too? What is the optimal dose of metformin for renoprotection?

CONFLICT OF INTEREST

None declared.

REFERENCES

- Nagi DK, Yudkin JS. Effects of metformin on insulin resistance, risk factors for cardiovascular disease, and plasminogen activator inhibitor in NIDDM subjects. A study of two ethnic groups. Diabetes Care. 1993;16:621-9.
- 2. Despres JP. Potential contribution of metformin to the management of cardiovascular disease risk in patients with abdominal obesity, the metabolic syndrome and type 2 diabetes. Diabetes Metab. 2003;29:6S53-61.
- Kurukulasuriya R, Banerji M, Chaiken R, Lebovitz H. Selective decrease in visceral fat is associated with weight loss during metformin treatment in African Americans with type 2 diabetes mellitus. Diabetes. 1999:48:A315.
- Standeven KF, Ariens RA, Whitaker P, Ashcroft AE, Weisel JW, Grant PJ. The effect of dimethylbiguanide on thrombin activity, FXIII activation, fibrin polymerization, and fibrin clot formation. Diabetes. 2002;51:189-97.
- Isoda K, Young JL, Zirlik A, et al. Metformin inhibits proinflammatory responses and nuclear factor-kappaB in human vascular wall cells. Arterioscler Thromb Vasc Biol. 2006;26:611-7.
- Bonnefont-Rousselot D, Raji B, Walrand S, et al. An intracellular modulation of free radical production could contribute to the beneficial effects of metformin towards oxidative stress. Metabolism. 2003;52:586-9.
- Martinez J, Moreno JJ. Role of Ca2+-independent phospholipase A2 on arachidonic acid release induced by reactive oxygen species. Arch Biochem Biophys. 2001;392:257-62.
- Morales AI, Detaille D, Prieto M, et al. Metformin prevents experimental gentamicin-induced nephropathy

by a mitochondria-dependent pathway. Kidney Int. 2010;77:861-9.

- Amini FG, Rafieian-Kopaei M, Nematbakhsh M, Baradaran A, Nasri H. Ameliorative effects of metformin on renal histologic and biochemical alterations of gentamicininduced renal toxicity in Wistar rats. J Res Med Sci. 2012;17:621-5.
- Rafieian-Kopaei M, Baradaran A, Merrikhi A, Nematbakhsh M, Madihi Y, Nasri H. Efficacy of Coadministration of Garlic Extract and Metformin for Prevention of Gentamicin-Renal Toxicity in Wistar Rats: A Biochemical Study. Int J Prev Med. 2013;4:258-64.
- Bruckbauer A, Zemel MB. Synergistic effects of metformin, resveratrol, and hydroxymethylbutyrate on insulin sensitivity. Diabetes Metab Syndr Obes. 2013;6:93-102.
- Kim J, Shon E, Kim CS, Kim JS. Renal podocyte injury in a rat model of type 2 diabetes is prevented by metformin. Exp Diabetes Res. 2012;2012:210821.
- Nasri H, Baradaran A, Ardalan MR, Mardani S, Momeni A, Rafieian-Kopaei M. Bright renoprotective properties of metformin: beyond blood glucose regulatory effects. Iran J Kidney Dis. 2013;7:423.
- Garber AJ, Duncan TG, Goodman AM, Mills DJ, Rohlf JL. Efficacy of metformin in type II diabetes: results of a double-blind, placebo-controlled, dose-response trial. Am J Med. 1997;103:491-7.
- Rocha A, Almeida M, Santos J, Carvalho A. Metformin in patients with chronic kidney disease: strengths and weaknesses. J Nephrol. 2013;26:55-60.

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Urine Examination in the Era of Modern Diagnostics

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Urine examination, dipstick, and urine microscopy, the fundamental, common, and inexpensive diagnostic methods in nephrology have been replaced relatively by more sophisticated procedures in many nephrology units and have become one of the laboratory personnel tasks and not nephrologists.¹ Tsai and colleagues, in their study of acute kidney failure, showed the increased chance of correct diagnosis from 19.2% to 69.3% when urinalysis report was done by a nephrologist.² Fogazzi and colleagues compared in their well-designed study the features of urine

