SAFDGS data. Genotypic data of rs7493 and rs12704795 were polymorphic and verified for Mendelian inconsistencies and allele and genotype frequencies were measured using the programme PEDSYS subroutines. The allele frequencies of rs7493 were 77% (C) and 23% (G). In regard to the rs12704795, the A and C allele frequencies were 76% and 24%, respectively. Genotypic data of rs7493 [CC (60%), CG (35%), GG (5%)] and rs12704795 [AA (55%), AC (39%), CC (6%)] were consistent with the Hardy–Weinberg Equilibrium expectations, and there was no evidence for hidden population stratification. Association analysis in our family data was carried out using the measured genotype approach within the variance components analytical framework implemented in SOLAR [5].

Of the phenotypes examined for association [T2DM, body mass index (BMI), blood pressure measures, total cholesterol, high density lipoprotein-cholesterol, triglycerides and ln ACR], the C/G variant (rs7493) exhibited significant association only with ACR ($P = 0.013$) after adjusting for the effects of age, age$^2$ x sex, diabetes, duration of diabetes, systolic blood pressure, and antihypertensive treatment with ACE inhibitors or AT1R antagonists. The mean ACR values in the two groups, the genetic influences attributable to this polymorphism and other covariates, and the association findings were found to be similar to ACR ($P = 0.019$). The analyses also indicate that the rs7493 variant explained ~8% of the total genetic variance in ACR. The biological relevance of our findings remains to be identified. Alterations in the activity of PON2 due to rs7493 are likely to influence the composition as well as the oxidation of lipoproteins [1]. For example, oxidized LDL may induce oxidative damage in kidney cells, therefore contributing to the pathogenesis of albuminuria. The precise biological relevance of this variant to albuminuria will therefore depend on its effect on the enzymatic activity of PON2. Our association analysis failed to show significant association between the phenotypes examined and genotypic data of rs12704795. The absence of relationships between the examined variants and the serum lipids and lipoproteins, BMI and blood pressure measures, is consistent with a previous report [1]. This study has limitation in that it focused on three variants in PON2 gene examined previously and has not attempted comprehensive tagging of all common variations within PON2 that could influence variation in ACR. In addition, it is possible that the associated variant is in linkage disequilibrium with (a) potential functional variant(s) in PON2 or with other variants in the flanking genes. These possibilities remain to be explored.

In conclusion, our data support the findings by others [1,2] that the Cys(311)Ser variant of PON2 may contribute to albumin excretion rate. Although the susceptibility to albuminuria due to this variant is found in different ethnic groups, the genetic influences attributable to this polymorphism appear to be rather minor in their magnitude.

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**Posterior reversible encephalopathy induced by intravenous immunoglobulin**

Sir, Intravenous immunoglobulin (IvIg) is commonly used in nephrology units for the treatment of auto-immune diseases, antibody-mediated renal allograft rejection and immune deficiencies. Minor adverse effects, such as myalgia, headache, shiver, nausea, vomiting or fever occur in <20% of patients. Major reactions including renal failure, thromboembolism, aseptic meningitis and anaphylaxis are less common. Whereas osmotic nephropathy resulting from maltose and saccharose toxicity is well-known to nephrologists, posterior reversible encephalopathy syndrome (PRES) is a rare and potentially severe adverse event of IvIg therapy.

