# REVIEW

# Clinical Problems in the Hospitalized Parkinson's Disease Patient: Systematic Review

Oliver H.H. Gerlach, MD,<sup>\*</sup> Ania Winogrodzka, MD, PhD, and Wim E.J. Weber, MD, PhD Section of Movement Disorders, Department of Neurology, Maastricht University Medical Centre, Maastricht, The Netherlands

### ABSTRACT:

The problems Parkinson's disease (PD) patients encounter when admitted to a hospital, are known to be numerous and serious. These problems have been inventoried through a systematic review of literature on reasons for emergency and hospital admissions in PD patients, problems encountered during hospitalization, and possible solutions for the encountered problems using the Pubmed database. PD patients are hospitalized in frequencies ranging from 7 to 28% per year. PD/parkinsonism patients are approximately one and a half times more frequently and generally 2 to 14 days longer hospitalized than non-PD patients. Acute events occurring during hospitalization were mainly urinary infection, confusion, and pressure ulcers. Medication errors were also frequent adverse events. During and after surgery PD patients had an increased incidence of infections, confusion, falls, and decubitus, and 31% of patients was dissatisfied in the way their PD was managed. There are only two studies on medication

## Introduction

Parkinson's disease (PD) is the second most common neurodegenerative disorder with a life-time risk of 2 percent in men and 1.3 percent in women.<sup>1</sup> Although the disorder is generally slowly progressive, it does have a major impact on disability and quality of life of affected patients.<sup>2,3</sup> One of the lesser studied aspects of PD is the spectrum of problems PD patients encounter,

Published online 31 January 2011 in Wiley Online Library (wileyonlinelibrary.com). DOI: 10.1002/mds.23449 continuation during surgery and one analyzing the effect of an early postoperative neurologic consultation, and numerous case reports, and opinionated views and reviews including other substitutes for dopaminergic medication intraoperatively. In conclusion, most studies were retrospective on small numbers of patients. The major clinical problems are injuries, infections, poor control of PD, and complications of PD treatment. There are many (un-researched) proposals for improvement. A substantial number of PD patients' admissions might be prevented. There should be guidelines concerning the hospitalized PD patients, with accent on early neurological consultation and team work between different specialities, and incorporating nonoral dopaminergic replacement therapy when necessary. © 2011 Movement Disorder Society

Key Words: Parkinson's disease; hospitalization; emergency room; perioperative

once they are admitted to a hospital. In our own and others' experience hospital admissions of PD patients are often problematic, especially so when patients are admitted on non-neurological wards.<sup>4,5</sup> Problem areas are exact timing or lack of drug administration, administration of contra-indicated drugs, complications due to immobilization, and psychiatric disorders triggered by the hospital admission.<sup>6–8</sup> As most non-neurologically educated health care personnel are unfamiliar with PD, protocols would be helpful to improve the care of PD patients in such environments.

We recently surveyed the majority of movement disorder specialist neurologists in the Netherlands and found that no specific guidelines or protocols exist to guide caregivers and PD patients in the hospital environment (manuscript in preparation). Before such guidelines can be formulated, one needs to know the prevalence and spectrum of the problems PD patients may experience during their hospital stay. To this end, we systemically reviewed all the

<sup>\*</sup>Correspondence to: Dr. Oliver H.H. Gerlach,, Section of Movement Disorders, Department of Neurology, Maastricht University Medical Centre, Maastricht, The Netherlands, PO Box 5800, 6202 AZ Maastricht, The Netherlands; o.gerlach@mumc.nl

Relevant conflicts of interest/financial disclosures: Nothing to report. Full financial disclosures and author roles can be found in the online version of this article.

Received: 11 December 2009; Revised: 16 July 2010; Accepted: 27 August 2010

TABLE 1. Pubmed search deta	ls
-----------------------------	----

Search	Number of hits
Parkinson* and emergenc*	422
Parkinson* and hospitali*	645
Parkinson* and healthcare	892
Parkinson* and hospital admission*	45
Parkinson* and hospital utilization*	4
Parkinson* and resource use	186
Parkinson* and perioperative	93
Parkinson* and preoperative	451
Parkinson* and intraoperative	453
Parkinson* and anesthes*	373
Parkinson* and surgery and cognit*	456
Parkinson* and surgical problems	260
Parkinson* and surgery and apomorphine	289
Parkinson* and postoperative	1231
Parkinson* and surgery and complication*	2241
Medication interruption or drug manipulations OR discontinuation or dose reduction and levodopa	576
Parkinson* and drug withdrawal	607
Parkinson* and lisuride	268
Parkinson* and medication error	59
Amantadine and intravenous*	106
Rotigotine	247
Parkinson* and fracture	236
Parkinson* and nurse	276
Parkinson* and nurse specialist	26
Parkinson* and orthopaedic	94
Parkinson* and orthopedic*	52

existing literature on the problems encountered by the hospitalized PD patients.

## **Methods**

We systematically reviewed the literature on reasons for emergency room (ER) and hospital admissions in patients with PD, problems encountered during hospitalization of this patient population, and possible solutions for the encountered problems, using the Pubmed database. Last research date was 17 June 2010.

To identify articles, we included (combinations of) keywords: See Table 1 for search details. Subsequently we analyzed the abstracts for relevant articles: as relevant articles we defined those as pertaining to the following four areas:

- 1. Analysis of prevalence and reasons for ER visits and subsequent admission.
- 2. Clinical problems during hospital stay.
- 3. Peri- and postoperative problems.

4. Suggestions for improvement of care for the hospitalized PD patient.

We also searched the reference list of each relevant article for other applicable articles.

In our search, there were no language limitations. We excluded articles concerning brain surgery.

