

Antiphospholipid syndrome-associated nephropathy in systemic lupus erythematosus

A letter in response to: Tektonidou MG. Antiphospholipid syndrome-associated nephropathy in systemic lupus erythematosus. *Int. J. Clin. Rheumatol.* 7(2), 131–134 (2012).

The article published by Maria G Tektonidou entitled 'Antiphospholipid syndrome-associated nephropathy in systemic lupus erythematosus' in the esteemed journal of *International Journal of Clinical Rheumatology* had some points that need further explanation [1].

In this article, Tektonidou has explained the morphologic lesions of antiphospholipid syndrome-associated nephropathy (APS-nephropathy). She also emphasizes that the renal pathologists should pay particular attention to the histologic lesions of APS-nephropathy during the examination of renal biopsies of systemic lupus erythematosus patients, especially those with positive antiphospholipid antibodies [1].

Tektonidou also fully explained morphologic lesions of APS-nephropathy [1]. In recent years, much progress has been made toward understanding vascular lesions in lupus nephropathy. Recently, Wu *et al.* conducted a study on 341 patients with lupus nephritis, and found that 279 were diagnosed with single or multiple renal vascular lesions that included 253 with vascular immune complex deposits, 82 with atherosclerosis, 60 with thrombotic microangiopathy, 13 with noninflammatory necrotizing vasculopathy and two with true renal vasculitis [2]. In this study, they suggested the inclusion of renal vascular lesions to the 2003 International Society of Nephrology/Renal Pathology Society system of lupus nephritis classification to improve renal outcome predictions [2]. However, vascular lesions in lupus nephritis have different etiologies and may result from lupus-related vasculopathy due to immune complex deposition in the vessel wall, vasculitis owing to rare association of ANCA-associated vasculitis with lupus and, finally, the most important is APS-nephropathy [2–6]. Thus, it is impossible to include the vascular lesions in the International Society of Nephrology/Renal Pathology Society 2003 lupus

nephropathy classification, while combining different vascular disorders into a general category of vasculopathy as this is incorrect. It is evident that the main etiologic factor of vascular lesions in systemic lupus erythematosus belongs to APS-nephropathy, which is also known as vaso-occlusive nephropathy [7–10], and it is well known that morphologic lesions of APS-nephropathy aggravate lupus nephropathy [11–15].

In fact, it seems judicious to suggest a distinct classification for APS-nephropathy, to emphasize more consideration of this disease and to avoid under-recognition of this nephropathy. This classification may be used together with the International Society of Nephrology/Renal Pathology Society 2003 of lupus nephropathy in the same report.

However, the main question is which morphologic lesions of APS-nephropathy have prognostic value and should, therefore, be included in this classification?

In answer to this, vascular lesions should be categorized into acute (thrombotic microangiopathy) and chronic (fibrose intimal hyperplasia and thrombus), glomerular lesions (glomerular ballooning) and tubule interstitial involvement (focal cortical atrophy and tubular thyroidization) [1,12,16,17].

In this regard, I draw attention to gathering pathologic lesions of this syndrome into a classification, such as Oxford classification for IgA-nephropathy for the below mentioned reasons [18–20].

First, in contrast to the morphologic lesions of lupus nephritis, which is usually additive, pathologic features of APS-nephropathy are not proliferative. In fact, in lupus nephropathy, pathologic lesions may evolve from class I to II, III, IV and, in the case of failure of treatment, class VI lupus nephritis will ensue [2,8,10]. However, in APS-nephropathy, pathologic damage



Hamid Nasri

Department of Nephrology, Division of Nephropathology, Isfahan University of Medical Sciences, Isfahan, Iran
hamidnasri@med.mui.ac.ir

is a vaso-occlusive disease, affects glomeruli, vessels and tubule interstitial.

Second, a suggested classification for APS-nephropathy, should be simple and practical, and resemble the Oxford classification [18,19].

However, the suggestion of a new classification for APS-nephropathy will involve a tremendous amount of work and will require a working group; thus, more studies on this topic is suggested.

Financial & competing interests disclosure

The author has no relevant affiliations or financial involvement with any organization or entity with a financial interest in or financial conflict with the subject matter or materials discussed in the manuscript. This includes employment, consultancies, honoraria, stock ownership or options, expert testimony, grants or patents received or pending, or royalties.

