### DISSERTATIONES MEDICINAE UNIVERSITATIS TARTUENSIS

# DISSERTATIONES MEDICINAE UNIVERSITATIS TARTUENSIS 136

# MANAGEMENT OF DEPRESSION IN FAMILY MEDICINE

### PILLE ÖÖPIK



Department of Polyclinic and Family Medicine, University of Tartu, Estonia

Dissertation is accepted for the commencement of the degree of Doctor of Medical Sciences on April 18, 2007 by the Council of the Faculty of Medicine, University of Tartu, Estonia

Supervised Professor Heidi-Ingrid Maaroos, Department of Polyclinic and

Family Medicine, University of Tartu, Estonia

Reviewed Professor Airi Värnik, Estonian-Swedish Institute of Mental

Health and Suicidology, Tallinn Estonia Karolinska Institute, Stockholm, Sweden

Professor Veiko Vasar, Department of Phychiatry,

University of Tartu, Estonia

Opponent: Professor Airi Värnik, Estonian-Swedish Institute of Mental

Health and Suicidology, Tallinn Estonia Karolinska Institute, Stockholm, Sweden

Professor Veiko Vasar, Department of Phychiatry,

University of Tartu, Estonia

Commencement: June 06, 2007

Publication of this dissertation is granted by the University of Tartu

ISSN 1024–395x ISBN 978–9949–11–603–4 (trükis) ISBN 978–9949–11–604–1 (PDF)

Autoriõigus Pille Ööpik, 2007

Tartu Ülikooli Kirjastus www.tyk.ee Tellimus nr. 161

To my family

### **CONTENTS**

1.	LIST OF ORIGINAL PUBLICATIONS	9
2.	ABBREVATIONS	10
3.	INTRODUCTION	11
4.	REVIEW OF THE LITERATURE	13
	4.1. Epidemiology of depression	13
	4.1.1. Prevalence of depression in population	13
	4.1.2. Prevalence of depression in primary care users	14
	4.1.3. Disability and depression	15
	4.2. Risk Factors for depression	16
	4.3. Screening of depression	17
	4.3.1. Characteristics of screening instruments	17
	4.3.2. Most common screening instruments	17
	4.4. Clinical forms and diagnosis of depression	18
	4.4.1. Classification of the clinical forms of depression	18
	4.4.2. Clinical symptoms of depression	19
	4.4.3. Composite International Diagnostic Interview	19
	4.5. Family doctors attitudes of depression	20
	4.6. Treatment of depression	20
	4.6.1. Treatment principles of depression	20
	4.6.2. Psychotherapy and other psychosocial treatments	21
	4.6.3. Common drugs in treatment of depression: antidepressants	21
5.	AIMS OF THE STUDY	22
	SUBJECTS AND METHODS	23
0.		23
	6.1. Study design	23 24
	6.2. Subjects	24
	6.2.1. Family doctors	
	6.2.2 Patients	25
	6.3. Methods	26
	6.3.1. Questionnaire for family doctors	26
	6.3.2. Patient interview with the Composite International	•
	Diagnostic Interview	26
	6.3.3. Questionnaire for assessment of sociodemographic and	~-
	health-related risk factors for depression	27
	6.3.4. Patient survey with the screening instrument EST-Q2	27
	6.3.5. Estonian Health Insurance Fund's data of drug prescriptions	
	for depression	28

	6.4.	Statistical methods
	6.5.	Ethics
7.	RES	SULTS
	7.1.	Factors influencing family doctors' readiness and motivation to manage with patients with depression symptoms
	7.2.	Family doctors' needs regarding problem solving in treatment of depression (motivating factors and problems related to treatment of depressive patients)
		Prevalence of depression in primary care in Estonia
		Factors associated with depression in primary care in Estonia
	7.5.	Suitability of the EST-Q2 screening scale depression subscale for
	7.6	screening depression in general practice
	7.0.	patients with depression from patients with other biomedical or
		psychosocial problems
	7.7.	Preferences and rationale of family doctors in pharmacological
	7.8.	treatment of depression
8.	DIS	CUSSION
9.	COl	NCLUSIONS
10	. RE	FERENCES
11	. AP	PENDICES
SU	JMM	IARY IN ESTONIAN
A	CKN	OWLEDGEMENTS
ΡĮ	IBLI	CATIONS

#### 1. LIST OF ORIGINAL PUBLICATIONS

- I Ööpik P, Aluoja A, Kalda R, Maaroos HI. Family doctors' problems and motivating factors in management of depression. BMC Fam Pract. 2006 Oct 30;7:64.
- II Aluoja A, Ööpik P, Kalda R, Maaroos HI. Prevalence of depression in primary care patients. Eesti Arst 2006; 85(12): 811–816 (in Estonian).
- III Ööpik P, Aluoja A, Kalda R, Maaroos HI. Screening for depression in primary care. Family Practice 2006; 23: 693–698.
- IV Ööpik P, Aluoja A, Kalda R, Maaroos HI. Treatment of depression in primary care. Eesti Arst 2005; 84(7): 481–487 (in Estonian).

#### 2. ABBREVIATIONS

BDI Beck Depression Inventory

CES-D Center for Epidemiologic Studies Depression Scale
CIDI Composite International Diagnostic Interview
DEPRES Depression Research in European Society

DSM-IV Statistical Manual of Mental Disorders. Fourth Edition

EST-Q Emotional State Questionnaire

EEK Emotsionaalne Enesetunde Küsimustik

FD Family Doctor

GDS Geriatric Depression Scale
GHQ General Health Questionnaire

ICD-10 International Classification of Diseases and Related Health

Problems, 10th Revision

ICPE International Consortium of Psychiatric Epidemiology Survey

LR Likelihood ratio

MINI Mini-International Neuropsychiatric Interview

NaSSA Noradrenergic and specific serotonergic antidepressants

NCS National Comorbidity Survey

NCS-R The National Comorbidity Survey Replication

NPV Negative predictive value

NSMHWB National Survey of Mental Health and Well-being

ODIN European Outcome of Depression International Network

PPV Positive predictive value

PRIME-MD The Primary Care Evaluation of Mental Disorders

PHQ Patient Health Questionnaire RIMA Monoamine oxidase inhibitor

SCAN Schedule of Clinical Assessment in Neuropsychiatry

SNRI Selective noradrenalin reuptake inhibitor SSRI Selective serotonin reuptake inhibitor

TCA Tricyclic antidepressant
WHO World Health Organization

#### 3. INTRODUCTION

There are several events in life, which make people seek help from a doctor because of problems and diseases appearing or exacerbating in different periods. Patients themselves can not distinguish what has made them feel worse and what kind of help they need. Approximately 20–40 patients attend my consultations daily but up to 10-20% of them have complaints that can be caused by mental problems and can also be symptoms of depression. Also, the results of a study performed in 1996–1997 indicate that 11.1% of the Estonian population suffer from significant depressive symptoms (Aluoja et al 2004). The author has increasingly diagnosed depression in her patients and prescribed antidepressants for treatment. Diagnosing depression and prescribing the right cure is not simple in family doctors' daily work. Some symptoms of depression can be confused with those of other medical conditions. For example, weight loss and fatigue may be associated with disorders such as diabetes, cancer, and thyroid disease. Other researchers have pointed out the same difficulty as well (Whooley et al 2000). Often, people with depression do not realize that their feelings are due to a medical condition and hence they do not seek medical care for depression. The proportion of patients with depression who reported only somatic symptoms was 45-95% (Simon et al 1999). However, depression is a medical condition, not a normal reaction to a life situation such as the death of a loved one or the loss of job.

A large number of persons suffering from depression do not evaluate their emotional health as poor, or they simply do not want to talk about the problem. Patients who rated their emotional health poorer were significantly more likely to discuss depression with their physicians. Female patients were almost 3.5 times more likely to bring up depression before their physicians did than male patients. More educated patients were significantly more likely to bring up depression before their physicians did during a medical visit (Sleath *et al* 2002).