## Results

### **Emergency Room and Hospital Admissions**

Patients with PD often need emergency treatment. There are four studies analyzing the reasons for ER admission,<sup>4,9-11</sup> including one case report<sup>11</sup> (totaling 327 PD patients, Table 2). A total of 16–45% of PD patients visit an ER once a year.<sup>9,10,12,13</sup> PD patients visit the ER more frequently than their matched reference group (0.6 vs. 0.4; P = 0.05).<sup>14</sup>

We found 12 studies on hospital admissions (Table 3): 11 studies totalling 3216 PD/parkinsonism patients,<sup>4,9,10,13,15-19,21,22</sup> and one study<sup>20</sup> on a database with 15304 PD/parkinsonism patients. In these studies, PD patients are hospitalized in frequencies ranging from 7–28% per year.<sup>9,10,15,19,20</sup>

PD/parkinsonism patients are hospitalized approximately one and a half times more frequently<sup>20,23,24</sup> and generally 2 to 14 days longer<sup>12,16,17,19,21,24–29</sup> than non-PD patients, although there is not a difference in every study.<sup>15,30</sup>

Reasons for emergency and hospital admission can be divided in:

- 1. Direct disease related morbidity: motor complications, psychiatric symptoms, autonomic dysfunction, sensory symptoms, sleep disorders, and side effects of anti-parkinsonian drugs.
- 2. Indirect disease related morbidity: traumas and pneumonia.
- 3. Non-PD related causes.

As most studies vary greatly in selection of patients, exact relative proportions of these 3 groups cannot be assessed (Tables 2 and 3). Some studies found that PD patients are more likely to be admitted to the ER and hospital for complications of the disease and its management than for primary motor problems.<sup>13,18</sup> A part (5–21%) of the patients were first diagnosed to have PD during a hospitalization.<sup>19,21</sup>

### **Problems During Hospitalization**

We found one prospective study on acute events occurring during hospitalization.<sup>4</sup> When admitted to a neurology ward (83 PD patients, mean age 69 years, mean disease duration 6 years), patients received an average of 0.6 non-neurological consultations. Reasons were: (aspiration) pneumonia, urinary infections

Study	Inclusion	Exclusion	Number of patients	Design	Control group	ER admissions (%)	Reasons ER visit
Vargas et al., 2007 <sup>9</sup>	PD	Hoehn and Yahr Stage 5	144	Retrospective	No	22% in 1 yr	Primarily side effects of anti-Parkinsonian drugs
		Parkinsonism					
		Severe cognitive dysfunction					
Cosentino et al., 2005 <sup>10</sup>	PD	Parkinsonism	130	Retrospective	No	22% in 1 yr	Injuries 61%, mainly fractures 37%
		Admissions					Abdominal pain 6%
		related to PD					Pneumonia, dysphagia, dyskinesia, epistaxis, hearing loss, pulp disease, teeth extraction, lumbago, pain in joint: all 3%
Martignoni et al., 2004 <sup>4</sup>	PD	Parkinsonism	48	Prospective	No	All selected patients	Cardiovascular 27%
							Trauma with fractures 19%
							Chest or abdominal problems 19%
							Neurological (both related and unrelated to PD) 17%
							Head injury 6%
							Hip prothesis displacement 2%
Factor and Molho, 2000 <sup>11</sup>	PD		5	Case report	No	All selected patients	Severe motor off periods, dyskinesia, psychosis, acute confusion, panic disorder, pain

#### TABLE 2. Emergency room admissions

ER, emergency room; PD, Parkinson's disease.

and retention, diarrhoea, atrial fibrillation, postural hypotension, low back pain, and TIA. Specialists consulted most frequently were cardiologists, internal medicine specialists and orthopaedic surgeons. Acute events observed during hospitalization on non-neurological department of 20 patients (mean age 80 years) after ER visits were: Urinary infection (33%), agitation, confusion and hyperthermia (28%), pressure ulcers and leg oedema (22%), bowel occlusion and hypotension (11%), and dysphagia (6%). Mortality rate during hospitalization of this last group was 20%. We found one retrospective study on this subject (173 PD patients), but this study is unclear on the presence or absence of complications before the hospital admission.<sup>21</sup> Problems identified in this study do accord however with those described above.<sup>4</sup>

Another retrospective study shows that in 74% of 35 emergency hospital admissions of PD patients, their medication was stopped, omitted or prescribed inappropriately with 61% of this group suffering significant sequelae. Nonadherence to medication schedules was a large problem and in 11% of cases antidopaminergic medication (metaclopramide) was prescribed.<sup>31</sup>

In a study analyzing inpatient falls, the use of anti-Parkinson's medication was an evident risk factor (OR 5.04).<sup>32</sup> Apart from the studies above, we found several opinionated views and reviews on problems during hospitalization of PD patients. These authors point to disease fluctuations, stress,<sup>7</sup> and medications like butyrophenones, phenothiazines, and metoclopramide, prescribed during hospitalization, as possible causes of PD exacerbation.<sup>6,33–38</sup> Additionally, sleep disorders, verbal communication problems, nutritional intake difficulties, and cognitive changes are mentioned as problem areas.<sup>7,19,39</sup>

#### **Perioperative Problems**

We found 15 retrospective studies on problems during and after surgery in patients with PD, with an emphasis on the postoperative period. Of these 15, two retrospective studies included a control group of non-PD patients.<sup>25,27</sup> One retrospective study focused on postoperative confusion<sup>33</sup> and two retrospective studies on medication errors.<sup>40,41</sup> Another study with 10 patients is actually a case report with 10 PD patients,<sup>42</sup> and the remaining 9 papers are retrospective studies on PD patients undergoing orthopaedic surgery.<sup>29,43-50</sup>

The first study compared 234 PD patients with a control group of 40,979, undergoing elective bowel

Study	Inclusion	Exclusion	Number	Design	Control group*	Hospital admissions	Five most frequent admission reasons
Vossius et al., 2010 <sup>15</sup>	PD		108 patients	Prospective	Yes	All selected patients	PD related symptoms 25%
							Vascular disorders 14%: significant less than control group
							Pulmonary disorders including pneumonia 12%
							Trauma 12%: significant more than control group
							Cancer 7%: significant less than control group
							Remark: discharge diagnosis instead of admission diagnosis
Klein et al., 2009 <sup>16</sup>	PD		143 patients	Retrospective	No	All selected patients	Motor complications 37%
	Emergency						Psychosis 24%
	admissions to Neurological Department						Somatic problems 14%
Guneysel et al., 2008 <sup>13</sup>	PD	Admissions resulting in death	76 patients	Prospective	No	All selected patients	Trauma 28%
	Emergency	PD diagnosis					UTI 20%
	hospital	during					Cardiovascular 15%
	admission	ER visit					Pneumonia, cerebrovascular both 12%
							GI 8%
Vargas et al.,	PD	Hoehn and Yahr stage 5	144 patients	Retrospective	No	28% programmed	Complication treatment
2007 <sup>9</sup>		Parkinsonism Severe cognitive dysfunction				admissions in 1 year	Drug adjustment
Louis et al., 2007 <sup>17</sup>	Young-onset PD	Obstetrical admissions	714 patients	Retrospective	Yes	All selected patients	Psychosis 23%: significant more than control group
	18-40 years old						Craniotomy 7%: significant more than control group
							Pneumonia, UTI both 6%
							Headache or seizure 4%
							Rehabilitation 3%: significant more than control group Remark: discharge diagnosis
							instead of
							admission diagnosis
							(Continu

