

Urinary pH as a target in the management of lithiasic patients in real-world practice: monitoring and nutraceutical intervention for non-lithogenic pH range

EULIS19
5th Meeting of the
EAU Section of Urolithiasis
3-5 October 2019, Milan, Italy

Juan Antonio Galán¹, Prevent-LitGroup^a, Jordi Cuñé²

1- Hospital General Universitario de Alicante, Alicante, Spain; 2- Medical Department, Devicare S.L., Cerdanyola del Vallès, Spain

^a Prevent-Lit group:

1. Juan Antonio Galán Llopis, Hospital General Universitario de Alicante, Alicante; 2. Montserrat Arzo Fàbregas, Hospital Germans Trias i Pujol, Badalona; 3. Jose María Banús Gassol, ICUN, Barcelona; 4. Joan María Benejam Gual, Hospital de Manacor, Islas Baleares; 5. Alberto Budía Alba, Hospital La Fe, València; 6. Enrique Cao Avellaneda, Hospital de Santa Lucía de Cartagena, Murcia; 7. Antonio Conte Visús, Policlínica Miramar, Palma de Mallorca; 8. Isabel Díaz Sánchez, Instituto Médico Tecnológico, Barcelona; 9. Enrique Argüelles Salido, Hospital Universitario Virgen del Rocío, Sevilla; 10. Juan Alberto Lancina Martín, Hospital Juan Canalejo, A Coruña; 11. María Pilar Luque Gálvez, Hospital Clinic de Barcelona, Barcelona; 12. Félix Millán Rodríguez, Fundació Puigvert, Barcelona; 13. José Francisco Morera Martínez, Hospital Universitario Dr. Peset, Valencia; 14. Paola Pardo, Hospital Universitario Dr. Peset, Valencia; 15. Manuel Carlos Reina Ruiz, Hospital Universitario Valme, Sevilla; 16. Miguel Ángel Rodríguez Cabello, Hospital Sanitas, Madrid; 17. Carlos Torrecilla-Ortiz, Hospital Universitario de Bellvitge, L'Hospitalet de Llobregat.

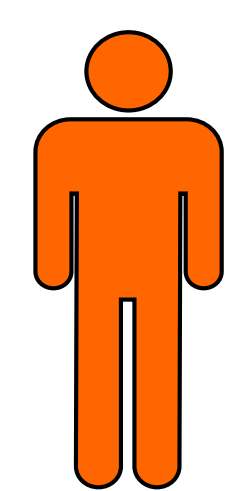
INTRODUCTION AND OBJECTIVE

pH influence on the formation and growth of certain type of stones is well-known and established. Low urine pH values promote uric acid (UA) stones, and those in the higher range do with calcium oxalate (CO) / calcium phosphate (CP) ones. Urine pH normalization helps preventing stone formation.

The objective of the study is to assess the effectiveness of the joint use of a pH meter in combination with nutraceuticals in restoring the urinary pH balance of patients with medical history of UA or CP/CO stones in real-world practice.

METHODS

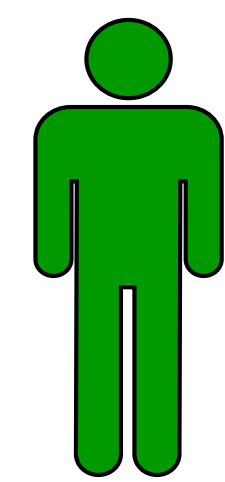
Interventional, prospective, open-label study, with 2 arms, in 143 lithiasic patients.



N= 78 (45,5%)
Acidifier group
pH at baseline > 6,2
history of **calcic lithiasis**



1 capsule/12hours
Lit-Control®-pH Down



N= 65 (54,5%)
Alkalinizer group
pH at baseline < 5,5
history of **uric lithiasis**



1 capsule/12h
Lit-Control®-pH Up

acidifier Lit-Control® pH Down		
substance	amount/capsule	mechanism
L-methionine	500 mg	decreases urinary pH
Phytate	170 mg	inhibits crystallization of CaOx & CaPO ₄

alkalinizer Lit-Control® pH UP		
substance	amount/capsule	mechanism
Magnesium-Potassium-Citrate	400 mg	calcium oxalate inhibitors increase urinary pH inhibit crystallization
Theobromine	150 mg	inhibit crystallization