At the same time, dealing with the patients with depression symptoms takes from me and other family practitioners more consultation time compared with other patients (Tähepõld *et al* 2003, Sleath *et al* 2002). Without additional easily applicable diagnostic means making of a correct diagnosis is complicated in primary health care. Because there are no laboratory tests for depression and no biological markers that can be routinely measured, the diagnosis of depression is made using a number of reliable depression scales and questionnaires that can help the physician rapidly identify symptoms of depression and assist in prescribing appropriate treatment. In addition, the primary care physician can explain the biochemical nature of depression and reassure the patient that the symptoms are not due to an inherent personality "weakness" (Freguson 2000). There is little information about which screening questions work best.

The above described practical need gave me the idea and stimulated me to investigate more profoundly the most important symptoms of depression and

the possibilities of identifying patients who need more thorough diagnostics in family practitioners' daily work. I was able to satisfy my interest owing to the fact that the PREDICT investigation had been initiated in Estonia which a great number of patients were enrolled. They passed the Composite International Diagnostic Interview and were asked about several risk factors of depression. I am glad to have made a contribution to the activity of this working group by adding my survey to their research.

#### 4. REVIEW OF THE LITERATURE

#### 4.1. Epidemiology of depression

#### 4.1.1. Prevalence of depression in population

Depression is one of the most prevalent psychiatric conditions in the community. Several surveys have been performed to find out the prevalence of depression among population, among them international surveys in different countries and continents (Lepine *et al* 1997, Ayuso-Mateos *et al* 2002, Andrade *et al* 2003). Other surveys have focused on the prevalence of depression in one country only (Blazer *et al* 1994, Kessler *et al* 2003, Wilhelm *et al* 2003, Patten *et al* 2006). The results from several studies demonstrate that lifetime prevalence estimates of depression varied widely, from 3% to 17.1%, while the point prevalence of major depression varied from 0.9% to 15.1% (Table 1).

Table 1. Prevalence of depression in community on the basis of epidemiological studies

Country	Study	Instrument	Reference	Major	Lifetime
-		(diagnostic		depressive	prevalence
		criteria)		episodes %	%
Finland Urban	ODIN	SCAN (DSM-	Ayuso-	4.7 *	
Rural		III/IV, ICD-	Mateos et al	4.1 *	
Ireland Urban		10)	2002	15.1 *	
Rural		,		5.5 *	
Norway Urban				7.0 *	
Rural				8.48 *	
Spain Urban				1.8 *	
UK Urban				7.8 *	
Rural				6.1 *	
Canada		CIDI	Patten et al	1.8 *	12.2
		(CCHS1.2)	2006		
US	NCS	CIDI (DSM-	Blazer et al	4.9 *	17.1
		III-R, ICD-10)	1994		
US	NCS-R	CIDI (DSM-	Kessler et al		16.2
		IV)	2003		
Australia	NSMHWB	CIDI (DSM-	Wilhelm et	3.2 *	
		IV, ICD-10)	al 2003		
Belgium	DEPRES	MINI (DSM-	Lepine et al	5.0 **	
France		III)	1997	9.1 **	
Germany		ŕ		3.8 **	
Netherlands				6.9 **	
Spain				6.2 **	
UK				9.9 **	

Country	Study	Instrument	Reference	Major	Lifetime
		(diagnostic		depressive	prevalence
		criteria)		episodes %	%
Brazil	ICPE	CIDI (DSM-	Andrade et	3.9 *	12.6
Canada		III-R, for	al 2003	1.9 *	8.3
Chile		Germany and		3.3 *	9.0
Czech Republic		Czech		1.0 *	7.8
Germany		Republic		1.3 *	11.5
Japan		DSM-IV)		0.9 *	3.0
Mexico				2.2 *	8.1
Netherlands				2.7 *	15.8
Turkey				3.1 *	6.3
US				4.6 *	16.9

<sup>\*</sup> point prevalence

Table 1 shows the difference between the estimates of prevalence of depression in rural and urban areas. The prevalence of depression depends upon the instrument and on the diagnostic criteria used. The prevalence of major depressive episode as diagnosed in Germany using two different instruments ranged from 1.3% to 3.8%; similar figures have been reported from the UK. An epidemiological study performed in the US found a difference in the prevalence of major depressive disorder for different racial/ethnic groups (Blazer *et al* 1994). An Estonian survey revealed that 11.1% of the population had symptoms of depression (Aluoja *et al* 2004).

#### 4.1.2. Prevalence of depression in primary care users

Depression is very common in primary care settings, as the prevalence rate of major depression has ranged from 1.5 to 27.3 (Simon *et al* 1999, Spizer *et al* 1999, Henkel *et al* 2004) (Table 2). Depressive disorders or significant depressive symptoms have been found in up to 55% of patients visiting general or family practitioners (Spizer *et al* 1999, Henkel *et al* 2004, Simon *et al* 2004) (Table 2).

<sup>\*\* 6-</sup>month prevalence

**Table 2.** Prevalence of depression in primary care

Country	Instrument	Reference	Major	Depressive
	(diagnostic		depressive	disorder %
	criteria)		episodes %	
Spain (Barzelona)	CES-D	Simon et al		31 *
Israel (Be'er Sheva)		2004		24 *
Australia (Melbourne)				52 *
Brazil (Porto Alegre)				52 *
Russia (St. Petersburg)				55 *
US (Seattle)				34 *
Germany	CIDI	Henkel et	10.2 **	18.3 **
	(DSM-IV)	al 2004		
Turkey (Ankara)	CIDI	Simon et al	10.8 **	
Greece (Athens)	(DSM-IV)	1999	7.1 **	
India (Bangalore)			8.5 **	
Germany (Berlin)			5.3 **	
Netherlands (Groningen)			14.4 **	
Nigeria (Ibadan)			4.1 **	
Germany (Mainz)			10.0 **	
UK (Manchester)			17.1 **	
Japan (Nagasaki)			1.5 **	
France (Paris)			13.6 **	
Brazil (Rio de Janeiro)			18.3 **	
Chile (Santiago)			27.3 **	
US (Seattle)			6.4 **	
China (Shanghai)			2.4 **	
Italy (Verona)			4.6 **	
US	PRIME-	Spizer et al	10 ***	16 ***
	MD PHQ	1999		

<sup>\*</sup> past week

The prevalence of depressive disorder is higher among primary care users than in population. This is expected because persons who have not health complaints not visit their family doctors (FDs).

#### 4.1.3. Disability and depression

Disability is associated with major depression. It has been found that the rates of mental disability were the highest in middle age and the rates of physical disability increased with age (Wilhelm *et al* 2003). Severity of depression and medical comorbidity are associated with longer absence from work (Chisholm

<sup>\*\*</sup> point prevalence

<sup>\*\*\*</sup> previous 2 weeks

et al 2003). By the year 2020, depression is prognosticated rank to second in disability calculated for all ages and for both sexes (World Health Organization (WHO)).

#### 4.2. Risk factors for depression

Risk factors for depression may be conceptualized as being either intrinsic to the individual, or are residing within the social environment.