## TABLE 3. Hospital admissions

Study	Inclusion	Exclusion	Number	Design	Control group*	Hospital admissions	Five most frequent admission reasons			
Temlett and Thompson, 2006 <sup>18</sup>	PD		761 hospital admissions	Retrospective	No	All selected patients	Primary for PD 15% Falls and fractures 11% Pneumonia, cardiac disease both 10% Gl 9%			
Cosentino et al., 2005 <sup>10</sup>	PD	Parkinsonism	130 patients	Retrospective	No	19% in 1 year	Organic brain syndrome 6% Diseases of digestive system 17%			
		related to PD					system, rehabilitation both 14% Cataract 10%			
							Injury, chest pain both 7% Sleep disturbance, epistaxis, abdominal pain, osteoarthrosis: all 3%			
Woodford and	PD	Parkinsonism	367	Retrospective	No	35% in 4 year	Cardiovascular 20%			
Walker, 2005 <sup>19</sup>	Emergency	Elective admissions	patients				Falls 13%			
	hospital	Day-case procedures					Pneumonia 11%			
	admission	Admissions resulting in					UTI 9%			
		death of patient					Decreased mobility/ dyskinesia, psychiatric both 8%			
Guttman	PD	< 25 years of age	15304	Retrospective	Yes	68%	Compared with controls			
et al., 2004 <sup>20</sup>	Use of PD drugs	) drugs	patients			in 6 years	First: aspiration pneumonia			
							Second: affective psychosis			
							Third: hip fractures			
							Fourth: urinary tract disorders including infections			
							Fifth: septicemia			
Martignoni et al., 2004 <sup>4</sup>	PD	Parkinsonism	132 patients	Prospective	No	All selected patients	Admission to neurological department			
							Poor control of PD symptoms 37%			
							Neurological 25%			
										Diagnosis confirmation, psychiatric complaints both 13%
							Sudden worsening of motor symptoms 7%			
							Head trauma and fracture 2%			
							Admission to non-neurological department			
							Medical and infectious illnesses 27%			
							Traumas with fracture 24%			
<b>T</b>		<b>.</b>	470			AU 1	Cardio-circulatory 22%			
Tan et al., 1998 <sup>21</sup>	PD	Drug induced parkinsonism	173 patients	Retrospective	No	All selected patients	Chest infection 22% Falls 13%			
							Control of PD symptoms 10%			
		Parkinson-plus syndromes					General medical problems 9%			
							(Continued)			

## **TABLE3.** (Continued)

TABLE	3.	(Continued)
-------	----	-------------

Study	Inclusion	Exclusion	Number	Design	Control group*	Hospital admissions	Five most frequent admission reasons
Urinary dysfunction 8%							
Kessler 1972 <sup>22</sup>	PD		468 patients	Retrospective	Yes	All selected	PD 19%
	Paralysis agitans					patients	Circulatory system 16%
	Parkinsonism						Digestive system 10%
							Accidents 9%
							Respiratory system 8%
							Remark: Neoplasm 4%: significant less than control group

\*Comparison with inclusion group concerning admission reasons.

PD, Parkinson's disease, UTI, urinary tract infection, GI, gastrointestinal.

resection, cholecystectomy, or radical prostatectomy, during a 6-year period. PD patients had a significantly increased incidence of aspiration pneumonia, urinary-tract infection, bacterial infections, with odds ratios of 3.8, 2.0, 1.7, respectively. Odds ratios for postoperative delirium and hypotension in PD patients were 2.6 and 2.5, with lesser significance. PD patients had a mortality rate of 7.3%, compared with a 3.8% in the control group.<sup>25</sup>

In the second study with a non-PD control group, 51 PD patients, treated on different surgical departments, were compared using matched-pair analysis. There were significantly more postoperative falls in the PD group (18% vs. 4%), and a higher although not significantly increased number of urinary tract infections (33% vs. 24%), pneumonia (10% vs. 4%), and also wound infection, urinary retention, respiratory insufficiency, and postoperative confusion. PD patients were hospitalized for more days and stayed on the intensive care unit longer.<sup>27</sup>

Another retrospective study showed that 60% of 25 PD patients with no pre-existent mental status abnormalities suffered postoperative confusion, some with hallucinations. The onset of the delirium was often delayed, 70% after 36 hours. In this study, there was no relationship between delirium and type of anti-Parkinson medication or anaesthetic procedure.<sup>33</sup>

In a retrospective study on pharmacological management during 51 surgical admissions of PD patients or patients with parkinsonian syndromes treated with PD medication, 71% had missed doses of their medication. Overall, antidopaminergic medication was prescribed in 41% and administrated in 22% although not allowed. 47% (69% for non-day-cases admissions) had complications: neuropsyciatric 41%, falls 8%, and worsening of motor symptoms 5%.<sup>40</sup> A second study on pharmacological management found that 30% of 92 PD patients had medication administration problems, leading to an increase in postoperative confusion or worsening of PD: 84% to 36% in the wellmanaged group. Overall 31% of patients was dissatisfied in the way their PD was managed in the perioperative period.<sup>41</sup>

In the smallest study (n = 10), all patients had complications. This paper identified the same problems as mentioned above, and additionally found pressure sores as an important problem.<sup>42</sup>

Next to the above studies, there are nine retrospective research articles on orthopaedic surgery and its complications. These studies (totalling 433 PD patients, Table 4) found pneumonia, urinary tract infections, confusion, and decubitus as the most frequent postoperative complications, in frequencies up to 49%<sup>44-50</sup> with an overall six month mortality up to 47%.<sup>44,45,47,48,50,51</sup> Apart from complications, several studies are contradictory regarding the outcome of orthopaedic procedures in PD patients,<sup>26,29,30,43-60</sup> and postoperative rehabilitation is reported to be slower.<sup>28,55</sup>

