

Cambridge University Hospitals

PKD Research Studies

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Objectives

• DRINK Water Intake Trial

• Pain in PKD



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The DRINK Water Intake Trial

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Vasopressin – Water Balance





Animal Studies – Yes!





Male

Female



 Less vasopressin made

- Less vasopressin receptors
- Fewer cysts
- Total Kidney weight smaller
- Kidney function preserved

HWI (1% Agar)

NWI





Human Studies - ??

- NOT RANDOMISED
- SMALL
- NOT DONE OVER LONG ENOUGH
 PERIOD

Benefits of Water

- Cheap
- Widely available
- Few side effects
- Alternative can't tolerate or not suitable for Tolvaptan
- Important question to patients PLRH/PKD Charity

Urine targets – Dilute enough to stop brain making vasopressin





Trial Design

• Randomised

• Open- label

• Feasibility

What are the criteria for taking part?

- Inclusion Criteria
 - ADPKD (Imaging ± Genetics)
 - $Age \ge 16$ years
 - $-eGFR \ge 20ml/min$
 - Able to perform urine dipstick test

What are the criteria for taking part?

- Exclusion Criteria
 - Fluid overload
 - Diuretics and other medications lower salt
 - Tolvaptan
 - eGFR <20ml/min (late CKD 4)</p>

What does taking part involve

- Clinic Visits
 - Physical examination
 - Blood pressure
 - Weight
- Bloods tests
 - Kidney function
 - Salt levels
 - Blood dilution levels

What does taking part involve

• Urine Tests







24 Hour Collection

Spot Collection Urine Dipstick Test

Fluid intake = $\frac{(uOsm)(uVol)}{Target uOsm}$ + Insensible Loss

Urine Specific Gravity



Urine Specific Gravity ≤1.010



Smartphone Apps - DRINK





Smartphone Apps- SPLASH







DRINK Trial - Schedule



What information will it give us?

- Is high water intake safe
- Can each of the groups stick to their required water intake and achieve urine targets
- Can we show that there is a significant difference between both groups
- Can participants use smartphone applications to record some of the data we need (urine dip and fluid intake)

What will be the next step?

- Large
- Long-term (3-5 years)
- Randomised
- Trial
- High versus normal water intake
- Outcomes: kidney and cyst size, kidney function



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Pain in PKD – future Research Studies





What is the problem?

Common

• Significantly impacts QoL

Not well understood

• Not well managed

Data From DRINK Trial

 Novel pain questionnaire – Pain Specialist (Dr Lee).

- Information
 - Nature/Characteristics
 - Site
 - Frequency
 - Patterns: PKD and non-PKD related

Pain Symposium

- When?
 - May 2017
- Where?
 - Cambridge
- Who is invited?
 - Pain specialists: PKD and general
 - PKD Patients
 - Specialists in clinical trials and research
 - Pain Psychologists

Pain Symposium

- What do we hope to achieve?
 - What we know now about PKD pain so far
 - What interventions are possible and would they be acceptable
 - How to design a pain trial
 - What measures of pain should we use
 - What trial outcomes/endpoints should we choose

-Willingness to participate in pain studies

If you would like any extra information about the research trials or would be interested in participating:

Dr Ragada El-Damanawi or Dr Thomas Hiemstra <u>Email:</u> add-tr.drinktrial@nhs.net <u>Website:</u>

https://cctu.medschl.cam.ac.uk/Trials/Drink/Dri nkTrial.htm

Acknowledgments

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Questions?