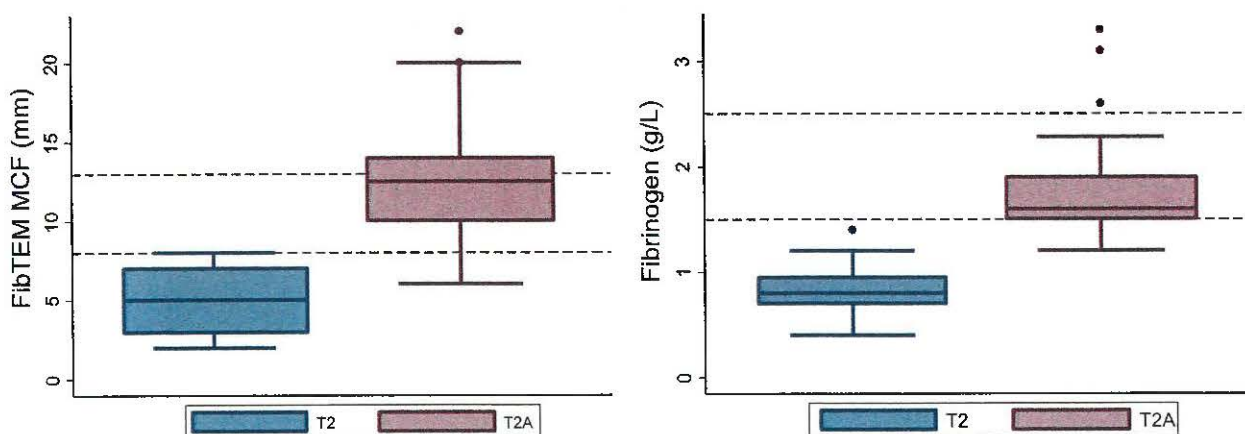


## 8. Primary endpoint: FibTEM MCF and Fibrinogen levels 5 minutes post dose (T2A)

The ROTEM-based dosing formula produced a consistent overshoot in FibTEM MCF (median achieved level 13mm, predicted level 9mm). However the precision was reasonable (interquartile range 10 to 14mm). A similar pattern was seen for achieved fibrinogen levels. Fibrinogen levels post-dosing were within the targeted range of 1.5 to 2.5 g/L in 43/60 (72%) patients receiving IMP. Of not, all patients achieved a fibrinogen level >1.0 g/L (minimum 1.2 g/L), and none would be deemed supratherapeutic (maximum 3.3 g/L).

This was achieved using an administered TOTAL fibrinogen concentrate dose of 51 to 218 mg/kg. However, this dose includes a factor to incorporate the extracorporeal circuit; the equivalent dose range for patients not receiving cardiopulmonary bypass would be 31 to 87 mg/kg (table).

stats	Fibrinogen (g/L)		FibTEM MCF (mm)		Fibrinogen dose (mg/kg)	
	T2	T2A	T2	T2A	TOTAL	PATIENT
max	1.4	3.3	8	22	218	87
p90	1.1	2.2	8	18	184	74
p75	1.0	1.9	7	14	138	69
p50	0.8	1.6	5	13	114	54
p25	0.7	1.5	3	10	81	41
p10	0.6	1.4	3	9	63	32
min	0.4	1.2	2	6	51	31
N	60	60	60	60	60	60



Fibrinogen (left) and FibTEM MCF (right) levels achieved pre- and post dosing (time point T2 and T2A). Dashed lines represent desired range at T2A, according to the protocol.