Next to these more or less formal surveys, case report and reviews mention the following perioperative problems: neuroleptic malignant syndrome, medication- or anaesthetic-induced exacerbation of PD, side-effects of PD medication, postextubation laryngeal spasm, bronchospasm, respiratory arrest, difficulty with salivation, gastrointestinal complications, deep vein thrombosis, urinary disturbances, temperature regulation problems, and tremor hampering eye surgery.<sup>6,8,34–38,61–80</sup>

#### Improvement of Care for the Hospitalized PD Patient

### Improvement During Hospitalization

There are no studies analyzing the effects of suggested recommendations/improvements. Some authors favour a multidisciplinary approach during hospitalization.<sup>4,7</sup> Other suggestions are: continuing the exact

Study	Inclusion	Exclusion	Number	Design	Intervention	Most frequent complications
Mehta et al., 2008 <sup>29</sup>	PD	Total knee arthroplasty revision	34 patients	Retrospective	Total knee arthroplasty	Confusion 35%
			39 knees			Superficial wound infection, aspiration pneumonia both 6%
Weber et al., 2002 <sup>48</sup>	PD		98 patients	Retrospective	Hip replacement	6 Month mortality 6%
			107 hips			Overall complications: 36%
						UTI 7%
						Dislocation 6%
						Postoperative confusion 4%
						Pneumonia, deep venous thrombosis both 3%
Duffy and Trousdale, 199643	PD		24 patients	Retrospective	Total knee arthroplasty	Confusion 20%
			33 knees			Deep venous thrombosis, superficial infections both 8%
						Myositis ossification, urinary retention, wound necrosis, respiratory tract infection all 4%
Turcotte et al., 1990 <sup>50</sup>	PD		87 patients	Retrospective	Hip fracture surgery	After 6 months:
			94 hips			Mortality 14%: myocardial infarction (n=5), infection (n=2), pulmonary embolism $(n=1)$ , unknown $(n=4)$
						Orthopaedic problem 14%
						Decubitus ulcers 5%
						Wound infections 4%
Vince et al., 1989 <sup>46</sup>	PD		9 patients	Retrospective	Total knee arthroplasty	Deep vein thrombus (n=4), UTIs (n=3), temporary disorientation (n=2), skin necrosis (n=1), intestinal ileus (n=1), pulmonary embolism (n=2)
			13 knees			
Staeheli et al., 198845	PD	Parkinsonism	49 patients	Retrospective	Hip fracture surgery	6 Months complication:
			50 hips			Mortality 20%: pneumonia 40%, congestive heart failure 20%, cerebrovascular accident 20%, pulmonary embolism 10%, breast cancer 10%
						UTI 20%, pneumonia 10%, decubitus ulcers 10%, pulmonary embolism 6%, cerebrovascular accident 6%, wound infection 4%
Eventov et al., 198344	PD	Impacted subcapital fractures	62 patients: 45 patients undergoing surgery	Retrospective	Hip fracture surgery	3 Month mortality surgery group 31% (1 year 38%): bronchopneumonia 43%, congestive heart failure 21%
	Ambulatory					3 Month mortality in patients not undergoing surgery 29% (35% 1 year)
						Survivors surgery group (n=31):
						UTI 23%, decubitus 23%, bronchopneumonia
						(Continued)

## TABLE 4. Orthopedic surgery, complications

Movement Disorders, Vol. 26, No. 2, 2011 203

#### **TABLE4.** (Continued)

Study	Inclusion	Exclusion	Number	Design	Intervention	Most frequent complications
						16%, contractures 6%, deep infection, cardiac arrythmias, myocardial infarction, dislocation, thrombophlebitis, paralytic ileus all 3%
						Survivors of patients not undergoing surgery (n=12):
						UTI 17%, decubitus 25%, bronchopneumonia 0%, contractures 17%
Coughlin and Templeton, 1980 <sup>47</sup>	PD		47 patients	Retrospective	Hip fracture surgery	6 Month mortality 47%
			49 hips			Decubitus 49%
	Ambulatory					Dislocation 37% (endoprosthesis)
Rothermel and Garcia, 1972 <sup>49</sup>	PD		23 patients: 16 without levodopa, 7 with levodopa	Retrospective	Hip fracture surgery	With levodopa: phlebitis n=1
						Without levodopa: debrided decubitus ulcers $n=2$ , phlebitis $n=2$ , deep hematoma $n=2$ , dislocation $n=2$ , urinary septicaemia $n=1$ , fatal myocardial infarction $n=1$

PD, Parkinson's disease; UTI, urinary tract infection.

personal medication regime, education of nurses and doctors, being attentive for early signs of complications like pneumonia to start early treatment, falling prevention, (temporary) medication adjustment, emotional support, good sleep hygiene by maintaining the home bedtime and trying to prevent sleeping during the day, sometimes consulting a sleep disorder specialist or start sleeping medication, exercise, speech therapy, sufficient nutritional intake high in fibre, adequate fluid intake, preventing a confusional state by limiting the number of care-givers and the amount of light and noise during night sleeping, avoiding certain medication harmful for the PD patient, and consulting a neurologist.<sup>7,21,31,35,37,67</sup>

#### Improvement of Perioperative Care

There are only two studies on medication continuation during surgery<sup>81,82</sup> and one study analyzing the effect of an early postoperative neurologic consultation,<sup>29</sup> and numerous case reports, and opinionated views and reviews.

During preoperative screening, some authors recommend extra attention to be paid to this group of patients, especially to respiratory status, urologic system, fluid status, cardiovascular system, gastrointestinal system, autonomic system, and cognition.<sup>6,25,35,37,66,73,74,78,83</sup> And, if necessary, supple-

mented with additional diagnostics like laboratory tests, pulmonary tests, electrocardiogram, and X-ray.<sup>25,66,73,78</sup> Most of the literature describes measures with regard to PD medication in the intraoperative period. To prevent large descents of dopamine levels intra- and postoperatively many authors advise continuation of PD medication as long as possible preoperatively and resume it as soon as possible postoperatively, and PD medication is preferably contin-ued.<sup>6,8,25,34,37,38,42,67–70,72–74,76,78,79,81,82,84–92</sup> Two studies on this topic describe PD patients using the rotigotine transdermal patch, a nonergot D1/D2/D3 dopamine agonist.<sup>81,82</sup> In the first prospective study, oral dopaminergic medication was easily switched to rotigotine before surgery and resumed afterwards in 14 PD patients undergoing surgery under general anaesthesia. Adverse events were two dopaminergic side effects namely nausea and hallucinations and one ventricular asystole.<sup>81</sup> The second study on this topic describes PD patients derived from two prospective clinical trails. PD patients undergoing surgery under general anaesthesia and who continued using rotigotine during surgery were retrospectively analyzed (n = 25). There was no worsening of PD symptoms, but (only) three complications: deep vein thrombosis, infection, and pain.<sup>8</sup>

