

# Cochlear Homeostasis Lab

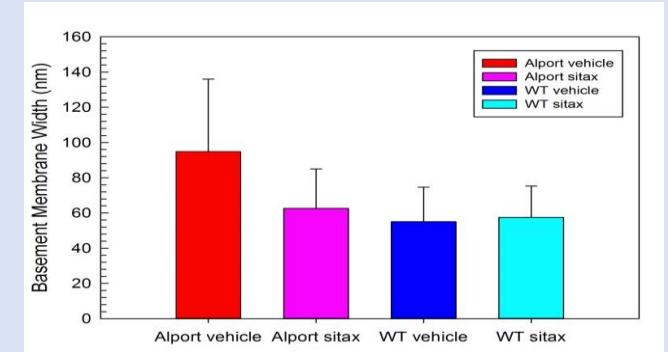
## Michael Anne Gratton, PhD



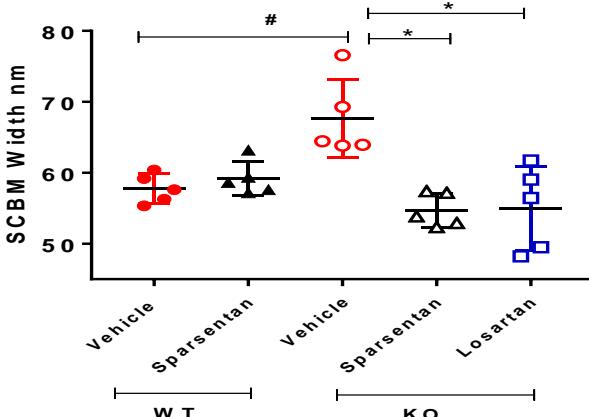
Cochlear basement membrane pathology with lateral wall (stria vascularis & spiral ligament) dysfunction resulting in hearing loss can occur in:

- Alport Syndrome
- Presbycusis
- Diabetes
- Autoimmune disease

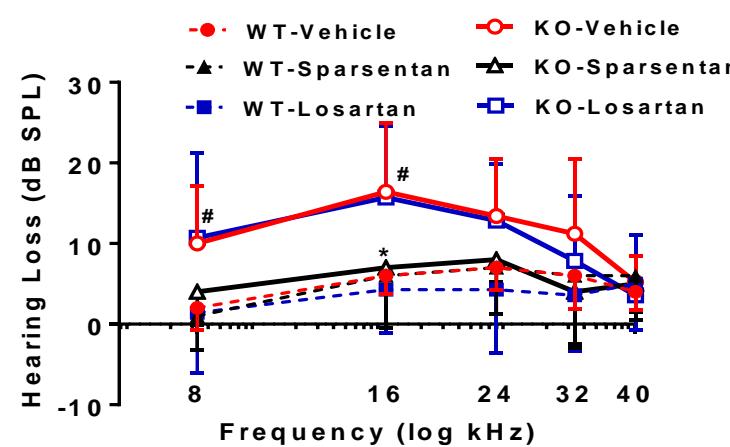
2018: Treatment with an  $\text{ET}_{\text{A}}$  blocker reversed basement membrane pathology in cochlear capillaries (right) and kidney glomerulus of a mouse modelling Alport Syndrome wh (Meehan et al Hear Res 2018)



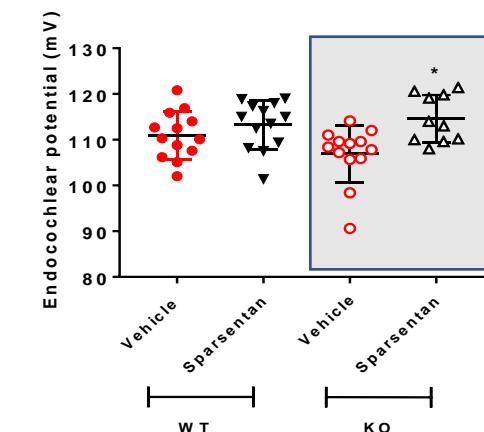
A **dual  $\text{ET}_{\text{A}}$ /AT<sub>1</sub> blocker**, Sparsentan, but not Losartan, SOC, rescues hearing loss in Alport mice



Sparsentan (open  $\Delta$ ) and Losartan (open  $\square$ ) treatment attenuated the progressive stria capillary basement membrane (SCBM) pathology in Alport mice. # and \* =  $p < 0.05$  n=5



Sparsentan (open black  $\Delta$ , but not losartan (open  $\square$ ), the current SOC, prevents hearing loss in Alport mice. # and \* =  $p < 0.05$  n=5



Sparsentan treated Alport mice have a higher endocochlear potential (open  $\Delta$ ), a marker for lateral wall function.

\* =  $p < 0.05$  n=13 KO-Vehicle vs KO SP