นิพนธ์ต้นฉบับ Original Article

The Survival and Its Associated Factors among Patients with Diabetic Kidney Disease on Continuous Ambulatory Peritoneal Dialysis: Three -Year Experience in Maharat Nakhon Ratchasima Hospital.

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Objective: To evaluate the patient survival and the influencing factors in continuous ambulatory peritoneal dialysis (CAPD) patients with diabetic kidney disease at Maharat Nakhon Ratchasima Hospital. Patients & Methods: Medical records of diabetic CAPD patients in Maharat Nakhon Ratchasima Hospital during January 1, 2008-December 31, 2010 were reviewed. The collected data included demographic details, underlying diseases, body mass index (BMI), laboratory findings, peritonitis episodes, and outcome of therapy on December 31, 2010. Results: One hundred and eleven diabetic CAPD patients were analysed. There were 57.7% male, mean age was 51.9±14.4 years (range 15.3-78.6 years). The CAPD patients had anemia (67.3%), hypokalemia (36.3%) and hypoalbuminemia (66.3%). In the follow up period, total observation period was 977.4 patient-months. Patient survivals at 1, 2 and 3 years were 64.4%, 47.5% and 47.5%, respectively. The mean patient survival time was 25.6 months. The influencing factors that lowered patient survival were age above 60 years, serum potassium below 3.5 mEq/L and serum albumin <3.5 g/dL. Peritonitis rate was 21.2 patient-months per episode. Streptococcus spp. and Staphylococcus aureus were common causative organisms. Mean patient survival in no-peritonitis group was higher than peritonitis group, without statistic significance. At the end of study, the patients continued on CAPD (60.2%), switched to hemodialysis (4.6%), were referred to another center (0.9%), and died (34.3%). Conclusion: Diabetic CAPD patients had had high incidences of hypokalemia and hypoalbuminemia that influenced the survival of these patients. The peritonitis was not significant influencing factors for patient survival. So there was importance to improve patient survival by correcting the influencing factors and nutritional status in diabetic CAPD patients.

Keywords: Continuous ambulatory peritoneal dialysis, Diabetic kidney disease, Survival, Influencing factors affecting survival

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บทคัดย่อ

การรอดชีวิตและปัจจัยที่มีผลต่ออัตรารอดชีวิตในผู้ป่วยโรคไตเรื้อรังจากเบาหวานที่ล้างไต ทางช่องท้อง: ประสบการณ์ 3 ปีในโรงพยาบาลมหาราชนครราชสีมา วราภรณ์ เลียวนรเศรษฐ, พ.บ.*, สุชาติ เจนเกรียงไกร, พ.บ.*, วารุณี เศวตมาลย์, พย.บ.*, สิริลักษณ์ สุทธรัตนกุล, พย.บ.* *หน่วยไต กลุ่มงานอายุรกรรม โรงพยาบาลมหาราชนครราชสีมา นครราชสีมา 30000 เวชสาร โรงพยาบาลมหาราชนครราชสีมา 2554; 35: 13-21.