As other substitutes for dopaminergic medication intraoperatively, continuous intravenous levodopa infusion, continuous subcutaneous infusion or immediate postoperative injections of apomorphine, and enteral levodopa/carbidopa via nasogastric tube or duodenostomy have been used by various authors, <sup>68,69,72,76,78,88,92,93</sup> but none have been studied in a controlled trial. The use of parenteral anticholinergic and antihistaminic medication as anti-Parkinson therapy is limited according to some authors, and may aggravate postoperative discomfort because of autonomic side effects and confusion. <sup>72,76,78</sup>

Some authors claim that general anaesthesia should be avoided when possible, and prefer local anaesthesia.<sup>37,66,67,73,75,90</sup> This is supported by a case report of a PD patient, undergoing surgery with regional anaesthesia, who was successfully given levodopa and carbidopa orally during the operation.<sup>86</sup> Regional anaesthesia also avoids postoperative nausea and vomiting.<sup>73</sup> Less invasive interventions, like laparoscopic surgery over abdominal surgery, are recommended.<sup>66</sup> Some authors advocate carefully considering the used operation technique.<sup>47,50</sup>

In a retrospective study, the postoperative period was analyzed in PD patients undergoing total knee arthroplasty receiving a preoperative or immediate postoperative neurologic consultation (n = 13) compared with patients receiving a delayed or no visit (n = 21). Only in the first group, there was a significant improvement in total Unified Parkinson's Disease Rating Scale with most improvement in activities of daily living but also an improvement of mood, mentation and behavior, and motor examination. In this group, there was also a shorter length of stay (P < 0.01) and less patients were confused: 15 to 48%.<sup>29</sup>

There are many more recommendations, without formal studies. Postoperatively lung expansion manoeuvres, noninvasive mechanical ventilation, percussion, adequate pain control, aspiration precautions, (early) mobilization, recognition of unique physical limitations and medication combinations, physiotherapy, turning regimens, maintenance of volume status, antiparkinsonian therapy adjustments, analyzing urine for urinary tract infections, and deep venous thrombosis prophylaxis may prevent complications like pulmonary emboli, infections, deep vein thrombosis and decubitus.  $^{6,27,42-45,50,55,59,60,66,72,74,78,83,91,94-96}$  In case of prolonged endotracheal intubation early tracheostomy is preferred.<sup>66</sup> Except for two case reports, there is no literature concerning the pre-emptive administration of drugs to prevent complications. In one case report, the cholinesterase inhibitor rivastigmine was given preoperatively to prevent delirium postoperatively in a PD patient successfully treated with this drug in two previous delirious episodes.<sup>97</sup> In another case report, prokinetics were administered in two patients to prevent paralytic ileus.<sup>92</sup> Some authors suggest preventive antibiotics to prevent infections.<sup>44,45,59</sup> As described before, the onset of the postoperative delirium is often delayed. If patients are discharged rapidly after surgery, there should be sufficient support in the home environment.<sup>33</sup> Overall, team work between different specializations, like surgery, neurology, geriatrics, and a rehabilitation unit is generally advocated.<sup>28,96,98</sup>

# Discussion

There are few studies analyzing the problems a PD patient encounters during hospitalization, and there are even less studies analyzing possible solutions. Most studies are retrospective and have small numbers of patients. In some studies, PD patients were diagnosed according to clear diagnostic criteria (like the United Kingdom Parkinson's Disease Society Brain Bank's clinical diagnostic criteria),<sup>4,9–11,15,19,48</sup> but in most studies, the diagnostic criteria were not clearly mentioned: a neurologist confirmed the diagnosis,<sup>13,43,45,46,48</sup> or medical record systems and/ or patients notes were used to identify PD patients,<sup>17,20–22,25,27,33,40</sup> or no information according to the diagnostic criteria, except the use of PD medication, was given.<sup>16,31,41,42,47,49,50,81,82</sup>

Overall PD patients are more frequently<sup>20,23,24</sup> and longer<sup>12,16,17,19,21,24–29</sup> hospitalized compared with controls. Generally the leading causes for admission are injuries (many with fractures), infections [mainly pneumonia and urinary tract infections (UTI)], poor control of PD and complications of PD treatment, psychiatric disturbances, and diseases of the circulatory and digestive system. A reduction of the number of admissions might be achieved by extra attention to fall prevention, adequate drug regulation with acuity for side effects, preventing and recognizing of early symptoms of infections and active monitoring in the home situation of both patients' vital parameters and therapy compliance.<sup>5,18,20</sup> When admitted to a hospital, most PD patients stay on a general medicine or surgery ward instead of a neurological one.<sup>5,15</sup> Apart from one small prospective study<sup>4</sup> and a retrospective study,<sup>31</sup> little is known about problems occurring during hospitalization of PD patients not undergoing surgery, mentioning the usual direct and indirect PD related problems and medication issues.

There are more studies on complications in PD patients hospitalized for surgery, particularly in the postoperative period. (Aspiration) pneumonia, UTI, bacterial infections, postoperative falls, postoperative delirium/confusion (often with a delayed onset<sup>33</sup>), and hypotension occur more frequently in this group of patients than in controls.<sup>25,27</sup> Pressure sores are also an important complication.<sup>42</sup> Mortality rates are also higher.<sup>25</sup> These data are confirmed by studies on PD patients undergoing orthopaedic surgery.<sup>29,43–50</sup> Many PD patients had postoperative medication

administration problems with more postoperative confusion or worsening of PD as a result.<sup>40,41</sup> It is striking that almost one third of the PD patients were dissatisfied concerning their PD treatment.<sup>41</sup>

### Suggestions for Improvement of Hospital Care

There are many proposals for improvement during hospitalization in PD patients with or without surgery. Suggestions on improvement do vary, but most authors agree that attention should be given to all aspects of PD and not only to motor function.