The consensus reached in most community-based epidemiological studies is the following: women are at greater risk for major depression than men; persons of a lower socioeconomic status are at greater risk than those who are more well-off economically; younger persons are at greater risk for major depression than older persons; persons who are separated or divorced show higher rates of major depression than persons who are married or have never been married; blacks are somewhat less at risk than whites (Patten et al 2006, Wilhelm et al 2003, Blazer et al 1994, 1998, Andrade et al 2003, Aluoja et al 2002, Williams et al 2007). Married people displayed the lowest prevalence, but the effect of marital status changed with age, the annual prevalence may increase with age in men who have never been married (Patten et al 2006). Female persons were found to have higher risk for minor depression (Blazer et al 1998). The prevalence of major depression was related to having chronic medical condition, and to unemployment (Patten et al 2006, Dutton et al 2004). Poverty and unemployment were associated with longer episodes of common mental disorders (Weich et al 1998). Medical chronic illness had the strongest association with depression (Wilhelm et al 2003). Strong association was found between depression and medical or physical comorbidity (Chisholm et al 2003). Depression is also associated with chronic physical illness and with relationship or financial difficulties, and sufferers are pessimistic about recovery (Tylee et al 1999). Up to half of the patients were categorized as currently depressed (Tylee 2000). Smoking cigarettes was more highly correlated with current major depression than drinking alcohol (Wilhelm et al 2003).

The consensus from most community-based epidemiological studies is that persons in urban areas are at greater risk for major depression than persons in rural areas (Patten *et al* 2006, Wilhelm *et al* 2003, Blazer *et al* 1994, 1998, Andrade *et al* 2003, Aluoja *et al* 2002). According to the NCS carried out among persons with major depression, male gender and older age were associated with higher prevalence with a seasonal pattern (Blazer *et al* 1998). The prevalence of major depression was related to global health-related quality of life (Herman *et al* 2002) and negative life events (Salokangas et al 1998). Strong association was found between depression and medical or physical comorbidity (Chisholm *et al* 2003).

#### 4.3. Screening of depression

#### 4.3.1. Characteristics of screening instruments

In evaluation of screening instruments, sensitivity, specificity, false negative and positive rates, positive and negative predictive values, and the likelihood ratio are important. Sensitivity is the chance of detecting a disease when present; specificity is the chance of ruling out a disease when absent. Predictive values indicate the relative frequency of a predictor being correct (Riffenburgh 1999). The likelihood ratio (LR) is a way to incorporate sensitivity and specificity of the test into a single measure. The LR shows how much we should shift our suspicion in a particular test result. A positive LR (sensitivy/1-specificity) indicates how much we have to increase the probability of a disease if the test result is positive. A negative LR (1-sensitivity/specificity) reflects how much we have to decrease the probability of a disease if test result is negative (Biggerstaff 2000).

#### 4.3.2. Most common screening instruments

Self-rate instruments vary in the number of symptoms, duration of symptoms and the scale of evaluation. Although each measure has a unique scoring system, higher scores consistently reflect more severe symptoms (Sharp et al 2002). For screening of depression, there are a number of different instruments (Table 3).

Table 3. Screening Measures for Depression

Measure	Number	Completion
	of items	time
		(approximate
		minutes)
Beck Depression Inventory (BDI) (Beck 1961)	21	5 to 10
Beck Depression Inventory-II (Beck et al 1996)	21	5 to 10
Beck Depression Inventory-PC (BDI-PC) (Beck et al	7	Less than 5
1997)		
Center for Epidemiological Studies Depression (CES-D)	20	5 to 10
(Radolff et al 1977)		
Center for Epidemiological Studies-Depression Scale for	20	5 to 10
Children (CES-DC) (Fendrich et al 1990)		
Zung Self-Rating Depression Scale (Zung 1965)	20	5 to 10
Geriatric Depression Scale (GDS) (Yesavage et al 1983)	30	10 to 15
Geriatric Depression Scale-short (Sheik et al 1986)	15	5 to 10
General Health Questionnaire (GHQ) (Goldberg)	12	5 to 10
Patient Health Questionnaire-9 (PHQ-9)	9	5 to 10

5 17

Different screening instruments differ in the number of items, symptoms and duration symptoms. Is has been attempted to screen depression with two questions about depressed mood (Arroll et al 2003). This method has high sensitivity accompanied by a high number of false positive results. The shortcoming of different screening instruments is occurrence of false positive and false negative results. There is no evidence that one method of screening for depression works better than another (Task Force 2002). Depression screening measures do not diagnose depression, but they provide an indication of severity within a given period of time (Sharp et al 2002). Depression occurs in children, adolescents, adults, and the elderly. The most commonly used screening measures for adults in primary care settings include the Beck Depression Inventory, the Zung Self-Depression Scale, the General Health Questionnaire (GHQ) and the Patient Health Questionnaire-9 (PHQ-9). The GDS 30- and 15-item versions for the elderly are the most highly evaluated screening instruments. Screening measures for children and adolescents have been primarily used for research, and their clinical use in primary care settings has not been established (Sharp et al 2002).

Screening instruments for different age groups are different because the main symptoms vary with age. A significant effect was exerted by ethnicity, with contributions from the items agitation and irritability (Carmody 2005). Women showed higher depression scores than men and women showed also higher scores than men for a number of items, as sadness, crying, energy and fatigue (Carmody 2005, Endler *et al* 1999).

#### 4.4. Clinical forms and diagnosis of depression

#### 4.4.1. Clinical symptoms of depression

Depression is mostly thought to be a steady feeling of sadness which is accompanied by other symptoms. The symptoms of depression as a disease are the following: depressed mood, loss of interest in nearly all activities, loss of capacity for enjoyment, insomnia or hypersomnia, waking in the morning several hours before usual time, depressed mood with a particular expression in the morning, reduced self-esteem and self-confidence, feeling of worthlessness or excessive guilt, fatigue or loss of energy, marked tiredness after even a minimum effort, diminished ability to think or concentrate, substantial change in appetite or weight, psychomotor agitation or retardation, recurrent thoughts of death or suicide and loss of libido.

#### 4.4.2. Classification of the clinical forms of depression

The most important bases of classifying mood disorders are nature of symptoms, severity of symptoms and duration of symptoms. Two different classifications are used in diagnosing depression. These are Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition (DSM-IV) and International Classification of Diseases, 10th Revision (ICD-10). The DSM-IV diagnostic categories for depression are: minor depression, dystymia, major depression. The diagnostic categories of ICD-10 for depression are: mild depressive episode, moderate depressive episode, severe depressive episode with psychotic symptoms and other depressive episode (atypical depression). Major depression according to DSM-IV corresponds to moderate depressive episode or severe depressive episode according to ICD-10.

**Table 4.** Definition of major depression by DSM-IV and ICD-10

Major depression	Criteria	Duration
DSM-IV	≥ 5 depressive symptoms, including depressed mood or loss of interest in nearly all activities	≥ 2 weeks
ICD-10	≥ 4 depressive symptoms, including two of the symptoms (lowering of mood, reduction in energy, and decrease in activity)	≥ 2 weeks

Diagnosis of major depression according to DSM-IV does not differ from diagnosis according to ICD-10, although in the former care diagnosis of depression requires more than one positive symptom.

The diagnostic categories of depression depend upon the number and severity of the symptoms. Higher depressive symptom scores in primary care patients were consistently associated with poorer health, functional status, global health-related quality of life, and increased use of health care, but not with demographic variables (Herman *et al* 2002). Treatment of patients depends mostly on the diagnostics according to classification.

#### 4.4.3. Composite International Diagnostic Interview

The Composite International Diagnostic Interview (CIDI) is a fully structured and standardized non-clinical psychiatric interview designed for use in general population surveys. The Interview is designed to assess major mental disorders including unipolar depression, bipolar disorder, panic disorder, social phobia and alcohol and drug dependence (WHO).

#### 4.5. Family doctors attitudes towards depression

Only a few studies have been conducted on the family doctors' opinions about depression related problems in their work.

