## Pathological observations in response to intra-renal administration method

**CD31** 

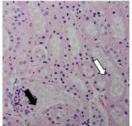
- Previous experiments by MedImmune/Moderna in rat and pig using intrarenal administration of KL10 VEGF-A mRNA did not reveal any pathological findings in kidney. However, no tissue blocks or raw data can be found to do comparative studies of cell specific markers.
- In collaboration with Jane Stubbe (University of Southern Denmark) a PBS control experiment was done (late April) to understand possible impact of method on kidney structures.

Time (min) 0 5 180

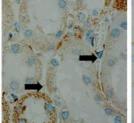
N=4

+/- FA perfusion fix
CD31 IHC (endothelium)
H&E pathology

## H&E 180 min post i.r. PBS administration



Black arrow: Glomerular injury White arrow: Tubular injury



Kidney from 180 min post PBS i.r. with loss of or patchy CD31 in peritubular capillaries (PTC) (black arrows)



Normal kidney with linear CD31 in PTC (white arrows)

## Conclusions

Experimental design acute studies

- Glomerular and tubular damage identified with PBS at same severity/degree as in a previous experiment
  - CD31 signal is disrupted, compatible with endothelial damage
- No damage identified at 5 min time point
- Pathology findings likely related to method of administration