Visits

- Baseline
- 30 days
- 60 days
- 90 days

Parameters

- Urinary pH
- Type of therapy
- Compliance
- Self-reported renal colic events

RESULTS

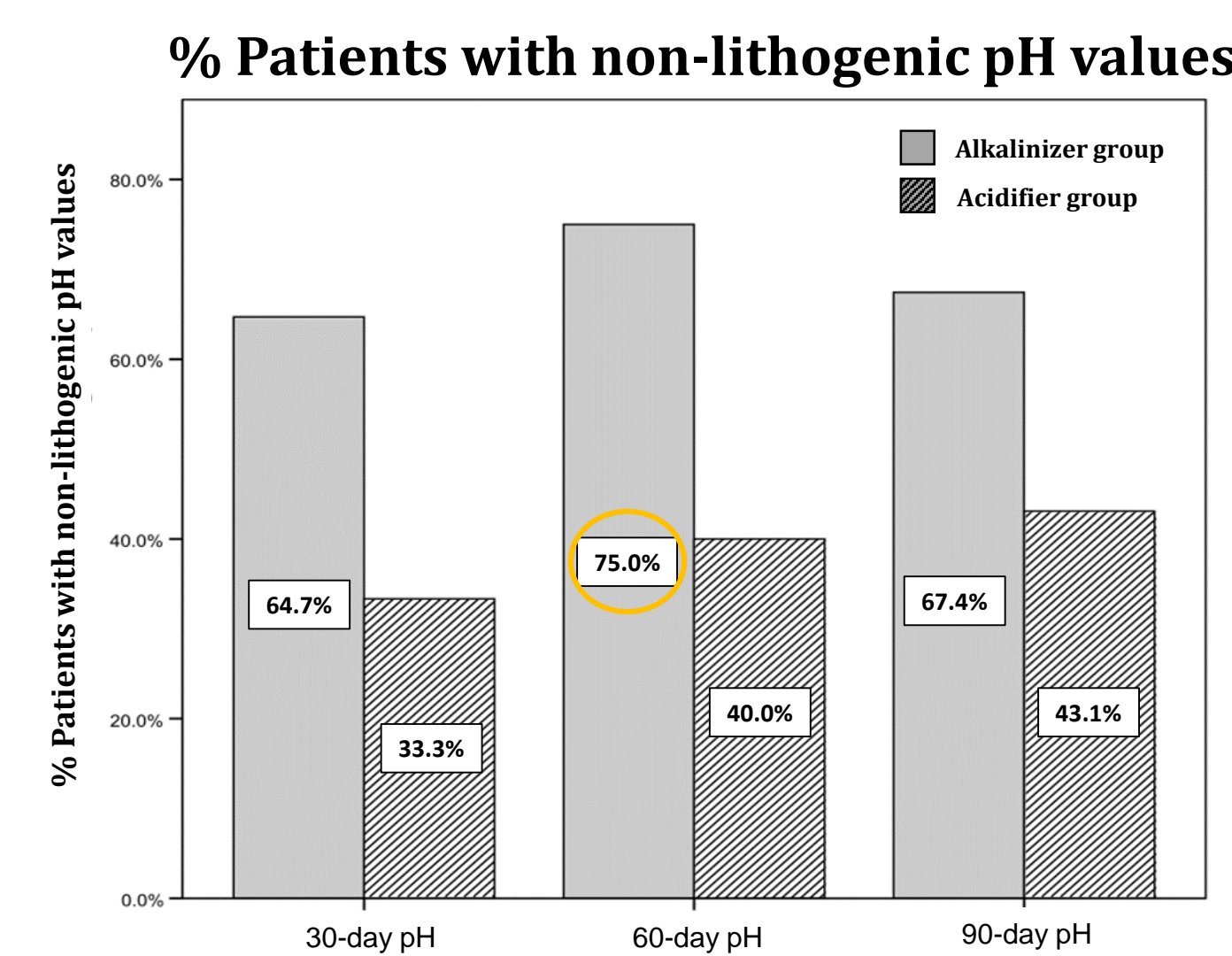
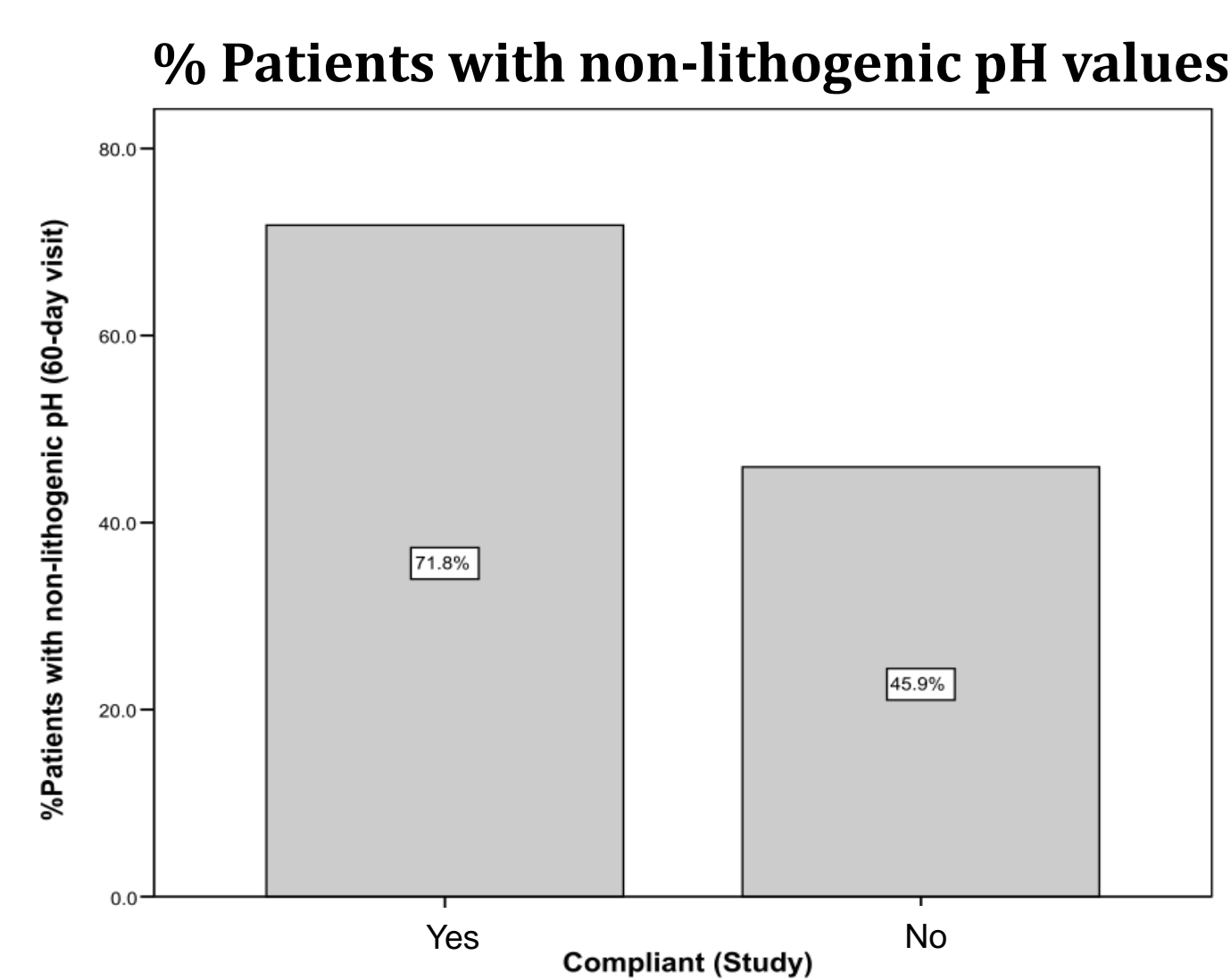
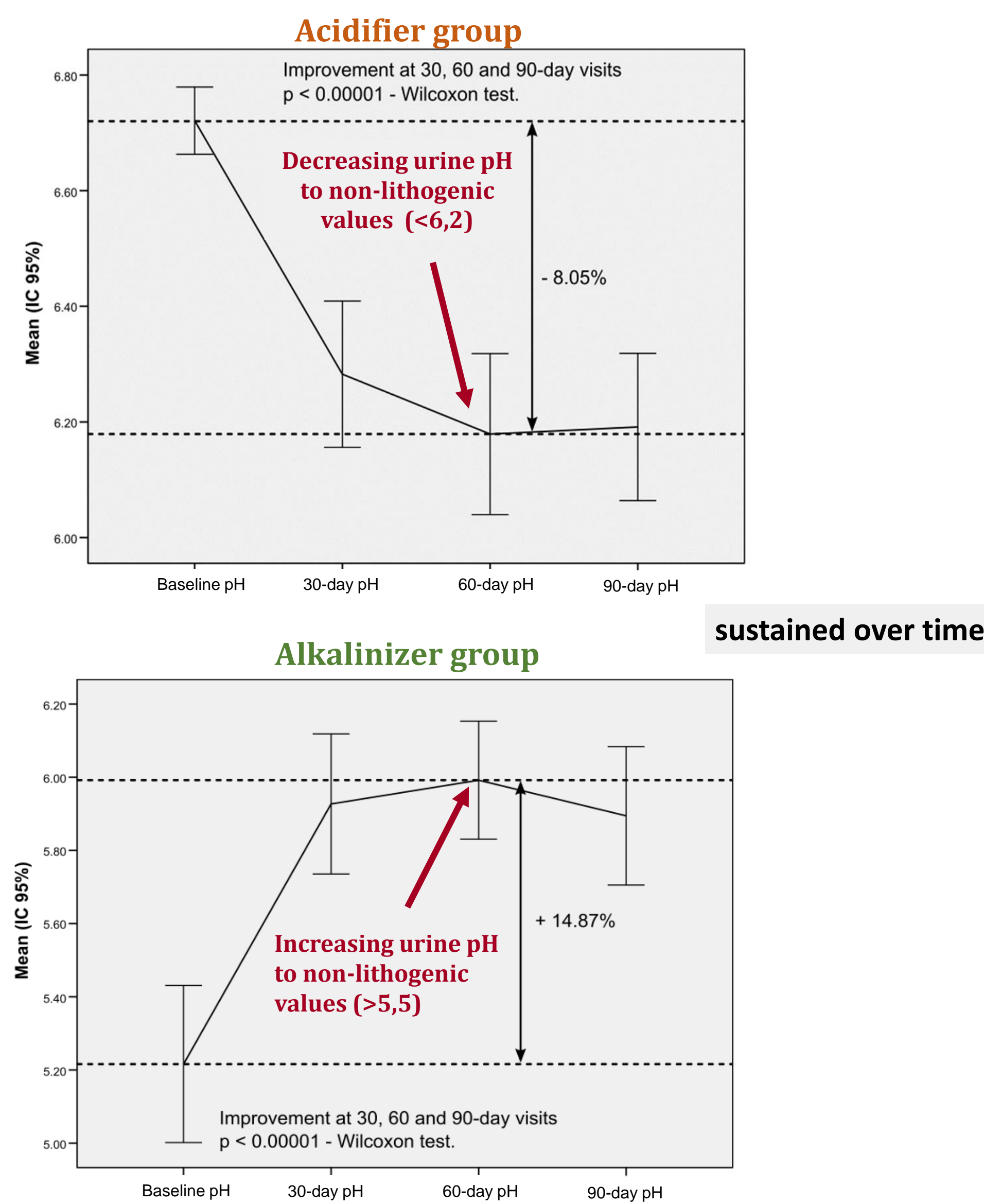