General practitioners have reported that the main obstacles to providing a good service for people with depression included not having enough time, too much work lack of services to refer to and difficulty in accessing services. The factors that influenced general practitioners to refer patients with depression to other services were risk to the patient, a clear need for specialist treatment and the need for assessment (Telford *et* al 2002, Ralition *et al* 1999). The ability of FD to diagnose and treat depression is directly related to their knowledge and further training (Rutz 2001). Most general practitioners believed that they could diagnose depression, almost half, of them had previously participated in a depression-related continuing medical education and the level of interest in psychiatry was significantly correlated with the treatment behaviour of general practitioners (Soykan *et al* 2003).

#### 4.6. Treatment of depression

For treatment of depression use is made of pharmacotherapy and other somatic treatments, as well as psychotherapy and other psychosocial treatments. Often combined therapy is applied.

#### 4.6.1. Treatment principles of depression

Family doctors' patients with major depression in different countries received antidepressant pharmacotherapy ranging from low (0%) in St. Petersburg to high (93%) in Australia (Wilson et al 2003, Simon et al 2004). The probably of treatment may be more influenced by the characteristics of a health care systems than by the clinical characteristics of individual patients (Simon et al 2004). Among FDs and psychiatrists were similar drug treatment prescription most often included antidepressants, while FDs often prescribed SSRIs (Wilson et al 2003, Ernst et al 2006). Compared with FDs, psychiatrists prescribed more often tricyclic and very novel antidepressants with longer duration, antipsychotics as well as mood stabilizers; also their patients received more psychotherapy (Tardieu et al 2006). Many clinicians reported their preference for an initial treatment that combined medication and psychotherapy as opposed to either modality alone (Kornbluh et al 2001). Study of trends in the rate of treatment during ten years data from the NCS show that the rate of treatment increased more in general medical service than in the psychiatric services. Trends in the rate of treatment were similar in two respects: severity of a disorder was significantly related to rate of treatment, and this

association did not change significantly over time (Kessler *et al* 2005). Also in Spain it was found that similar proportions of patients in specialist care and general medical care received minimally adequate treatment (Fernandez *et al* 2006). Most patients were satisfied with the care that they received from their primary care physician and approximately 65% of patients considered their physician's knowledge of depression and treatment to be excellent or very good (Schwenk *et al* 2004).

#### 4.6.2. Psychotherapy and other psychosocial treatments

The clinical practice guidelines contained in *Depression in Primary Care* (Clinical Practice Guidelines) recommend that psychotherapy and patient education should be considered when treating patients with major depressive disorder. Psychotherapy can take many forms, including cognitive therapy, behavioural therapy, and interpersonal therapy. A few studies have reported the efficacy of psychosocial treatment approaches, including problem-solving treatment (Dowrick *et al* 2000, Mynors-Wallis *et al* 2000), group psycho education (Dowrick *et al* 2000), and the cognitive behavioural analysis system of psychotherapy (Keller *et al* 2000). Psychological therapy was more effective treatment for depression than usual general practitioner's care in short term (Ward *et al* 2000). Generic counselling seems to be as effective as anti-depressant treatment for mild to moderate depressive illness (Chilvers *et al* 2001). A combination of an active drug and simple psychological treatment was more effective than simple psychological treatment alone (Malt *et al* 1999).

#### 4.6.3. Common drugs in treatment of depression: antidepressants

The factors to be considered when choosing an antidepressant include the spectrum of adverse effects, long-term tolerability, dosing schedule, clinically significant drug interactions, underlying medical conditions, earlier response to therapy, and medicine-economics (Cohen 1997). More recent antidepressants are clearly effective in treating depressive disorders in diverse settings (Malt et al 1999, Thase 1999, Williams et al 2000, Petersen et al 2002). Most FDs and psychiatrists indicated SSRIs as their first-line treatment preference as they have fewer side effects (Petersen et al 2002, Dording et al 2002, Wilson et al 2003). Patients who had the largest number of symptoms were more likely to be taking antidepressants compared with other patients (Tylee 2000). The most common barrier to receiving treatment was concern about costs and about the adverse effects of a medication (Simon et al 2004). Ample evidence shows that treating depression with counselling, medications, or both improves patient outcomes (US Task Force 2002). Combining pharmacotherapy and psychotherapy can be more effective than use of either modality alone (Mynors-Wallis et al 2000, Sutherland et al 2003).

#### 5. AIMS OF THE STUDY

- 1. To find out the family doctors' readiness, motivation, problems and needs in management of patients with depression symptoms (Paper I).
- 2. To estimate the prevalence of depression in family practice (Paper II).
- 3. To investigate the relationship of depression with some sociodemographic factors, life events and general health of patients (Paper II).
- 4. To establish the suitability of the EST-Q2 depression subscale for screening of depression in general practice (Paper III).
- 5. To find out a combination of symptoms allowing family doctors to distinguish patients with depression from patients with other biomedical or psychosocial problems (Paper III).
- 6. To study the preferences and rationale of family doctors in pharmacological treatment of depression (Paper IV).
- 7. To examine antidepressant prescribing patterns among family doctors (Paper IV).

### **6. SUBJECTS AND METHODS**

### 6.1. Study design

Overview of the study designs, subjects and methods is presented in table 5.

Table 5. Study design, subjects and methods

Aim of the study	Study design	Study subjects	Methods	Paper
To find out the FDs' readiness, motivation, problems and needs in management of	cross- sectional study	FDs	questionnaire	I
patients with depression symptoms				
To estimate the prevalence of depression in family practice	cross- sectional study	FDs' consecutive patients aged 18 to 75, who were recruited in the study (sample I)	face-to-face structured interview CIDI	II
To investigate the relationship of depression with some sociodemographic factors, life events and general health of patients	cross- sectional study	FDs' consecutive patients aged 18 to 75, who were recruited in the study (sample I)	questionnaire	П
To establish the suitability of the EST-Q2 screening scale depression subscale for screening of depression in general practice	cross- sectional study	FDs' consecutive patients aged 18 to 75, who were recruited in the study (sample II)	face-to-face structured interview CIDI and self-rate instrument EST-Q2	III
To find out a combination of symptoms allowing FDs to distinguish patients with depression from patients with other biomedical or psychosocial problems	cross- sectional study	FDs' consecutive patients, aged 18 to 75, who were recruited in the study (sample II)	face-to-face structured interview CIDI and self-rate instrument EST-Q2	III

Aim of the study	Study design	Study subjects	Methods	Paper
To study the	cross-	family doctors	questionnaire	IV
preferences and	sectional			
rationale of FDs in	study			
pharmacological				
treatment of				
depression				
To examine	retrospective	family doctors	data of the Estonian	IV
antidepressant	analysis	and	Health Insurance	
prescribing patterns	study	psychiatrists	Fund for 2003	
among psychiatrists	_			
and FDs				

#### 6.2. Subjects

#### 6.2.1. Family doctors

Questionnaires were sent by post to 500 (89% of all FDs) certified practicing FDs in October 2005. Two hundred and five questionnaires were returned. Of the respondents 84(41%) worked in rural and 121(59%) worked in urban areas. The background of the FDs who received the questionnaire is presented in table 6.

Table 6. Background characteristics of the respondents

Location	Solo	Group	Average	Average length	Average length
	practice n	practice n	age, years	of service as a	of service as a
	(%)	(%)	(±SD)	physician, years	FD, years (±SD)
				(±SD)	
Rural	63(75)	21(25)	45.9(±8.4)	19.1(±7.7)	5.1(±1.3)
(n=84)					
Urban	35(29)	86(71)	45.7(±8.5)	19.5(±9.2)	4.5(±2)
(n=121)					
Total	98(48)	107(52)	45.8(±8.5)	19.4(±8.6)	4.8(±1.5)
(n=205)	, í	, í	, , ,		

Practising FDs in Estonia and the FDs participating in this study were similar regarding the characteristics of gender and location of practice (Paper I Table 2).