Most publications refer to the intraoperative period mainly to prevent a decline in dopamine levels because of discontinuation of dopaminergic medication, including two studies favouring the use of a rotigotine patch.<sup>81,82</sup> Most authors agree that anesthesiologists and surgeons should take the increased vulnerability of PD patients into account when planning and selecting procedures.<sup>37,47,50,66,67,73,75,86,90</sup> One small study shows that early consultation by a neurologist may prevent complications and reduce length of hospital stay.<sup>29</sup>

## Conclusions

Most studies were retrospective and had small numbers. Prospective studies with large numbers of PD patients, defined according to clear diagnostic criteria and preferable diagnosed by a specialist with special interest in PD, would be preferred for future research. Generally patients with PD are hospitalized much more frequently and longer than control groups. The leading causes are injuries, infections, poor control of PD and complications of PD treatment. The inclusion of hospitalization data into patient registries for PD could be a major improvement in identifying the important problems.

There are many (un-researched) proposals for improvement during hospitalization in PD patients. A substantial number of PD patients' admissions might be prevented. There should be guidelines concerning the hospitalized PD patients, with accent on early neurological consultation or consultation of another specialist like a geriatrician with a special interest in PD and team work between different specialities, and on sufficient training of all people involved in the treatment and recovery of this patient group. This protocol should include dopaminergic replacement therapy in case PD patients are not allowed or able to take their oral medication. Preferably this therapy should not be laborious and not invasive so that it is easily applicable.

#### References

 Poos MJJC, Gijsen R. Ziekte van Parkinson. Omvang van het probleem. Achtergronden en details bij cijfers uit huisartsenregistraties. Nationaal Kompas Volksgezondheid, versie 319 2009. Accessed 1 November 2009. Available at: [http://wwwrivmnl/vtv/object\_document/o3307n17760html].

- Chapuis S, Ouchchane L, Metz O, Gerbaud L, Durif F. Impact of the motor complications of Parkinson's disease on the quality of life. Mov Disord 2005;20:224–230.
- Rahman S, Griffin HJ, Quinn NP, Jahanshahi M. Quality of life in Parkinson's disease: the relative importance of the symptoms. Mov Disord 2008;23:1428–1434.
- Martignoni E, Godi L, Citterio A, et al. Comorbid disorders and hospitalisation in Parkinson's disease: a prospective study. Neurol Sci 2004;25:66–71.
- Antonini A, Miro L, Castiglioni C, Pezzoli G. The rationale for improved integration between home care and neurology hospital services in patients with advanced Parkinson's disease. Neurol Sci 2008;29(Suppl 5):S392–S396.
- Dorton DM. The care of patients with Parkinson's disease. J Post Anesth Nurs 1995;10:102–106.
- Vergenz S. Caring for the Parkinson's patient: a nurse's perspective. Dis Mon 2007;53:243–251.
- Stotz M, Thummler D, Schurch M, Renggli JC, Urwyler A, Pargger H. Fulminant neuroleptic malignant syndrome after perioperative withdrawal of antiParkinsonian medication. Br J Anaesth 2004;93:868–871.
- 9. Vargas AP, Carod-Artal FJ, Nunes SV, Melo M. Disability and use of healthcare resources in Brazilian patients with Parkinson's disease. Disabil Rehabil 2008;30:1055–1062.
- 10. Cosentino M, Martignoni E, Michielotto D, et al. Medical healthcare use in Parkinson's disease: survey in a cohort of ambulatory patients in Italy. BMC Health Serv Res 2005;5:26.
- 11. Factor SA, Molho ES. Emergency department presentations of patients with Parkinson's disease. Am J Emerg Med 2000;18: 209–215.
- Huse DM, Schulman K, Orsini L, Castelli-Haley J, Kennedy S, Lenhart G. Burden of illness in Parkinson's disease. Mov Disord 2005;20:1449–1454.
- 13. Guneysel O, Onultan O, Onur O. Parkinson's disease and the frequent reasons for emergency admission. Neuropsychiatr Dis Treat 2008;4:711–714.
- Parashos SA, Maraganore DM, O'Brien PC, Rocca WA. Medical services utilization and prognosis in Parkinson disease: a population-based study. Mayo Clin Proc 2002;77:918–925.
- Vossius C, Nilsen OB, Larsen JP. Parkinson's disease and hospital admissions: frequencies, diagnoses and costs. Acta Neurol Scand 2010;121:38–43.
- Klein C, Prokhorov T, Miniovitz A, Dobronevsky E, Rabey JM. Admission of Parkinsonian patients to a neurological ward in a community hospital. J Neural Transm 2009;116:1509–1512.
- Louis ED, Henchcliffe C, Bateman BT, Schumacher C. Youngonset Parkinson's disease: hospital utilization and medical comorbidity in a nationwide survey. Neuroepidemiology 2007;29:39–43.
- Temlett JA, Thompson PD. Reasons for admission to hospital for Parkinson's disease. Intern Med J 2006;36:524–526.
- 19. Woodford H, Walker R. Emergency hospital admissions in idiopathic Parkinson's disease. Mov Disord 2005;20:1104–1108.
- Guttman M, Slaughter PM, Theriault ME, DeBoer DP, Naylor CD. Parkinsonism in Ontario: comorbidity associated with hospitalization in a large cohort. Mov Disord 2004;19:49–53.
- Tan LC, Tan AK, Tjia HT. The profile of hospitalised patients with Parkinson's disease. Ann Acad Med Singapore 1998;27: 808–812.
- Kessler, II. Epidemiologic studies of Parkinson's disease. II. A hospital-based survey. Am J Epidemiol 1972;95:308–318.
- Pressley JC, Louis ED, Tang MX, et al. The impact of comorbid disease and injuries on resource use and expenditures in parkinsonism. Neurology 2003;60:87–93.
- 24. Guttman M, Slaughter PM, Theriault ME, DeBoer DP, Naylor CD. Burden of parkinsonism: a population-based study. Mov Disord 2003;18:313–319.
- 25. Pepper PV, Goldstein MK. Postoperative complications in Parkinson's disease. J Am Geriatr Soc 1999;47:967–972.
- Idjadi JA, Aharonoff GB, Su H, et al. Hip fracture outcomes in patients with Parkinson's disease. Am J Orthop 2005;34:341–346.