้ วั**ทถุประสงค์:** เพื่อประเมินอัตรารอดชีวิตและปัจจัยที่มีผลต่ออัตรารอดชีวิตในผู้ป่วยโรคเบาหวานและใด้รับการรักษา ด้วยการล้างใตทางช่องท้องในโรงพยาบาลมหาราชนครราชสีมา **ผู้ป่วยและวิธีการ:** ศึกษาข้อมูลผู้ป่วยโรคเบาหวานที่ ใด้รับการรักษาด้วยการถ้างใตทางช่องท้องในโรงพยาบาลมหาราชนครราชสีมา ตั้งแต่ 1 มกราคม 2551 ถึง 31 ธันวาคม 2553 โดยเก็บข้อมูลพื้นฐาน, ดัชนีมวลกาย, ผลตรวจทางห้องปฏิบัติการ, อัตราการติดเชื้อในช่องท้องและผลการรักษา เมื่อวันที่ 31 ธันวาคม 2553 **ผลการศึกษา:** ผู้ป่วยโรคเบาหวานที่ถ้างใตทางช่องท้องจำนวน 111 ราย เป็นเพศชาย ร้อยละ 57.7, อายูเฉลี่ย 51.9±14.4 ปี (พิสัย 15.3-78.6 ปี) ผู้ป่วยมีภาวะซีคร้อยละ 67.3, ภาวะ โปแตสเซียมต่ำร้อยละ 36.3 และภาวะ ไข่ขาวในเลือดต่ำร้อยละ 66.3 ระยะเวลาในการศึกษาทั้งหมด 977.4 ผู้ป่วย-เดือน อัตรารอดชีวิตผู้ป่วยที่ 1 ปี, 2 ปี และ 3 ปีเท่ากับร้อยละ 64.4, 47.5 และ 47.5 ตามลำดับ ค่าเฉลี่ยอัตรารอดชีวิตเท่ากับ 25.6 เดือน ปัจจัยที่มีผลต่ออัตรา รอดชีวิตต่ำได้แก่ อายุมากกว่า 60 ปี, ระดับ โปแตสเซียมต่ำกว่า 3.5 มิลลิอิคิวาเลนท์/เคซิลิตร, ภาวะไข่ขาวในเลือดต่ำ กว่า 3.5 กรัม/เคซิลิตร อัตราการติดเชื้อในช่องท้องเท่ากับ 21.2 ผู้ป่วย-เคือนต่อครั้ง พบว่าเชื้อสเต็ป โตคอคคัสและเชื้อ สแตฟไฟโลคอคคัส ออเรียส เป็นสาเหตุของการติดเชื้อในช่องท้องที่พบได้บ่อย ผู้ป่วยที่ไม่มีการติดเชื้อในช่องท้อง ้มีอัตรารอดชีวิตสูงกว่ากลุ่มที่มีการติดเชื้อในช่องท้องแต่ไม่แตกต่างอย่างมีนัยสำคัญ เมื่อสิ้นสุดการศึกษาพบว่าผู้ป่วย ้ที่ยังคงรักษาด้วยการล้างใตทางช่องท้องร้อยละ 60.2, เปลี่ยนไปฟอกเลือดร้อยละ 4.6, ส่งต่อไปรักษาที่อื่นร้อยละ 0.9 และเสียชีวิตร้อยละ 34.3 สรป: ผู้ป่วยเบาหวานที่ล้างไตทางหน้าท้องมือบัติการณ์การเกิดภาวะ โปแตสเซียมและไข่ขาว ในเลือดต่ำสูงซึ่งปัจจัยดังกล่าวมีผลต่ออัตรารอดชีวิตในผู้ป่วยล้างไตทางช่องท้อง อุบัติการณ์การติดเชื้อในช่องท้องไม่ ้ มีผลต่ออัตรารอดชีวิตในผู้ป่วยโรคเบาหวานที่ล้างไตทางช่องท้องอย่างมีนัยสำคัญ ดังนั้นการแก้ไขปัจจัยดังกล่าวและ ภาวะโภชนาการมีความสำคัญต่ออัตรารอดชีวิตในผู้ป่วยเบาหวานที่ล้างใตทางหน้าท้อง **คำสำคัญ:** การล้างใตทางหน้าท้อง, โรคไตเรื้อรังจากเบาหวาน, อัตรารอดชีวิต, ปัจจัยที่มีผลต่ออัตรารอดชีวิต

Introduction

Incidence of end stage renal disease (ESRD) has increased every year in the world^(1,2). The standard treatment of ESRD is renal replacement therapy (RRT) such as hemodialysis (HD), continuous ambulatory peritoneal dialysis (CAPD) and kidney transplantation (KT)⁽³⁻⁵⁾. Maximum annual growth of CAPD in the

world has been reported from Asian countries. Several reports have shown patient survival rate in peritoneal dialysis is better than HD in the first 2 years, and then they are similar in long term survival⁽⁶⁻⁸⁾. There were many factors including old age, comorbid diseases especially diabetes mellitus (DM), malnutrition had affected morbidity and mortality in CAPD patients. DM

is common cause of ESRD and one of influencing factors for patient survival⁽⁸⁻¹⁰⁾.

In Thailand, ESRD patients who underwent CAPD modality were increased every year. In Maharat Nakhon Ratchasima Hospital, we had started 'PD first' program since 2008, all of diabetic-ESRD patients under UCS who were registered to CAPD in Maharat Nakhon Ratchasima Hospital were recruited for the analysis of factors that influenced the patient survival, during January 1, 2008-December 31, 2010.