#### 6.2.2. Patients

The sample was recruited from April to June 2003 by 23 family doctors (15 from urban and 8 from rural areas) who had shown interest in participating in the study. The recruitment of patients and the design of the study were carried out according to the PREDICT project (King *et al* 2006). The FDs were specially instructed to recruit patients proceeding from the project criteria.

The inclusion criteria were:

- 1) consecutive attendees of FDs' consultations
- 2) patients from urban and rural areas
- 3) patients aged 18 to 75 years The exclusion criteria were:
- 1) non-Estonian speakers
- 2) presence of a severe organic mental illness
- 3) presence of a terminal illness

After the participants had given their informed consent, a subsequent detailed interview was carried out either at their home or at general practices within two weeks. Then the patients completed the EST-Q2 and a questionnaire for assessment of sociodemographic and health-related risk factors of depression on their own and the interviewers administered the CIDI. The FDs invited 1370 patients, of whom 1175 agreed to take part in the study. A total of 1100 interviews were completed as 75 patients could not be contacted or had changed their mind about participation. Further, 6 interviews of 1100 were excluded due to the incomplete data of the questionnaire for assessment of sociodemographic and health-related risk factors of depression and 42 interviews of 1100 were excluded due to the incomplete data of EST-Q2. Study sample I for investigation of the prevalence of depression in family practice and the relationship of depression with some sociodemographic factors, life events and general health of patients consisted of 1094 persons. Study sample II for establishment of the suitability of the EST-Q2 and for finding out a combination of symptoms for screening of depression in family practice best consisted of 1058 persons: 776(73%) women (mean age  $40.5\pm15.4$ ) and 282(37%) men (mean age  $42.7\pm$ 16.2). The formation and characteristics of the samples is shown in figure 1.

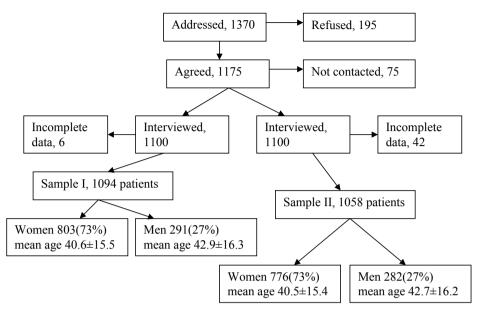


Figure 1. Sample characteristics

#### 6.3. Methods

#### 6.3.1. Questionnaire for family doctors

The questionnaire was compiled and tested by the author of this thesis Pille Ööpik. The tailor-made questionnaire included both closed and open questions. The closed questions required yes/no answers and the open questions required a description of an opinion. The questionnaire included questions about the FDs' background (location of the practice, solo or group practice, age, gender, length of service). The questionnaire consisted of 12 questions (Appendix 1). A questionnaire-based survey was conducted from October to November 2002.

### 6.3.2. Patient interview with the Composite International Diagnostic Interview

The Composite International Diagnostic Interview (CIDI) was selected for comparison because the reliability and validity of this instrument has been established. The CIDI is a fully structured diagnostic interview providing current (and lifetime) psychiatric diagnoses according to ICD-10 and DSM-IV, which was developed by the World Health Organization. A depressive episode

was established using the Depression Section of CIDI. In this study we used one-month depression determined according to the criteria of ICD-10. The interviews were carried out by trained instructed interviewers. The interview was carried out after the participants had given their informed consent, at their home or at the general practice, within two weeks.

### 6.3.3. Questionnaire for assessment of sociodemographic and health-related risk factors for depression

Selection of presumed risk factors was based on previous research (Anderson *et al* 1993, Weich *et al* 1997) and on a systematic review of the literature by the work group of PREDICT. Where possible, they used published self-reported measures of established reliability and validity. In some cases, questions were developed for the particular study or adapted from available standardised instruments. The work group PREDICT addressed the risk factors that are intrinsic either to the individual or to the social context, while remaining aware that there is inevitable overlap in such categorisation. A set of risk factors was established specifically for the PREDICT study and was tested for reliability (King *et al* 2006).

In this study we used a questionnaire of socio-demographic factors, economic coping, life events and self-rated of problems physical health.

#### 6.3.4. Patient survey with the screening instrument EST-Q2

A new modification of EST-Q consisted of 32 items, which performs well in psychiatric patients and general population, was developed in 2002 (Aluoja *et al* 1999). The items, which did not belong to any subscale, were omitted. The EST-Q2 contained the subscales of Depression, Anxiety, Agoraphobia-Panic, Fatigue and Insomnia, reflecting the symptoms of depressive and anxiety disorders according to ICD-10 and DSM-IV. Each item was rated on a 5-point scale ranging from 0 to 4 (0 = not at all; 1 = seldom; 2 = sometimes; 3 = often; 4 = all the time). The participants were asked to report how much the various problems had troubled them during the past four weeks, using the scale. The EST-Q2 version consisted of 28 items, the Depression subscale consisted of 8 items encompassing cognitive and affective symptoms of depression. The cutoff point for depression was >11 (Appendix 2).

### 6.3.5. Estonian Health Insurance Fund's data of drug prescriptions for depression

First, data were inquired from the Estonian Health Insurance Fund about the medicines for treatment of depression prescribed by FDs and psychiatrists according to the prescriptions sent to the Estonian Health Insurance Fund (by pharmacies). As the Estonian Health Insurance Fund receives information only about the medicines compensated by them, it was possible to obtain data about antidepressants (ANDP), antipsychotics (ANPS) and mood stabilizers (MST).

Second, data were inquired about the frequency of new diagnoses of depression made by FDs and psychiatrists according to the treatment invoices sent to the Estonian Health Insurance Fund (by doctors). All data were inquired as of 2003. The data were drawn for the diagnoses with codes F32-F33 according to the criteria of ICD-10.

#### 6.4. Statistical methods

The results of the questionnaire for FDs were analysed with the use of frequency distribution tables. The differences between the groups were tested using the Chi-square test. The open questions were analysed using thematic analysis. All answers to the open-ended questions were recorded. The subsequent statements were first analysed by the first author Pille Ööpik. For identifying any statements, related to the FDs' motivation to deal with depressive patients, and any problems arising during work with depressive patients, all statements expressing motivation for, or indicating problems with working with depressive patients were coded and categorized according to their content and the categories were labelled in order to verify that the described findings reflected the database adequately.

A depressive episode for sample I was assessed using the Depression Section of CIDI, which provides present, six-month and lifetime psychiatric diagnoses according to ICD-10. In these study was assessed present and six-month depression. Analyses were performed with the software package SPSS for Windows 10.0.

Relationship between depression and background factors was estimated by  $\chi^2$ -test. When calculating the odds-ratios we took as a reference category the level of background variable to which correspond the lowest rate of depression.

Two-by-two tables were constructed for sample II, displaying screening instrument (EST-Q2) diagnosis (positive/negative) versus CIDI diagnosis (positive/negative). Sensitivity, specificity, false negative and false positive rates, and positive and negative predictive values were calculated to assess the ability of the screening instruments to render the diagnosis of depression according to CIDI. Further, the positive and negative likelihood ratios of the test

were assessed. The likelihood ratio (LR) is a way to incorporate the sensitivity and the specificity of the test into a single measure. The LR indicates how much we should shift suspicion in the case of a particular test result. A positive LR (sensitivy/1-specificity) indicates how much we have to increase the probability of the disease if the test result is positive. A negative LR (1-sensitivity/ specificity) reflects how much we have to decrease the probability of the disease if the test result is negative. Stepwise logistic regression was used to find out the best combination of symptoms for screening depression. We developed two regression models. In Model 1 we used the symptoms of the EST-O2 Depression scale as the predictors of CIDI-diagnosed depression. In Model 2 we added to the predictor variables the somatic and the behavioural symptoms of EST-Q2 depression, plus one anxiety symptom, worrying too much. According to our assessment, the scale in which all arguments were statistically significant (p<0.05) was the best. Different cut-off points were used to compare the scales. All analyses were performed with the software package SAS 8.1.