- Mueller MC, Juptner U, Wuellner U, et al. Parkinson's disease influences the perioperative risk profile in surgery. Langenbeck's archives of surgery/Deutsche Gesellschaft fur Chirurgie 2009;394:511–515.
- Jonsson B, Sernbo I, Johnell O. Rehabilitation of hip fracture patients with Parkinson's disease. Scand J Rehabil Med 1995;27: 227–230.
- 29. Mehta S, Vankleunen JP, Booth RE, Lotke PA, Lonner JH. Total knee arthroplasty in patients with Parkinson's disease: impact of early postoperative neurologic intervention. Am J Orthop 2008;37: 513–516.
- Londos E, Nilsson LT, Stromqvist B. Internal fixation of femoral neck fractures in Parkinson's disease. 32 patients followed for 2 years. Acta Orthop Scand 1989;60:682–685.
- Magdalinou KN, Martin A, Kessel B. Prescribing medications in Parkinson's disease (PD) patients during acute admissions to a District General Hospital. Parkinsonism Relat Disord 2007;13: 539–540.
- 32. Tanaka M, Suemaru K, Ikegawa Y, Tabuchi N, Araki H. Relationship between the risk of falling and drugs in an academic hospital. Yakugaku Zasshi 2008;128:1355–1361.
- Golden WE, Lavender RC, Metzer WS. Acute postoperative confusion and hallucinations in Parkinson disease. Ann Intern Med 1989;111:218–222.
- Severn AM. Parkinsonism and the anaesthetist. Br J Anaesth 1988; 61:761–770.
- Mason LJ, Cojocaru TT, Cole DJ. Surgical intervention and anesthetic management of the patient with Parkinson's disease. Int Anesthesiol Clin 1996;34:133–150.
- Harada T, Mitsuoka K, Kumagai R, et al. Clinical features of malignant syndrome in Parkinson's disease and related neurological disorders. Parkinsonism Relat Disord 2003;9(Suppl 1):S15–S23.
- Kalenka A, Hinkelbein J. Anaesthesia in patients with Parkinson's disease. Der Anaesthesist 2005;54:401–409; quiz 410–401.
- 38. Goldberg LI. Anesthetic management of patients treated with antihypertensive agents or levodopa. Anesth Analg 1972;51:625–632.
- Ong KC, Chew EL, Ong YY. Neuroleptic malignant syndrome without neuroleptics. Singapore Med J 2001;42:85–88.
- Derry CP, Shah KJ, Caie L, Counsell CE. Medication management in people with Parkinson's disease during surgical admissions. Postgrad Med J 2010;86:334–337.
- 41. Barber M, Stewart D, Grosset D, MacPhee G. Patient and carer perception of the management of Parkinson's disease after surgery. Age Ageing 2001;30:171–172.
- 42. Simon D, Shapira OM, Mor E, Pfefferman R. Parkinson syndrome. A significant risk factor in the patient with acute surgical disorder. Int Surg 1992;77:313–316.
- 43. Duffy GP, Trousdale RT. Total knee arthroplasty in patients with parkinson's disease. J Arthroplasty 1996;11:899–904.
- 44. Eventov I, Moreno M, Geller E, Tardiman R, Salama R. Hip fractures in patients with Parkinson's syndrome. J Trauma 1983;23: 98–101.
- 45. Staeheli JW, Frassica FJ, Sim FH. Prosthetic replacement of the femoral head for fracture of the femoral neck in patients who have Parkinson disease. J Bone Joint Surg Am 1988;70:565–568.
- Vince KG, Insall JN, Bannerman CE. Total knee arthroplasty in the patient with Parkinson's disease. J Bone Joint Surg Br 1989;71: 51–54.
- 47. Coughlin L, Templeton J. Hip fractures in patients with Parkinson's disease. Clin Orthop 1980;148:192–195.
- 48. Weber M, Cabanela ME, Sim FH, Frassica FJ, Harmsen WS. Total hip replacement in patients with Parkinson's disease. Int Orthop 2002;26:66–68.
- Rothermel JE, Garcia A. Treatment of hip fractures in patients with Parkinson's syndrome on levodopa therapy. J Bone Joint Surg Am 1972;54:1251–1254.
- Turcotte R, Godin C, Duchesne R, Jodoin A. Hip fractures and Parkinson's disease. A clinical review of 94 fractures treated surgically. Clin Orthop 1990;256:132–136.
- Pritchett JW, Bortel DT. Parkinson's disease and femoral neck fractures treated by hemiarthroplasty. Clin Orthop 1992;279: 310-311.