Materials and Method

Medical records of 111 adult diabetic patients with newly diagnosed ESRD who underwent CAPD under UCS in Maharat Nakhon Ratchasima Hospital during January 1, 2008-December 31, 2010 were reviewed.

Inclusion criteria: All of adult diabetic ESRD patients under UCS who were registered, performed CAPD and followed up since January 1, 2008 at Maharat Nakhon Ratchasima Hospital until death, change to hemodialysis or kidney transplantation modalities, referral to other centers, or until December 31, 2010. Technique failure was defined by changing the treatment to hemodialysis or removal of the Tenchkoff catheter.

Exclusion criteria: CAPD patients' age below 15 years, no diabetes mellitus and refusal to follow up at CAPD clinic in Maharat Nakhon Ratchasima Hospital.

Data collection consisted of demographic details, underlying diseases; diabetes mellitus, body mass index (BMI) classification by Asian-pacific criteria (21,222), laboratory findings including hemoglobin, hematocrit, serum potassium, serum calcium, serum phosphate, serum albumin, parathyroid hormone (PTH) and peritonitis episode.

Statistical analysis

All data were expressed as percentage, mean and standard deviation (SD). Comparisons were performed using the Student's t or Chi-squared tests. Patient survival was performed using Kaplan-Meier analysis. Comparison of the survival curves was made using log rank test and it would be considered statistical significant if p was < 0.05 (p < 0.05).

Results

One hundred and eleven diabetic CAPD patients were analysed. Sixty-four patients were male (57.7%) and forty-seven patients were female (42.3%), mean age was 51.9±14.4 years (range 15.3-78.6 years). Mean body mass index (BMI) was 23.0±3.5 kg/m² (range 16.2-31.9). Sixty-six patients (67.3%) had hemoglobin below 10 g/dL. Thirty-three patients (36.3%) had serum potassium less than 3.5 mEq/L. Sixty-Three patients (66.3%) had serum albumin levels below 3.5 g/dL. Six patients (9.4%) had parathyroid hormone levels below 150 pg/dL while nineteen patients (29.7%) had parathyroid hormone levels above 300 pg/dL (Table 1). Total observation period was 977.7 months (mean 9.1+7.9 months). The patient survival in diabetic CAPD patients was 64.4%, 47.5% and 47.5% (at 1 year, 2 years and 3 years, respectively), mean duration of patient survival was 25.6 months.

The subgroup univariate analysis for factors affecting patient survival in diabetic CAPD patients was shown as Table 2. The factors that lowered patient survival in diabetic CAPD patients were old age (mean patient survival in group age \geq 60 years vs. group age \leq 60 years was 11.7 vs. 28.1 months: p=0.015) (Figure 1), hypokalemia (mean patient survival in group serum

(mg/dL) <8.9

9.0-10.9

Serum phosphate: mean±SD

Parathyroid hormone (pg/ml)

≥11.0

(mg/dL) <4.5

4.5-5.5

≥5.5

<150

≥300

150-299.9

Table 1 Laboratory findings in diabetes mellitus (DM) group.

Variables Patients (%) Range BMI: mean±SD (kg/m²) 23.0 ± 3.5 16.2-31.9 <18.5 9 (11.2%) 18.5-23.0 34 (42.5%) >23.0 37 (46.2%) Hemoglobin: mean \pm SD (g/dL) 9.3 \pm 1.7 4.5-12.8 <10 66 (67.3%) ≥10 32 (32.7%) Serum potassium: mean±SD 3.7 ± 0.8 2.2-6.0 (mEq/L)< 3.5 33 (36.3%) ≥3.5 58 (63.7%) Serum albumin: mean±SD 1.9-4.1 3.2 ± 0.5 (g/dL)<3.5 63 (66.3%) ≥3.5 32 (33.7%) Serum calcium: mean±SD 8.7 ± 1.0 5.4-12.7

60 (61.9%)

36 (37.1%)

1 (1%)

 4.4 ± 1.4

54 (56.8%)

22 (23.2%)

19 (20.0%)

 243.8 ± 235.9

6 (9.4%)

39 (60.9%)

19 (29.7%)

1.6-8.2

11.2-1343.5

Table 2 Factors affecting mean patient survival time in diabetes mellitus group (DM)