To find out preference for medicines the numbers for the reported preferences were summarized. For analysing preference for medicines, all sentences about the preference were marked and similar preferences were further coded and categorized according to their content. The second author analysed the texts independently in a similar way. A few ambiguities in the analyses were discussed to reach consensus.

The data of the prescribed medicines were analysed with the use of frequency distribution tables. Statistical significance was tested using the Chisquare test. The level of statistical significance was set at  $p \le 0.05$ .

#### 6.5. Ethics

The Committee of Ethics of the University of Tartu has approved the study protocol and the form of informed consent of the study.

#### 7. RESULTS

### 7.1. Factors influencing FDs' readiness and motivation to manage with patients with depression symptoms

Of the family doctors 185(90%) considered dealing with depressive patients being within their competence, while 20(10%) did not. The latter were of the opinion that the problems of such patients were only the psychiatrists' responsibility and that psychiatrists had more time for them than FDs. Of the respondents 180(98%) were willing to deal with depressive persons and 200(88%) had to deal with depression. The opinions of the FDs about depression management were not significantly different depending on the location and type of the practice (p>0.05) (Paper II, Table 3).

A large number of FDs 150(73%) use screening tests in the case suspected depression. The most frequently mentioned test was EST-Q2, while the other tests were mentioned less often (Figure 2).

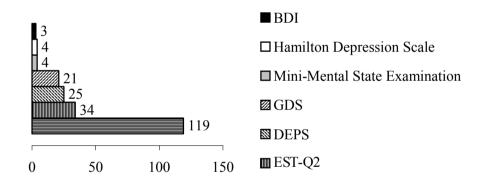


Figure 2. The screening instruments used by family doctors

The FDs pointed out many motivation factors for management of patients with depression symptoms. The motivation factors for the FDs were grouped into five topics according to the content (Table 7).

**Table 7.** Motivation factors for the family doctors

Category	Example of the FDs statement						
High prevalence of	"Depression is widespread."						
depression in primary health	"There is a great need for depression treatment."						
care	"Depression often accompanies the main disease."						
FDs' feeling of commitment	"I hope I can help patients."						
	"We cannot be dispatchers sending people to various						
	places."						
	"Patients refuse to see a psychiatrist; FDs have to manage						
	on their own."						
Positive results of treatment	"After effective treatment the patient seems reborn."						
	"Earlier positive experience in depression management."						
	"If patients receive help, further co-operation will be						
	good."						
	"If treatment is effective, the patient will not demand						
	clinical investigations any more."						
	"Several somatic complaints disappear during the						
	treatment of depression."						
	"Patient's recovery gives much satisfaction."						
FDs' advantages	"Patients' trust is important."						
	"We know our patients better than psychiatrists do."						
	"FDs are better informed of concomitant diseases."						
	"It is much easier for the patient to consult the FD."						
Convenience from the	"A bedridden patient at home cannot go anywhere else."						
patient's point of view	"Patients do not want to see the psychiatrist. Psychiatrists'						
	offices are located far from the patients' homes."						
	"Specialists' waiting lists are long."						

# 7.2. Family doctors' needs regarding problem solving in treatment of depression

The problems that the FDs described regarding management of depressive patients' were grouped into four categories (Table 8).

 Table 8. Problems of family doctors

Category	Example of FDs` statement
High rate of depression	"Patients' depressive disorders are a daily problem."
in primary health care	"We can see a patient with a depressive background
	almost every day."
	"Depression has become more widespread over the
	years."
High cost of management	"The depressive patient requires more consultation time
of depression	to focus on psychological problems."
	"Depression is often accompanied by multiple somatic
	complaints and patients place high expectation on the
	investigations performed with the use of apparatuses."
Patients' difficulties with	"It is difficult to explain to the patient that depression is
accepting the diagnosis	the cause of all his/her complaints."
and with the subsequent	"Patients feel that somatic diseases are "respectable"
treatment	diseases and are afraid to accept the diagnosis of
	depression."
	"Patients refuse to see the psychiatrist because they
	think of them as shrinks who treat insane persons."
	"Patients do not recognize the cause of depression; they
	ignore it and will not do anything to change the
	situation."
	"Many patients stop taking their medication or do not
	start altogether because drugs are expensive."
	"Psychological counselling is expensive and
	psychotherapy is unavailable for many persons due to
	the location of their home."
Physicians' inadequate	"Sometimes FDs do not recognize depression."
resources/skills to help	"It is difficult or impossible for the physician to
patients	eliminate the causes of depression."
	"In addition to drugs, patients need psychotherapy,
	behavioural therapy, family therapy, etc.; however, we
	do not have such skills."
	"It is difficult to refer patients to psychiatrists due to
	their long waiting lists."
	"Seeing the psychiatrist often involves additional costs
	for patients as psychiatric aid may not be available in
	the neighbourhood."
	"There is no co-operation between the FD and the
	psychiatrist or the psychologist."

In addition to these problems, the FDs noted the persistence of frequent depression risk factors as unemployment, problems related to work, low income, insecure future, absence of security, unorganized social work.

Of the FDs 115(56%) had sufficient knowledge to diagnose and treat depression, and 90(44%) respondents considered their knowledge inadequate. The opinion of 181(88%) physicians was that they definitely needed further training.

#### 7.3. Prevalence of depression in primary care in Estonia

Proceeding from the diagnostic categories of ICD-10 for depression, 6-month depressive episode was diagnosed in 258(23.6%) and 1-month depressive episode in 169(15.4%) participants of sample I. Among the participants who were diagnosed with 6-month depressive episode, it was more often moderate and severe. However, mild depressive episode was more frequent among the men ( $\chi^2 = 12.13$ ; p<0.001) compared with the women (Table 9).

Table 9. Grade of depression

	Total n(%)	Women n(%)	Men n(%)
Mild episode	36(14)	17(8)	8(17)
Moderate episode	111(43)	88(42)	16(34)
Severe episode	111(43)	105(50)	24(49)

## 7.4. Factors associated with depression in primary care in Estonia

The factors associated with depression in primary care were feminine gender, lower education, negative life events in the preceding 6 months, lower socioeconomic status, chronic medical illness, disability (Paper IV, Table 1).

Marital status was not significant for the women with depressive episode. Among the men depressive episodes were more frequent for widowers, divorced men and for men who had never been married. Logistical regression revealed that unemployment was a higher risk factor for depression than employment (OR 2.3 CI 1.1–4.8; p<0.05). In this study there was no association between depression and age.

# 7.5. Suitability of the EST-Q2 screening scale depression subscale for screening depression in general practice

The CIDI interview diagnosed 1-month depressive episode in 162(15.3%) participants of sample II, while 300(28.4%) were screened depressive by EST-

Q2. Of the participants who were diagnosed with depressive episode, mild depression was diagnosed in 10%, moderate in 38% and severe in 52%. The EST-Q2 classified 18.8% of the subjects differently in comparison to CIDI (Table 10).

Table 10. Depression diagnosed by CIDI and EST-Q2

EST-Q2 CIDI	Negative n(%)	Positive n(%)	Total n(%)
Negative n(%)	728(68.8)	168(15.9)	896(85.7)
Positive n(%)	30(2.8)	132(12.5)	162(15.3)
Total n(%)	758(71.6)	300(28.4)	1058(100)

The 168(15.9%) persons who did not have depression by CIDI, but whom EST-Q2 screened as depressive, were classified as "false positive". Thirty (2.8%) persons who were diagnosed to be depressive by CIDI, but were not depressive according to EST-Q2, were classified as "false negative".

Table 11 present the results of sensitivity, specificity, false-negative rate, predictive values and LR for different cut-off points for one-month depression.