No writing assistance was utilized in the production of this manuscript.

References

- 1 Tektonidou MG. Antiphospholipid syndrome-associated nephropathy in systemic lupus erythematosus. *Int. J. Clin. Rheumatol.* 7(2), 131–134 (2012).
- 2 Wu LH, Yu F, Tan Y *et al.* Inclusion of renal vascular lesions in the 2003 ISN/RPS system for classifying lupus nephritis improves renal outcome predictions. *Kidney Int.* 83(4), 715–723 (2013).
- 3 Nasri H. Hypertension and renal failure with right arm pulse weakness in a 65 years old man. *J. Nephropathol.* 1(3), 130–133 (2012).
- 4 Mubarak M. Catastrophic antiphospholipid syndrome presenting with sudden renal failure: the lesson lies in vascular lesions. *J. Nephropathol.* 2(2), 135–138 (2013).
- 5 Daugas E, Nochy D, Huong DL *et al.* Antiphospholipid syndrome nephropathy in systemic lupus erythematosus. *J. Am. Soc. Nephrol.* 13(1), 42–52 (2002).
- 6 Mardani S, Nasri H. Catastrophic antiphospholipid syndrome presenting with sudden renal failure and past history of long-lasting psychosis and hypertension in a woman. *J. Nephropathol.* 2(2), 110–113 (2013).
- 7 Tektonidou MG, Sotsiou F, Moutsopoulos HM. Antiphospholipid syndrome (APS) nephropathy in catastrophic, primary, and systemic lupus erythematosus-related APS. *J. Rheumatol.* 35(10), 1983–1988 (2008).
- 8 Ali A, Al-Windawi S. Tubulointerstitial lupus nephritis. *J. Nephropathol.* 2(1), 75–80 (2013).
- 9 Tektonidou MG, Sotsiou F, Nakopoulou L, Vlachoyiannopoulos PG, Moutsopoulos HM. Antiphospholipid syndrome nephropathy in patients with systemic lupus erythematosus and antiphospholipid antibodies: prevalence, clinical associations, and long-term outcome. *Arthritis Rheum.* 50(8), 2569–2579 (2004).
- 10 Mubarak M. Hidden face of lupus nephritis exposed: isolated tubulointerstitial lupus nephritis. *J. Nephropathol.* 2(1), 71–72 (2013).
- 11 Asherson RA, Klumb EM. Antiphospholipid syndrome nephropathy in different scenarios. *J. Rheumatol.* 35(10), 1909–1911 (2008).
- 12 Hill GS, Nochy D. Antiphospholipid syndrome in systemic lupus erythematosus. *J. Am. Soc. Nephrol.* 18(9), 2461–2464 (2007).
- 13 Serrano F. Antiphospholipid syndrome: a complex disease. *J. Nephropathol.* 2(1), 73–74 (2013).
- 14 Anis S, Ahmed E, Muzaffar R. Prevalence of anti-beta2GPI antibodies and their isotypes in patients with renal diseases and clinical suspicion of antiphospholipid syndrome. *J. Nephropathol.* 2(3), 181–189 (2013).
- 15 Nishimura M, Nii T, Trimova G *et al.* The NF- κ B specific inhibitor DHMEQ prevents thrombus formation in a mouse model of antiphospholipid syndrome. *J. Nephropathol.* 2(2), 114–121 (2013).
- 16 Tektonidou MG. Renal involvement in the antiphospholipid syndrome (APS)-APS nephropathy. *Clin. Rev. Allergy. Immunol.* 36(2–3), 131–140 (2009).
- 17 Ardalan MR, Vahedi A. Antiphospholipid syndrome: a disease of protean face. *J. Nephropathol.* 2(1), 81–84 (2013).
- 18 Cattran DC, Coppo R, Cook HT *et al.* The Oxford classification of IgA nephropathy: rationale, clinicopathological correlations, and classification. *Kidney Int.* 76(5), 534–545 (2009).
- 19 Nasri H, Mortazavi M, Ghorbani A *et al.* Oxford-MEST classification in IgA nephropathy patients: a report from Iran. *J. Nephropathol.* 1(1), 31–42 (2012).
- 20 Nasri H. Antiphospholipid syndrome-associated nephropathy: current concepts. *J. Ren. Inj. Prev.* 2(1), 1–2 (2012).