Factors	Mean patient survival (Months)	<i>p</i> -value
Mean patient survival	25.6	
Age:		
<60 years	28.1	0.015
>60 years	11.7	
BMI:		
$<18.5 \text{ kg/m}^2$	11.4	0.117
$18.5-23.0 \mathrm{kg/m^2}$	24.5	
$>23.0 \text{ kg/m}^2$	18.7	
Hemoglobin:		
<10 g/dL	13.2	0.113
≥10 g/dL	29.2	
Serum potassium:		
<3.5 mEq/L	10.6	0.001
≥3.5 mEq/L	31.6	
Serum albumin:		
<3.5 g/dL	12.9	0.015
≥3.5 g/dL	34.9	
Serum calcium:		
<8.9	12.4	0.110
9.0-10.9	29.6	
≥11.0	9.1	
Serum phosphate:		
<4.5 mg/dL	15.5	0.609
4.5-5.5 mg/dL	14.7	
\geq 5.5 mg/dL	22.0	
Peritonitis		
absence	29.6	0.123
presence	12.8	

potassium <3.5 mEq/L vs. group serum potassium \geq 3.5 mEq/L was 10.6 vs. 31.6 months: p=0.001) (Figure 2) and hypoalbuminemia (mean patient survival in group serum albumin <3.5 g/dL vs. group serum albumin \geq 3.5 g/dL was 12.9 vs. 34.9 months: p=0.015). (Figure 3)

There were 46 peritonitis episodes. Twenty-eight episodes (60.9%) had negative culture, eleven episodes showed gram positive cocci and seven episodes showed gram negative bacilli. The most common causative organism was gram positive cocci such as *Streptococcus*

Survival Functions

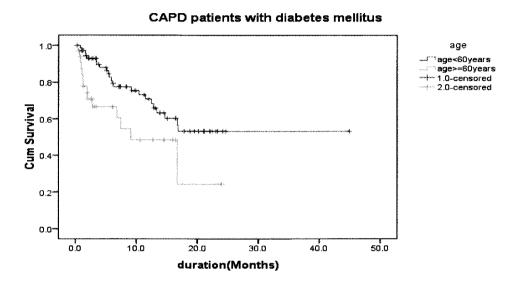


Figure 1 The survival in diabetic CAPD patients according to age group by Kaplan-Meier

spp. (13.0%) and Staphylococcus aureus (10.8%) in this study. (Figure 4) Mean peritonitis rate was one episode per 21.2 patient-months. The peritonitis group had shorter mean patient survival than no-peritonitis group, but there was no significant difference. (Table 2)

At the end of study, sixty-five patients (60.2%) were ongoing CAPD modality, five patients (4.6%) shifted to hemodialysis modality, thirty-seven patients (34.3%) died and only one patient was referred to another center. (Figure 5)

Survival Functions

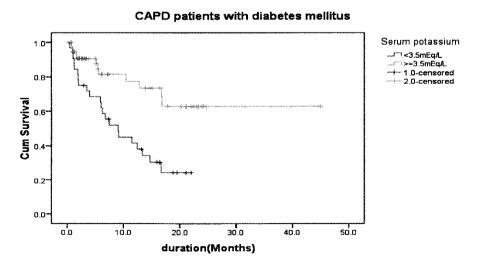


Figure 2 The survival in diabetic CAPD patients according to serum potassium by Kaplan-Meier

Survival Functions

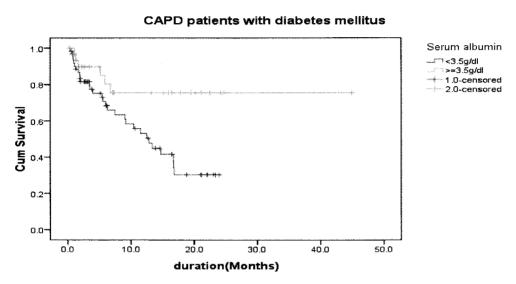


Figure 3 The survival in diabetic CAPD patients according to serum albumin by Kaplan-Meier

Discussion

Incidence of end stage renal disease (ESRD) has increased every year in the world^(1,2). DM is common cause of ESRD and one of influencing factors to patient survival⁽⁸⁻¹⁰⁾. The factors that increased mortality rate were diabetes mellitus, old age, hypoalbuminemia⁽¹⁰⁻¹²⁾.