**Table 11.** Comparison of the test characteristics for the EST-Q2 at cut-off points >11, >10, >12

Screening	Sensiti-	Specifi-	FN	FP	PPV	NPV	Positive	Negative
instrument	vity	city					LR	LR
EST-Q2>11	0.81	0.81	0.19	0.19	0.44	0.96	4.3	0.23
EST-Q2>10	0.86	0.77	0.13	0.23	0.4	0.97	3.7	0.18
EST-Q2>12	0.79	0.84	0.2	0.15	0.49	0.96	4.9	0.25

FN — false negative rate

FP — false positive rate

PPV — positive predictive value

NPV — negative predictive value

LR — likelihood ratio

The EST-Q2 had good specificity, sensitivity, positive predictive value and positive LR for the screening of depression at the present cut-off point >11. By decreasing cut-off by one point, the sensitivity and the false-negative rate improved, but the positive predictive value and the positive likelihood ratio decreased. By increasing cut-off by one point, the sensitivity decreased, while the specificity, the positive predictive value and the positive likelihood value improved.

# 7.6. Combination of symptoms allowing general practitioners to distinguish patients with depression from patients with other biomedical or psychosocial problems

To find out a combination of symptoms distinguishing depressive patients from healthy persons in the best way, two models were developed according to EST-Q2. Model 1 considered 8 most characteristic symptoms included in the EST-Q2 depression scale: feeling of sadness, loss of interest, feeling of worthlessness, self-accusation, thoughts of suicide, feeling lonely, hopelessness, impossible to enjoy things. Model 2 considered 17 symptoms included in the EST-Q2 depression scale plus excessive worrying about several different things, feeling so restless that it is hard to sit still, fatigue or loss of energy, diminished ability to think or concentrate, rest does not restore strength, being easily fatigued, difficulty in falling asleep, restless or disturbed sleep, waking up too early. The significance of the association between the symptoms of EST-Q2 identified and CIDI-identified episodes is presented in Table 12.

**Table 12.** Association between the symptoms of EST-Q2 and CIDI-identified depressive episode: logistic regression model 1 and model 2

Symptoms	Logistic	regression model 1	Logistic regression model 2			
	Estimate	OR (95% CI)	Estimate	OR (95% CI)		
Feeling of sadness	-0.46**	0.63 (0.46 to	-0.50**	0.60 (0.45 to 0.82)		
		0.86)				
Loss of interest	-0.81**	0.45 (0.34 to	-0.64**	0.52 (0.39 to 0.70)		
		0.59)				
Feeling of worthlessness	ns		-0.29**	0.75 (0.61 to 0.92)		
Self-accusations	-0.33**	0.72 (0.58 to	ns			
		0.90)				
Feeling lonely	-0.22*	0.80 (0.65 to	ns			
		0.98)				
Impossible to enjoy	-0.35**	0.70 (0.56 to	-0.26*	0.77 (0.60 to 0.97)		
things		0.89)				
Excessive worry about			-0.27*	0.76(0.60 to 0.97)		
several different things				·		
Rest does not restore			-0.35**	0.70 (0.57 to 0.87)		
strength						

<sup>\*</sup> p<0.05

<sup>\*\*</sup> p<0.01

ns – no significant

Out of the symptoms of Model 1, feelings of sadness, loss of interest, self-accusations, loneliness and inability for enjoyment were the best identifiers of depressive patients. In combination with feeling of sadness, feeling no interest or pleasure in things, feeling of worthlessness, impossibility to enjoy things, excessive worrying about several different things and rest does not restore strength were significantly associated with having depressive disorder in Model 2. The symptoms feeling of sadness, loss of interest and impossible to enjoy things were the identifiers of depressive patients in both models. The best identifier of depressive episode was loss of interest.

We compiled two new scales from the symptom combinations, which enabled us to distinguish between depressive and non-depressive persons on the basis of Model 1 and Model 2. The first scale, EST-QNew1, consists of the following symptoms: feeling of sadness, feeling no interest, self-accusations, feeling lonely and no enjoyment. The second scale, EST-QNew2, consists of the following symptoms: feeling of sadness, feeling no interest, worthlessness, no enjoyment, excessive worrying and rest does not restore strength. We examined the ability of both scales to screen depression at different cut-off points in comparison with the original EST-Q2 depression scale (Table 13).

**Table 13.** Comparison of the test characteristics of the two new models and EST-Q2 at the cut-off point >8 for EST-Qnew1 and at the cut-off point >11 for EST-Qnew2

Screening instruments	Cut-off point	Sensi- tivity	Speci- ficity	FN	FP	PPV	NPV	Positive LR	Negati- ve LR
EST-Q2 Depression Scale	>11	0.81	0.81	0.19	0.19	0.44	0.96	4.3	0.23
EST-Qnew1	>8	0.81	0.82	0.19	0.18	0.45	0.96	4.5	0.23
EST-Qnew2	>11	0.81	0.85	0.19	0.15	0.5	0.96	5.4	0.22

FN – false negative rate

FP – false positive rate

PPV – positive predictive value

NPV – negative predictive value

LR – likelihood ratio

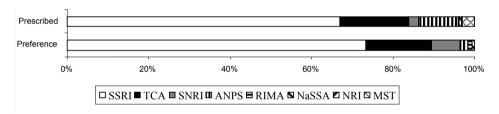
The test characteristics of EST-QNew1 at the cut-off point >8 were comparable to those of EST-Q2. The sensitivity was the same as for EST-Q2, but the specificity increased from 0.81 point to 0.82 and FP and PPV improved by the order of 0.01 and positive LR improved by the order of 0.02.

The test characteristics of EST-QNew2 at the cut-off point >11 were better than those of EST-Q2 and EST-QNew1 at the cut-off point >8. The sensitivity of EST-Qnew2 did not change in comparison with that of EST-Q2, while the specificity increased from 0.81 to 0.85; FP, negative LR and PPV increased from 4.3 to 5.4; at the same time, none of the characteristics became worse. In 50% of the persons who were screened as depressive, depressive disorder had also been diagnosed by CIDI.

# 7.7. The preferences and rationale of family doctors in pharmacological treatment of depression

The FDs indicated selective serotonin reuptake inhibitors as their first-line treatment preference. The medication groups preferred by the FDs for treatment of depression were SSRI (69%), TCA (15%), SNRI (7%), RIMA (1%), NaSSA (1%), tranquillizers (5%) and antipsychotics (2%).

In the list of the first-line drugs noted by the FDs these particular medication groups were also preferred most frequently (Figure 3).



**Figure 3.** Comparison of preference and the classes of antidepressant drugs prescribed by family doctors

The first-line drug in the list of the FDs was Cipramil which was mentioned almost twice more often than the others. Cipramil was followed by Cipralex, Seroxat and Nycoflox which were noted with almost equal frequency. Among them Cipramil, Cipralex and Nycoflox were prescribed more often (Figure 4).

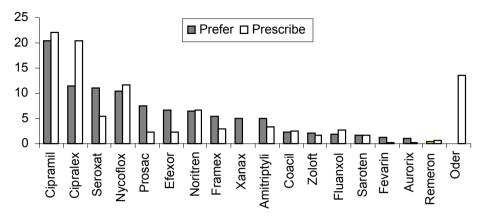


Figure 4. Preference and the drug prescribed by family doctors

The FDs prescribed also medicines, which they did not mention as their preference, these are presented in figure 3 under "Other". This list includes anti-depressants, antipsychotics, and mood stabilizers. The FDs did not mention mood stabilizers as their preference, which actually made up only 0.6% of the whole amount of the medicines prescribed by them.

To the question why they preferred these medicines, the FDs responded providing different reasons (Table 14).

