

- In vitro investigation of vas deferens (VD) function from male normal rats in organ baths.
- Useful to investigate the effect of drugs developed on epididymal VD to evaluate the smooth muscle reactivity
- Evaluation of the ability of drugs at modulating epididymal VD smooth muscle tone, organ bath studies:
  - on adrenergic contractile responses elicited by alpha-adrenergic pharmacological stimulation: phenylephrine/norepinephrine (prostatic VD is much less reactive to adrenergic agents)
  - on contractions induced by electrical field stimulation (EFS) (stimulation of efferent nerve terminals)
  - on KCl contractile responses.
- Evaluation of mRNA by RT-PCR
- Evaluation of protein expression: by immunohistochemistry (IHC) or western-blot (WB) in parallel of organ bath studies.

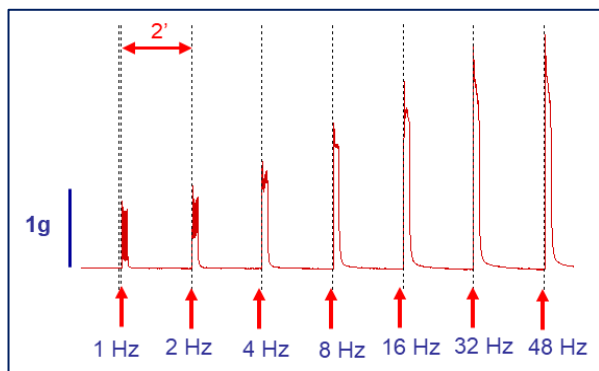


Figure 1: Original tracing showing a frequency-response curve to EFS (300 mA, 10 s, 0.5 ms, 1 to 48 Hz) in rat vas deferens tissue. (Pelvipharm, internal data).

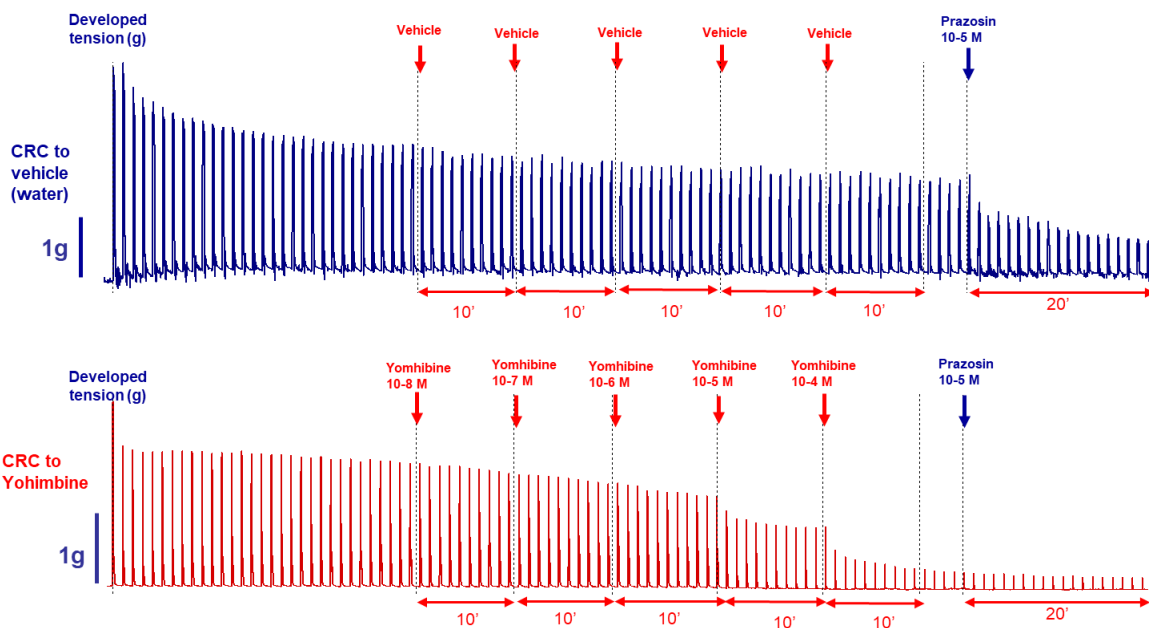


Figure 2: Original tracing showing a continuous EFS stimulation (300 mA, 10 s, 0.5 ms, 10 Hz) and concentration response curve to an antagonist for alpha<sub>2</sub>-adrenoceptors (yohimbine)  $10^{-8}$  M to  $10^{-4}$  M or its vehicle in rat epididymal vas deferens strips. Maximal relaxation was induced by a reference of alpha-1 blocker (Prazosin,  $10^{-5}$  M). (Pelvipharm, internal data).

## Endpoints

- Evaluation of the capacity of a drug to inhibit human or rat vas deferens smooth muscle contractions.
- Determination of potency (**EC<sub>50</sub>**) and efficiency (**E<sub>max</sub>**) of a drug.
- Determination of the affinity (**pA<sub>2</sub>**) of a drug for a human or rat vas deferens receptor.

**NB: Pelvipharm will gladly study the feasibility to fit this experimental model to its client's needs.**