## **Online Supplemental**

Short title: Portal Osmopressor Mechanism Linked to TRPV4

## Portal Osmopressor Mechanism Linked to TRPV4 and Blood Pressure Control

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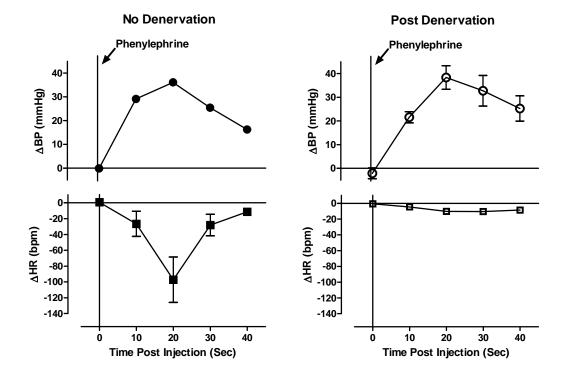
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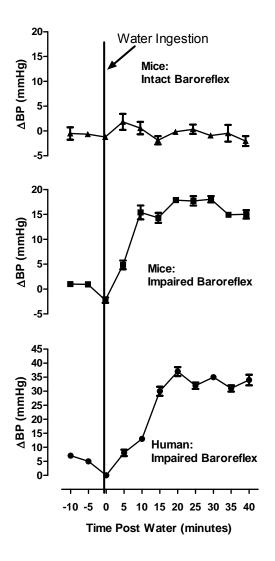
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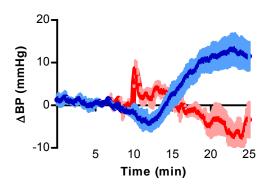
## S1. Blunted HR response to phenylephrine after baroreflex deafferentation.

Representative BP and HR changes after i.v. phenylephrine (t = 0), pre and post baroreflex deafferentation. Phenylephrine challenge was used to validate successful baroreflex deafferentation in mice.

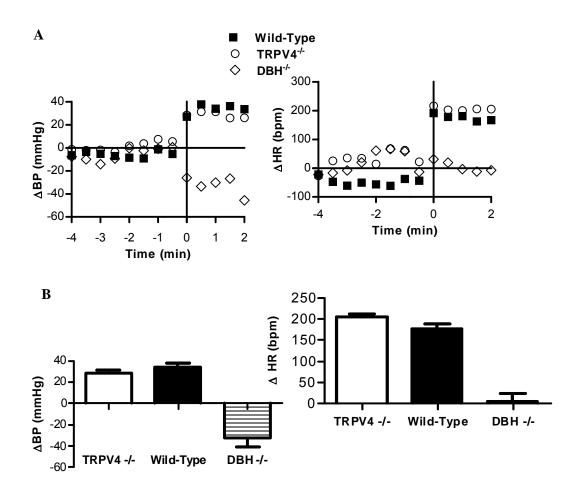


## S2. BP profile during water consumption in humans and mice.

Mice with intact baroreflexes show little BP response to water infusion, since baroreflex buffering attenuates it. Surgically baroreflex-impaired mice, as well as patients with baroreflex impairment, show a robust increase in BP that is sustained well beyond the period of water ingestion. This enhanced pressor response facilitates mechanistic studies of water's cardiovascular effects.



**S3.** Role of plasma volume in pressor response. 150 µL intravenous saline (*red*), given at 10 minutes elicited only a small, transient pressor response. Duodenal (*blue*) infusion of water is shown for comparison.



**S4.** *Trpv4<sup>-/-</sup>* have intact sympathetic efferents. A, Representative tracings of the change in BP and HR during restraint for wild-type ( $\blacksquare$ ), *Trpv4<sup>-/-</sup>* ( $\circ$ ), and *Dbh<sup>-/-</sup>* ( $\diamond$ ) mice. B, Average change in BP and HR during 2-minute restraint.