A 42-year-old man was admitted for end-stage renal failure secondary to myeloma cast nephropathy in the context of plasma cell leukaemia, diagnosed 2 months previously. His past medical history was otherwise unremarkable. Haematological tests showed increased white blood cells (12380/mm$^3$) with 27% of circulating plasma...
of 450 mg/l revealed diffuse osteolytic bone lesions. Serum creatinine was massive plasma cell infiltration (67%). X-ray examination revealed diffuse osteolytic bone lesions. Serum creatinine was 450 μmol/l with severe hyperkalaemia requiring periodic haemodialysis. Vincristine, adriamycin and dexamethasone therapy was initiated. One month after the first course of chemotherapy, circulating plasma cells were no longer detected, while bone marrow plasmacytosis (10%) and serum-free light chains of 9760 mg/l. Gammaglobulin level was 13 g/l. Bone marrow smears and biopsy demonstrated massive plasma cell infiltration (67%). X-ray examination revealed diffuse osteolytic bone lesions. Serum creatinine was 450 μmol/l with severe hyperkalaemia requiring periodic haemodialysis. Vincristine, adriamycin and dexamethasone therapy was initiated. 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At the onset of neurological symptoms, laboratory data showed: glucose 1 g/l, sodium 140 mmol/l, potassium 3.5 mmol/l, serum creatinine 450 μmol/l, total protein 66 g/l, albumin 28 g/l, calcium 2.2 mmol/l, phosphorus 1.53 mmol/l, CRP 7 mg/l, white blood cell 14 500/mm3, haemoglobin 9.7 g/dl, haematocrit 30% and platelet 379 000/mm3. Fundoscopic examination was normal. Brain MRI revealed increased signal in the white matter of left parieto-occipital lobe on T2 weighted sequences. Unfortunately, lumbar puncture and plasma viscosity was not performed. After 24 h, complete resolution of neurological symptoms was achieved.

In summary, PRES should be suspected in patients with neurological signs, especially visual disturbance after IV Ig infusion and adequately confirmed by MRI. Nephrologists need to be aware of this unusual neurological complication, as early recognition may improve prognosis.

Fig. 1. Brain MRI showing reversible unilateral left parieto-occipital white matter hypersignal on T2 weighted sequences.
A supernumerary parathyroid gland located in an unusual site, parapharyngeal space, in a patient with persistent renal hyperparathyroidism

Sir,

Sometimes it is difficult to detect pathological parathyroid glands both in primary and secondary hyperparathyroidism (HPT). We are reporting a case of persistent renal HPT due to a supernumerary gland located in the parapharyngeal space.

The patient was a 58-year-old female, who had continued haemodialysis (HD) for 25 years and referred to our department due to advanced renal HPT. She underwent total PTx with forearm autograft and four parathyroid glands could be removed. However, a high intact parathyroid hormone (PTH) level was persistent on the first day after PTx. To detect missed glands, image diagnostic examinations, including MIBI scitigram, ultrasonography (US), computed tomography (CT) and magnetic resonance imaging (MRI) were performed several times. Finally, we recognized an abnormal uptake of MIBI scintigram in her face, and we detected a mass located in the parapharyngeal space by CT (Figure 1). The mass was removed by an operation from the oral cavity. Histopathological examination confirmed that the mass represented a parathyroid gland. The high intact PTH level decreased dramatically. It is suggested that the gland may be an undescended parathyroid gland.

This case may be the first to report a parathyroid gland located high up in the neck at the level of the parapharyngeal space. In 1995, Simeone et al. [1] reported a case with an undescended parathyroid gland located within or in close approximation to the parapharyngeal wall, but the parapharyngeal space has a higher position in the neck than that area. The inferior parathyroid glands arise from the third pharyngeal pouch along with the thymus. During the course of normal development, the inferior parathyroid glands descend through the neck with the thymus. The loss of expression of some genes that induce descending of the parathyroid glands may be a cause why the gland remained in the parapharyngeal space. It has been confirmed that a gene, Hoxa3, is involved in the descent of the parathyroid glands [2].

An undescended parathyroid gland is one of many pitfalls that may cause the missing of an abnormal parathyroid glands. Undescended parathyroid glands are most commonly found within the carotid sheath at the level of the carotid bifurcation [3]. In our large series, the frequency of undescended glands was 0.97%, and half of them could be removed at reneck exploration [4]. In 1998, we reported that the frequency of supernumerary glands in renal HPT was 92 of 570 glands (16.1%) and of these 92 glands, 12 were removed at reoperation [5].

The risk of persistent/recurrent HPT due to remaining parathyroid tissue is not negligible, especially in patients who require long-term maintenance HD after PTx. Therefore, supernumerary glands have special significance during surgery for renal HPT, either as a cause for persistent or recurrent HPT.

We conclude that one should keep in mind that parathyroid glands can be located in the parapharyngeal space and that one should look for this possibility, when pathological glands in the usual sites are not detected.