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findings in proliferative and nonproliferative glomerular diseases and found higher frequencies of erythrocyte, leukocyte, tubular epithelial cell, and different casts in proliferative glomerular disease.³ Those findings provided 80.8% sensitivity and 79.2% specificity for clinical diagnosis.³ Such an approach may help to identify risk factors in different diseases such as Henoch-Schonlein nephritis in children to justify kidney biopsy indications and aggressive treatment.⁴ Interestingly, there is a case report of adenine phosphoribosyl transferase deficiency, diagnosed by urine examination and unusual crystals which drew nephrologists' attention in a patient with poor graft function.⁵

Furthermore, 16% prevalence of decoy cells in kidney transplant patients emphasizes the consideration of inexpensive urine examination, although decoy cells are not either sensitive or specific for BK virus infection.^{6,7}

Re-introduction of this test into clinical practice looks necessary from time to time. The importance of this test application seems more serious when other facilities for diagnosis is lacking, especially in many countries of developing world with restricted human and financial resource.^{8,9}

Many argue that the nephrologists are still needed to take their role in interpretation of what is seen, because they may know what to look for based on the clinical setting. Urine examination should be an integral part of nephrology educational curriculum. Hami and coworkers enthusiastically studied the attitude of Iranian nephrologists towards the subject.¹⁰ By a well-prepared questionnaire, under the supervision of an outstanding international urine examination expert, professor Fogazzi, renal centers were questioned about involvement of nephrologists in performing urine microscopy. They once again put emphasis on the necessity for clinical nephrologists to roll up their sleeves to take the responsibility of this task more because of the superiority of nephrologist-performed urinalysis to laboratory staff-performed urinalysis in finding out the diagnosis.

To achieve as much as possible data from this diagnostic tool all aspects of it should be taken into account. Proper cleansing procedures, sample collection, handling, slide preparation, equipment, and knowledgeable interpreter are all important.¹¹

Thanks to professor Fogazzi, the highly regarded expert in the field, the second workshop was held in Isfahan, Iran in 2012 and most mentioned items were reviewed comprehensively.

CONFLICT OF INTEREST

None declared.

REFERENCES

- Verdesca S, Brambilla C, Garigali G, Croci MD, Messa P, Fogazzi GB. How a skillful (correction of skilful) and motivated urinary sediment examination can save the kidneys. Nephrol Dial Transplant. 2007;22:1778-81.
- Tsai JJ, Yeun JY, Kumar VA, Don BR. Comparison and interpretation of urinalysis performed by a nephrologist versus a hospital-based clinical laboratory. Am J Kidney Dis. 2005;46:820-9.
- Fogazzi GB, Saglimbeni L, Banfi G, et al. Urinary sediment features in proliferative and non-proliferative glomerular diseases. J Nephrol. 2005;18:703-10.
- Assadi F. Childhood Henoch-Schonlein nephritis: a multivariate analysis of clinical features and renal morphology at disease onset. Iran J Kidney Dis. 2009;3:17-21.
- Stratta P, Fogazzi GB, Canavese C, et al. Decreased kidney function and crystal deposition in the tubules after kidney transplant. Am J Kidney Dis. 2010;56:585-90.
- Kroth LV, Henkin CS, Peres LD, et al. Prevalence of urinary decoy cells and associated risk factors in a Brazilian kidney, pancreas, and kidney-pancreas transplant population. Transplant Proc. 2012;44:2394-6.
- Yassari F, Parvin M, Ahmadpoor P. Nephroquiz 4: a 43-year-old woman with kidney allograft dysfunction due to BK virus nephropathy. Iran J Kidney Dis. 2010;4:164-7.
- Fogazzi GB, Attolou V, Kadiri S, Fenili D, Priuli F. A nephrological program in Benin and Togo (West Africa). Kidney Int Suppl. 2003;:S56-60.
- 9. Fogazzi GB, Bosan IB, Garigali G, Arogundade F. A theoretical and practical course on urine microscopy in Nigeria: a model for nephrological skill transfer. Nephron Clin Pract. 2011;117:c398-402.
- Hami M, Shahidi S, Nouri-Majalan N, Atapour A, Fogazzi GB. A workshop on urinalysis and a survey on urine microscopy among kidney centres of Iran. Iran J Kidney Dis. 2013;7:432.
- Fogazzi GB, Grignani S, Colucci P. Urinary microscopy as seen by nephrologists. Clin Chem Lab Med. 1998;36:919-24.

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