vs. 78% HTK).

Chatauret *et al.* remind us that in experiments designed to compare two conditions only one parameter must change between the two conditions. An experimental animal DCD model with long, warm ischemia times and preservation times seems to change more than one parameter, compared with our clinical study.

The authors also see a "clear superiority of MP over CS in their study independent of the solution used."

We do not see a reason to question our conclusions concerning MP.

Although new solutions belong to the most interesting developments in transplantation, from our point of view multicenter trials comparing different solutions in machine perfusion are not realizable in the next few years without additional data.

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