Figure 2. Percentage of patients with non-lithogenic pH values depending on whether they are compliant with dietary supplements .

- 75% of patients in the alkalinizer group reach **non-lithogenic urine pH values in 60 days of treatment**
- The non-lithogenic pH at 90 days (HR: 0.428, 95% Confidence interval [CI]: 0.193-0.947) and compliance at 60 days (HR: 0.428, 95% CI: 0.189-0.972) were independently associated with colic complaints-free survival

Figure 1. Mean of urinary pH at baseline and after 30, 60 and 90 days of dietary supplements intake.

CONCLUSION

In patients with medical history of lithiasis, monitoring of pH in combination with the product may be useful in maintaining non-lithogenic pH values, yielding very high ratios of success specially in compliant patients. Besides this main outcome, a reduction in self-reported colic complaints associated to pH balance was also observed.

The nutraceuticals were shown to be safe and well tolerated (in more than 90% of the cases, the investigators reported a good or very good tolerance), registering 8.84% of adverse events (13/147 patients), none of them serious.

BIBLIOGRAPHY

1. Türk C, et al. Guidelines on Urolithiasis. European Association of Urology 2016. Update 2018 <https://uroweb.org/guideline/urolithiasis/#4> 2. Grases F, et al. Effects of phytic acid on renal stone formation in rats. Scand J Urol Nephrol (1998) 32: 261-265. 3. Grases F, et al. Studies on calcium oxalate monohydrate crystallization: influence of inhibitors. Urol Res (1994) 22: 39-43. 4. Schlemmer U, et al. Phytate in foods and significance for humans: Food sources, intake, processing, bioavailability, protective role and analysis (Review). Mol. Nutr. Food Res. 2009; 53: S330-S375. 5. Saw NK, et al. Effects of inositol hexaphosphate (phytate) on calcium binding, calcium oxalate crystallization and in vitro Stone growth. J Urol (2007) Jun; 177(6): 2366-70. 6. Grases F, et al. Phytotherapy and renal stones: the role of antioxidants. A pilot study in wistar rats. Urol Res (2009) 37:35-40. 7. Grases F, et al. Effects of polyphenols from grape seeds on renal lithiasis. Oxid Med Cell Longev. 2015; 2015:813737. 8. Grases F, et al. Efficacy of mixtures of magnesium, citrate and phytate as calcium oxalate crystallization inhibitors in urine. J Urol. 2015 Sep; 194(3):812-9. 9. Massey L. Magnesium therapy for nephrolithiasis. Magnesium Research 2005; 18 (2): 123-6. 10. Fernandez-Concha Schwalb J, et al. Randomized clinical trial on urinary pH monitoring and nutraceutical intervention in the prevention of ureteral stent calcification. Poster accepted at EAU19, December 2018. AM19-4171. 11. Grases F, et al. Simplified methods for the evaluation of the risk of forming renal stones and the follow-up of stone-forming propensity during the preventive treatment of stone-formation. Urolithiasis. 2016 Feb;44(1):77-82. 12. De Coninck V, et al. Evaluation of a portable urinary pH meter and reagent strips. Journal of Endourology, 2018 Jul; 32(7): 647-652.