Volumetric changes in the hippocampus and relationship to memory indices

in children with Hyperinsulinaemic hypoglycaemia and Ketotic hypoglycaemia

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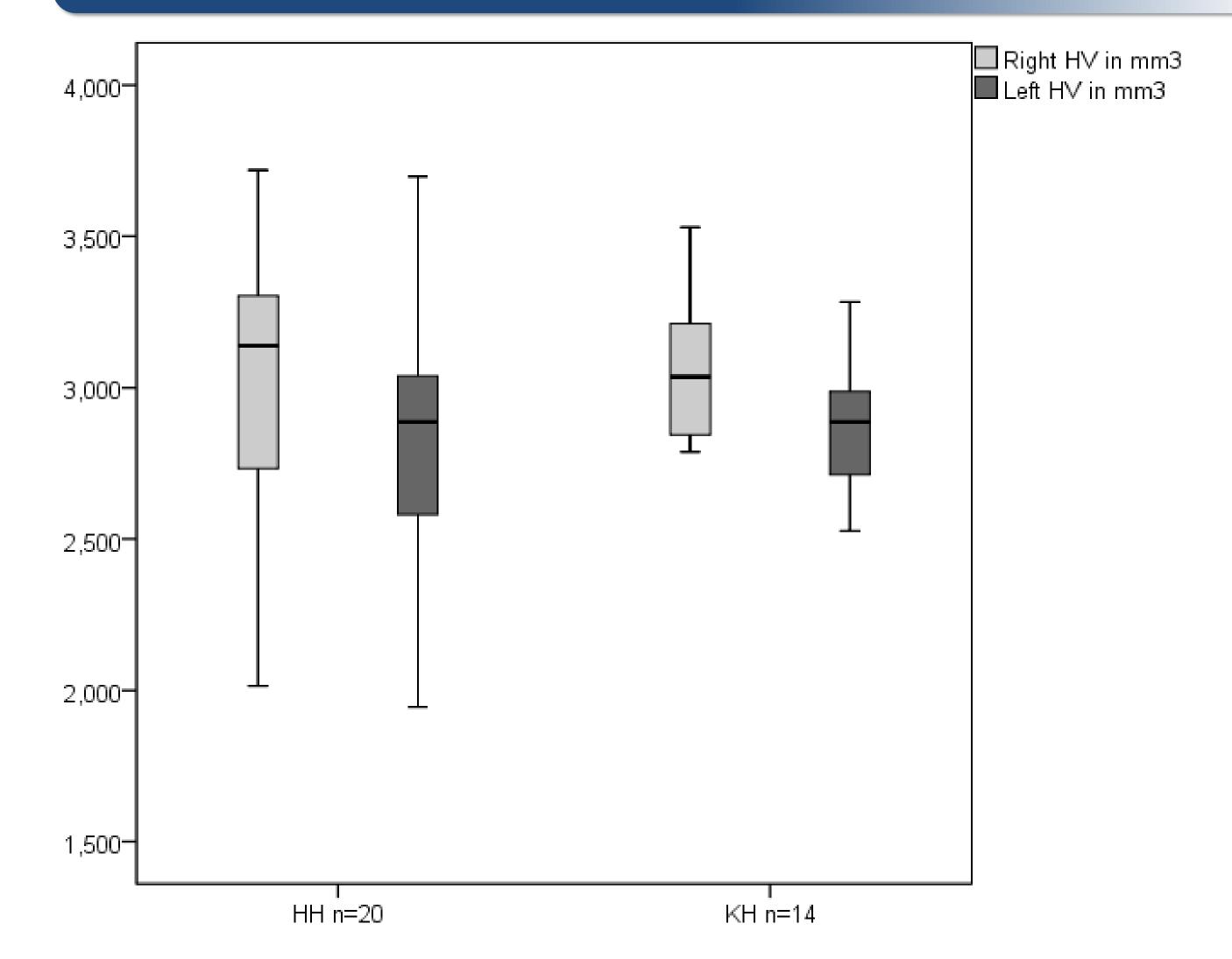
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Background and Objective

Children with hyperinsulinaemic hypoglycaemia (HH) are at a high risk of brain injury, while children with ketotic hypoglycaemia (KH) are believed to be neurologically normal. Hippocampus is known to be susceptible to hypoglycaemia, and is one of the key structures in the memory system. Our objective was to ascertain if children with HH sustain greater hippocampal injury and memory deficits in comparison to children with KH.

Comparison of Hippocampal volumes (HV) between children with HH and KH





Great Ormond Street NHS Hospital for Children **NHS Trust**

Methods

20 neurologically normal children between 5-16 years of age with HH and 14 children with KH were recruited from the endocrine and metabolic outpatient clinics from 2009-2012. Cognitive assessment was performed using Wechsler Intelligence Scale for Children Fourth edition and Children Memory Scale. Conventional (T1, T2 weighted) MRI for visual inspection of hippocampus and fast low angle shot (FLASH) three-dimensional MRI for manual hippocampal volume measurement were acquired. The FLASH data was analyzed with MEDx, version 3.3 (Sensor Systems Inc., VA). Each hippocampal slice, 1 mm in thickness, was manually traced.

On conventional MRI, small hippocampi were seen in 28.5% (n=6) with HH and 7% (n=1) with KH

| Hippocampus | HH | KH | (KH vs HH) |
|-----------------|---------|---------|-----------------|
| Volume | (n=20) | (n=14) | <i>p-</i> value |
| Right (median) | | | |
| mm ³ | 3138.36 | 3035.19 | 0.959 |
| Left (median) | | | |

Results

Baseline characteristics of the patients are as follows

| Patient characteristics | HH | KH | p-value |
|-------------------------------|-------------|-------------|---------|
| | | | |
| Number of patients | 20 | 14 | |
| Age at scan in years | 10.3 | 8.7 | |
| (mean ± SEM) | (2.6) | (2.1) | 0.073 |
| Gender (Male/female) | 13M/7F | 9M/5F | 0.96 |
| Gestational age in weeks | 39.3 | 38.7 | 0.42 |
| Birth weight in grams (mean ± | 3870.5 | 3337.2 | |
| SEM) | (898.7) | (747.4) | 0.069 |
| Handedness (Right/Left) | 15R/5L | 11R/3L | 1.000 |
| Height sds (median) | -0.06 (0.8) | -0.05 (0.6) | 0.703 |
| Weight sds (median) | 0.09 | -0.17 | 0.287 |
| Seizures at diagnosis (n) | 9 | 3 | 0.27 |

Memory indices and IQ

Comparison of mean scores, Children Memory scale (CMS) and Intelligence Quotient (IQ) between HH and KH, are as follows

| CMS Index scores | нн | KH | <i>p</i> value | | |
|---|------|-------|----------------|--|--|
| Visual Immediate | 94.8 | 106.3 | 0.022 | | |
| Visual Delayed | 97.5 | 104.5 | 0.093 | | |
| Verbal Immediate | 94.9 | 110.5 | 0.006 | | |
| Verbal Delayed | 97.2 | 108.7 | 0.055 | | |
| General Memory | 93.1 | 110 | 0.002 | | |
| Learning index score | 91.3 | 107 | 0.005 | | |
| Full scale IQ | 89.3 | 100.5 | 0.028 | | |
| Hippocampal volumes did not correlate to memory | | | | | |

indices in HH group. 3 children in the HH group had > 20-25% reduction in bilateral hippocampal volumes, however only one child had impaired memory.

Conclusions

Children with HH manifest significant impairment of memory, however they do not correlate with hippocampal volumes. Further studies are required to determine the neural substrate underlying these memory impairments.

References

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