A rare case of neonatal hypocalciuric hypercalcemia complicated by cardiac arrhythmia

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Introduction
Familial hypocalciuric hypercalcemia is a rare, lifelong, and usually benign hereditary disorder which is usually asymptomatic. Here we present a rare case of neonatal hypocalciuric hypercalcemia complicated by cardiac arrhythmia.

Patients

History of presenting illness
A healthy male infant weighing 2636g was delivered by spontaneous vaginal delivery at term. The pregnancy had progressed normally, but neonatal arrhythmia was found by physical examination after birth. Electrocardiography demonstrated isolated premature atrial ectopics. The baby was otherwise well and was discharged on day 6 with outpatient follow-up. On day 10, he represented with fever and hypercalcemia was coincidently discovered. Further investigation was subsequently carried out. There was no family history of hypercalcemia.

Physical Examination
The only abnormal finding was an irregular pulse. BW:2.6 kg. BT 37.2, HR 80-140 bpm, RR 30, SpO2 98%(RA)

Imaging study

ECG
Premature atrial contractions (PAC) with block
QTc (Bazett) 0.42 s (normal range; 0.35-0.44 s)

Laboratory data

<table>
<thead>
<tr>
<th>Blood test</th>
<th>Serum measurement</th>
<th>Normal range</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ca</td>
<td>3.06 mmol/L</td>
<td>2.10-2.50 mmol/L</td>
</tr>
<tr>
<td>iCa</td>
<td>1.64 mmol/L</td>
<td>1.22-1.32 mmol/L</td>
</tr>
<tr>
<td>P</td>
<td>1.39 mmol/dL</td>
<td>0.70-1.40 mmol/L</td>
</tr>
<tr>
<td>ALP</td>
<td>1868 U/L</td>
<td>500-1600 U/L</td>
</tr>
<tr>
<td>PTH-intact</td>
<td>89 pg/mL</td>
<td>10-65 pg/mL</td>
</tr>
<tr>
<td>25-OH VitD</td>
<td>&lt;4 ng/mL</td>
<td>10-30 ng/mL</td>
</tr>
</tbody>
</table>

Urine test
24-hour Fraction excretion of Ca 0.97 %
(Diagnostic criteria for hypocalciuric hypercalcemia is less than 1%)

Clinical course

ECG following Vit D supplementation

Course of laboratory data

<table>
<thead>
<tr>
<th></th>
<th>Day 10</th>
<th>Day 30</th>
<th>Day 60</th>
<th>Day 120</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ca</td>
<td>3.06</td>
<td>2.91</td>
<td>3.17</td>
<td>3.24</td>
</tr>
<tr>
<td>ALP</td>
<td>1863</td>
<td>2852</td>
<td>2285</td>
<td>1496</td>
</tr>
<tr>
<td>PTH-intact</td>
<td>89</td>
<td>85</td>
<td>53</td>
<td>32</td>
</tr>
</tbody>
</table>

- He was diagnosed with hypocalciuric hypercalcemia on the basis of clinical and laboratory findings.
- The arrhythmia did not worsen until day 30 and the total calcium level remained stable.
- Vitamin D supplementation was prescribed on day 30 because he suffered from vitamin D deficiency (persistent high levels of ALP and PTH), which was also diagnosed with her mother after his admission.
- The arrhythmia gradually disappeared corresponding to decline in ALP and PTH levels following treatment.

Discussion
- Hypocalciuric hypercalcemia itself rarely causes arrhythmia without shortening of QT interval. However, high levels of PTH concomitant with vitamin D deficiency could induce premature atrial contraction as seen in this case.
- PTH is reported to exert a direct action on cardiac myocytes and influence cardiac contractility linked to arrhythmia. Furthermore, previous literatures showed hypercalcemia in patients with primary hyperparathyroidism increased the occurrence of arrhythmia such as premature atrial contraction, which often disappeared following parathyroidectomy.
- Therefore, it is possible that normalization of PTH levels by vitamin D blunted susceptibility to arrhythmia.

Conclusion
- Clinicians should consider electrolyte abnormalities including parathyroid hormone which regulates calcium homeostasis in the differential diagnosis of neonatal arrhythmia.