- 52. Tabamo RE, Fernandez HH, Friedman JH, Lucas PR. Spinal surgery for severe scoliosis in Parkinson's disease. Med Health R I 2000;83:114–115.
- Heckmann SM, Heckmann JG, Weber HP. Clinical outcomes of three Parkinson's disease patients treated with mandibular implant overdentures. Clin Oral Implants Res 2000;11:566–571.
- 54. Bryant C, Harley J. Re: Parkinson's disease patients who fracture their neck of femur: a review of outcome data. Injury 2007;38:1222.
- 55. Fast A, Mendelsohn E, Sosner J. Total knee arthroplasty in Parkinson's disease. Arch Phys Med Rehabil 1994;75:1269–1270.
- Kryzak TJ, Sperling JW, Schleck CD, Cofield RH. Total shoulder arthroplasty in patients with Parkinson's disease. J Shoulder Elbow Surg 2009;18:96–99.
- Babat LB, McLain RF, Bingaman W, Kalfas I, Young P, Rufo-Smith C. Spinal surgery in patients with Parkinson's disease: construct failure and progressive deformity. Spine (Phila Pa 1976) 2004;29:2006–2012.
- 58. Oni OO, Mackenney RP. Total knee replacement in patients with Parkinson's disease. J Bone Joint Surg Br 1985;67:424-425.
- 59. Clubb VJ, Clubb SE, Buckley S. Parkinson's disease patients who fracture their neck of femur: a review of outcome data. Injury 2006;37:929–934.
- 60. Koch LD, Cofield RH, Ahlskog JE. Total shoulder arthroplasty in patients with Parkinson's disease. J Shoulder Elbow Surg 1997;6: 24-28.
- Bohmdorfer W, Schwarzinger P, Binder S, Sporn P. Temporary suppression of tremor by remifentanil in a patient with Parkinson's disease during cataract extraction under local anesthesia. Anaesthesist 2003;52:795–797.
- 62. Clemente ML, Federici L, Brogi A, Civeli L, Giuntini L. Althesin -Fentanyl combination as anesthetic in patients with Parkinson's disease. Minerva Anestesiol 1980;46:675–680.
- 63. Stone DJ, DiFazio CA. Sedation for patients with Parkinson's disease undergoing ophthalmologic surgery. Anesthesiology 1988; 68:821.
- 64. Sehgal A, Panchal ID. Postoperative bronchospasm in a patient with Parkinson's disease. Can J Anaesth 2001;48:214–215.
- 65. Backus WW, Ward RR, Vitkun SA, Fitzgerald D, Askanazi J. Postextubation laryngeal spasm in an unanesthetized patient with Parkinson's disease. J Clin Anesth 1991;3:314–316.
- Oliveira E, Michel A, Smolley L. The pulmonary consultation in the perioperative management of patients with neurologic diseases. Neurol Clin 2004;22:277–291.
- 67. Burton DA, Nicholson G, Hall GM. Anaesthesia in elderly patients with neurodegenerative disorders: special considerations. Drugs Aging 2004;21:229–242.
- Furuya R, Hirai A, Andoh T, Kudoh I, Okumura F. Successful perioperative management of a patient with Parkinson's disease by enteral levodopa administration under propofol anesthesia. Anesthesiology 1998;89:261–263.
- 69. Broussolle E, Marion MH, Pollak P. Continuous subcutaneous apomorphine as replacement for levodopa in severe parkinsonian patients after surgery. Lancet 1992;340:859-860.
- Easdown LJ, Tessler MJ, Minuk J. Upper airway involvement in Parkinson's disease resulting in postoperative respiratory failure. Can J Anaesth 1995;42:344–347.
- Onofrj M, Thomas A. Acute akinesia in Parkinson disease. Neurology 2005;64:1162–1169.
- 72. Galvez-Jimenez N, Lang AE. The perioperative management of Parkinson's disease revisited. Neurol Clin 2004;22:367–377.
- 73. Nicholson G, Pereira AC, Hall GM. Parkinson's disease and anaesthesia. Br J Anaesth 2002;89:904–916.
- 74. Doyle SR, Kremer MJ. AANA journal course. Update for nurse anesthetists. Parkinson disease. AANA J 2003;71:229–234.
- 75. Brindle GF. Anesthesia in the patient with parkinsonism. Prim Care 1977;4:513–528.
- Rosin AJ, Devereux D, Eng N, Calne DB. Parkinsonism with 'onoff' phenomena. Intravenous treatment with levodopa after major abdominal surgery. Arch Neurol 1979;36:32–34.
- Liu EH, Choy J, Dhara SS. Persistent perioperative laryngospasm in a patient with Parkinson's disease. Can J Anaesth 1998;45(5 Pt 1):495.

- Galvez-Jimenez N, Lang AE. Perioperative problems in Parkinson's disease and their management: apomorphine with rectal domperidone. Can J Neurol Sci 1996;23:198–203.
- 79. Koganei H, Tanaka S, Iijima K, Horikawa Y, Tanaka O. Postoperative respiratory failure resulting from perioperative withdrawal of antiparkinsonian medications. Masui 1999;48:70–72.
- Bevan DR, Monks PS, Calne DB. Cardiovascular reactions to anaesthesia during treatment with levodopa. Anaesthesia 1973;28:29–31.
- 81. Wullner U, Kassubek J, Odin P, et al. Transdermal rotigotine for the perioperative management of Parkinson's disease. J Neural Transm 2010;117:855-859.
- 82. Korczyn AD, Reichmann H, Boroojerdi B, Hack HJ. Rotigotine transdermal system for perioperative administration. J Neural Transm 2007;114:219–221.
- 83. Merli GJ, Bell RD. Preoperative management of the surgical patient with neurologic disease. Med Clin North Am 1987;71:511–527.
- Sobolewski P. Treatment of Parkinson disease in patients with surgical problems. Neurologia i neurochirurgia polska 2003;37(Suppl 5):183–188.
- 85. Kalenka A, Schwarz A. Anaesthesia and Parkinson's disease: how to manage with new therapies?Curr Opin Anaesthesiol 2009;22: 419–424.
- Reed AP, Han DG. Intraoperative exacerbation of Parkinson's disease. Anesth Analg 1992;75:850–853.
- Ngai SH. Parkinsonism, levodopa, and anesthesia. Anesthesiology 1972;37:344–351.
- Horai T, Nishiyama T, Yamamoto H, Hanaoka K. High dose Ldopa infusion during general anesthesia for gastrectomy in a patient with parkinsonism. Masui 2002;51:42–45.

- Frucht SJ. Movement disorder emergencies in the perioperative period. Neurol Clin 2004;22:379–387.
- Spiegel J. Perioperative management of patients with Parkinson's disease. Orthopade 2009;38:843–846.
- Shipton EA, Roelofse JA. Anaesthesia in a patient with Parkinson's disease. A case report. South African medical journal = Suid-Afrikaanse tydskrif vir geneeskunde 1984;65:304–305.
- Fujii T, Nakabayashi T, Hashimoto S, Kuwano H. Successful perioperative management of patients with Parkinson's disease following gastrointestinal surgery: report of three cases. Surg Today 2009;39:807–810.
- 93. Mizuno J, Kato S, Watada M, Morita S. Perioperative management of a patient with Parkinson's disease with intravenous infusion of levodopa. Masui 2009;58:1286–1289.
- 94. Toledo LW. The postanesthesia patient with Parkinson's disease. J Post Anesth Nurs 1992;7:32–37.
- 95. O'Brien B, Harmon D. Severe postoperative nausea and vomiting in a parkinsonian patient. Hosp Med 2000;61:747.
- Zuckerman LM. Parkinson's disease and the orthopaedic patient. J Am Acad Orthop Surg 2009;17:48–55.
- Dautzenberg PL, Wouters CJ, Oudejans I, Samson MM. Rivastigmine in prevention of delirium in a 65 years old man with Parkinson's disease. Int J Geriatr Psychiatry 2003;18: 555–556.
- 98. Segatore M. Managing the surgical orthopaedic patient with Parkinson's disease. Orthop Nurs 1998;17:13–20; quiz 21–12.