Median survival in diabetic CAPD patients was significantly inferior to that in non-diabetic patients. Patient survival of diabetics was 85%, 62%, 48% at 1, 2, and 3 years, respectively⁽¹⁴⁾. In the previous study, we reported the overall outcome of CAPD therapy. There was rather poor outcome in the first year of 'PD'

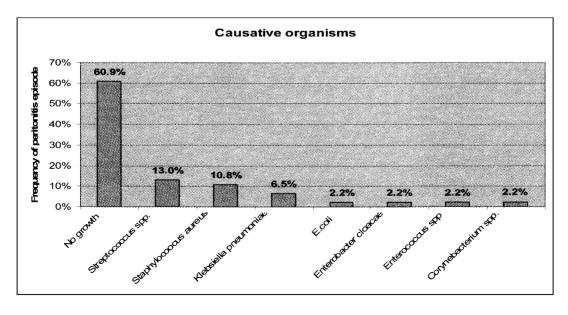


Figure 4 Result of peritoneal dialysate fluid culture in diabetic-CAPD patients with peritonitis

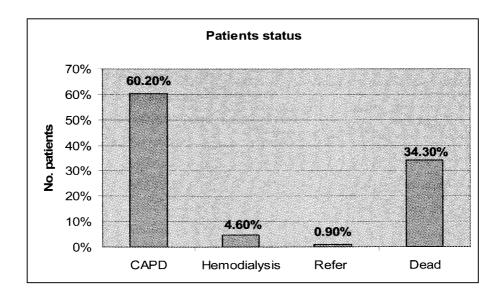


Figure 5 Status of diabetic-ESRD Patients on CAPD (at the end of study)

program in our center⁽¹¹⁾. We studied the factors affecting the patient survival in ESRD patients with DM undergoing CAPD at our center for 3 year-experience in this study. We found that the patient survival rates in diabetic CAPD patients were 64.4%, 47.5% and 47.5% at 1, 2 and 3 years, respectively. There were lower survival rate in our CAPD patients in the first year and similar survival rate at 3 years of treatment. In subgroup univariate analysis, elderly patients (age above 60 years), hypokalemia (serum potassium below 3.5 mEq/L) and hypoalbuminemia (serum albumin below 3.5 g/dL) were significant factors that lowered mean patient survival more than others. As several previous studies, diabetes mellitus, old age, hypoalbuminemia were risk factors that increased mortality rate (10-12). There was high prevalence of high transporters in DM patients and it was the significant predictor for patient survival in the previous study⁽²⁸⁾. But peritoneal equilibrium test (PET) was not performed in this study, so the prevalence of high transporter was not performed in our patients.

The previous study presented that lower BMI could increase relative risk of death in CAPD patients (7,27) while the obesed patients had equivalent survival to the patients with normal BMI (15,16). In our study, there was no significant difference in patient survival due to BMI variable, it may be from the difference in BMI category and in ethnics in each study (Caucasians and Asians population (21,22))

In the past, peritonitis was a leading cause of mortality (about 0.8%-12.5%) and one of causes of technique failure in CAPD patients⁽¹⁷⁾. The factors that were associated with increased risk of peritonitis were low baseline serum albumin^(12,17,18) and anemia⁽¹⁹⁾. Mutiple studies have shown high prevalences of malnutrition and hypoalbuminemia in CAPD patients^(12,20,24). Espinosa et al found that DM patients showed a significantly poorer nutritional status than non-DM patients⁽²³⁾. Nakamoto et al showed the incidences of hypoproteinemia and protein loss in DM were higher than that in non-DM patients, probably due to high

permeability of peritoneal membrane in DM patients^(25,26).

In our study, peritonitis rate was one episode per 21.2 patient-months. The most common causative organism was gram positive cocci, no evidence of fungal peritonitis in this study. Mean patient survival of CAPD patients with peritonitis group was shorter than that of no-peritonitis group but there was no statistic significance. This result was differed from other study. Because almost all care-givers were PD exchangers. But in this study, comparison between DM and non-DM groups, data of care-givers were not analysed.

Conclusion

This study was performed among diabetic ESRD patients who were on CAPD, it was shown that the old age (more than 60 years), poor nutritional status (hypoalbuminemia and hypokalemia) were significant factors that lowered the patient survival than other factors. The elderly diabetic CAPD patients require special attention to improve nutritional status more than non-DM group. For our CAPD patients, we need to evaluate the peritoneal function and to improve nutritional status in the further study.

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