**Table 9.** Rationale of the family doctors regarding preference of medications

Reasons	Example called by FDs
Effectiveness	Highly effective
	Highly effective for the elderly
Side effects	Good tolerance
	Few side effects
	Good for men
Active substance	Fast active substance
	Transient treatment
Ease of administration	Suitable state of medication
	Low dosage
	Low dosage for start
Incremental complaint	For panic and anxiety
	For bulimia
	For insomnia
	For chronic pain
	For patients with somatic complaints, for addicts
Price	Cheap
Other	More available information

The factors that influenced antidepressant selection were good effect, ease of administration, presence of specific clinical symptoms, presence of co-morbid psychiatric disorders and price of medicaments, which are also commonly listed in the literature (Gitlin 2002). In addition, the FDs considered speed of the effect, previous knowledge of effectiveness and ample information to be important.

# 7.8. Antidepressant prescribing patterns among the psychiatrists and the family doctors

Depression was diagnosed in 37 029 patients in 2003: in an outpatient setting by FDs in 19 521 cases and by psychiatrists in 13 444 cases. Altogether 70 039 psychotropic drug visits (drug visit was defined as a visit during which at least 1 drug was prescribed) were made, of which 44 125(63%) were made to FDs and 25 914(37%) were made to psychiatrists. Of these visits 63 987(89%) were antidepressant drug visits, 5741(8%) were antipsychotic drug visits and 311(3%) were mood stabilizer drug visits. Altogether 80 911 medicines were prescribed, of which 71 486(88%) were antidepressants, 8769(11%) were antipsychotics and 656(1%) were mood stabilizers. Antidepressant drug visits and antipsychotic drug visits were made more often to FDs and mood stabilizer drug visits were made more often to the psychiatrists, while in both cases the prescribed medicines were the same (Table 15).

**Table 15.** Drug visits and drug prescribing patterns among the family doctors and the psychiatrists

Patterns	Drug visits n(%)			Prescribed drug n(%)		
	Total	FD	Psychiatrist	Total	FD	Psychiatrist
Antidepressants	63 987(100)	41 527(65)	22 460(35)	71 486(100)	46 158(65)	25 329(35)
Antipsychotics	5741(100)	3723(65)	2018(35)	8769(100)	5170(59)	3599(41)
Mood stabilizers	311(100)	132(42)	179(58)	656(100)	298(42)	358(58)

Altogether 25 410 patients with depression were treated in an outpatient setting during a year. Monotherapy was used for 24 313(95.7%) patients. Antidepressants were used in 88% of the cases, antipsychotics in 7.5% of the cases and mood stabilizers in 0.2% of the cases. Combined treatment was applied in 4.3% of the cases, while a combination of an antidepressant and an antipsychotic was common.

Most often the FDs prescribed SSRIs for treatment of depression; the second choice was TCA, while the drugs of the other groups were prescribed in less

than 3.5 % of the cases. The psychiatrists often prescribed the same classes of antidepressant drugs. In comparison with the psychiatrists, the FDs prescribed more SSRI and the psychiatrists prescribed more TC, SNRI, NaSSA, RIMA and NRI (p=0.0001) (Paper I, Table 2). The most frequently prescribed drugs were fluoxetine (SSRI), escitalopram (SSRI), citalopram (SSRI), paroxetine (SSRI) and nortriptylin (TCA). The FDs prescribed fluoxetine, escitalopram, citalopram and nortriptylin more often compared with the psychiatrists (p=0.0001) and the psychiatrists prescribed the other antidepressants more frequently compared with the FDs (Paper I, Table 3). Only in the case of prescribing amitriptyline there was no difference between the FDs and the psychiatrists. The psychiatrists also prescribed seldom used drugs more often (less than 1%) such as fluoxetine (SSRI), imipramine (TCA), moclobemide (RIMA), milnaciprane (SNRI) and reboxetine (NRI). The FDs and the psychiatrists prescribed 16 different antidepressants from the classes of antidepressant drugs with 28 different names. More frequently were prescribed Cipralex (escitalopram), Nycoflox (fluoxetine), Cipramil (citalogram) and Seroxat (paroxetine). The FDs and the psychiatrists prescribed similar antidepressants for treatment of depression.

### 8. DISCUSSION

Depressive disorders are a common problem in many countries (Satcher et al 2001, Meltzer et al 1995, Aluoja et al 2004), yet they are often not recognized in primary care (Katon et al 1992). Owing to the EU study PREDICT, data about the prevalence of depression among primary health care attendees are available (King et al 2006). In Estonia the psychiatrist was the person who treated depressive patients fifteen years ago. In 1991 training of FDs was started in Estonia, which changed medical service in primary care. Every patient is free to choose his/her FD (Maaroos 2004, Maaroos et al 2004). Usually the FD is the first person to see patients with depressive symptoms. However, as patients can visit the psychiatrist without referral, management of depression can be believed not to be the FDs task. It is therefore important to know how well the FD is prepared for the task of managing patients with depression symptoms.

Like in countries with different social and cultural backgrounds (Ralition *et al* 2000, Soykan *et al* 2003) in our study, most FDs considered depression management to be their task. The readiness of the FDs to deal with depression patients did not depend on the location or association of their practice, which is understandable as all practices in Estonia have the same features-patient lists and accessibility (Maaroos *et al* 2004).

It is highly promising that the motivating factors are evidently based on the FDs' sense of duty. FDs feel that depression is a highly prevalent condition and an important problem in primary care. They acknowledge the great need for its treatment and feel primary responsibility for their patients' treatment. FDs have also experienced that successful treatment of depression improves the patient's health as well as doctor-patient relationship. A study of the FDs consultation style in Estonia (Tähepõld *et al* 2003) showed the same approach: FDs helped patients with psychological problems more than patients expected. In agreement with other studies (Sleath *et al* 2002, Harman *et al* 2001), FDs in Estonia feel that patients have more trust in them than in an unknown specialist. According to our data and other authors opinions (Sleath *et al* 2002, Harman *et al* 2001), it is possible to conclude that in treating depression, FDs are mainly motivated by patient- and relationship-oriented factors.

One should bear in mind that there exist several factors hindering FDs' management of depression. The major issues are related to the FDs' time and knowledge resources as well as the patients' low compliance. Although the frequency of depression appeared as an important motivating factor, it was also mentioned as a problematic factor, as depressive patients increase the workload of FDs. Several studies have shown that 10–25% of patients who visit primary health care specialists; suffer from depression from time to time (Meltzer *et al* 1995, Goldberg *et al* 1992). One of the most disturbing factors for FDs in dealing with depressive patients is the short consultation time per patient. Although an average visit to the FD lasted 9.0 min the longest consultation

lasted 36.3 min in the case of a psychological problem (Tähepõld *et al* 2003). Longer visits of depressive patients to their FDs, compared with visits of other patients have been described by other authors (Sleath *et al* 2002, Harman *et al* 2001). Even the outcome of depression treatment is dependent on the consultation time (Ralition *et al* 2000). Evidently, ordinary consultation is too short for dealing with problems of mental health. The advantage of FDs is the opportunity after the first visit to reserve more time for consultation for patients with a psychological problem.

The studied FDs considered it important to have specialized knowledge and cooperation with other specialists. They wanted to be more trained in diagnosis and treatment of depression, which is consistent with results from other countries (Lecrubier 2001, Simson *et al* 1999, Dowrick *et al* 2000). According to the FDs' opinions, some problems arise from patients' compliance. It is hard for patients to see the true reasons for their problems and to accept the diagnosis of depression, as was shown also by Ralition et al 2000.

Treatment of depression was considered to be the task of the FD, according to our study, and was associated with some problematic issues. Treatment is often time and resource consuming, patients tend to stop taking the prescribed medicine, or they do not procure it altogether; this finding was supported also by other authors (Paykel *et al* 1992). Treatment of depression should be complex and the efficacy of psychotherapy in the treatment in primary care is evident (Ward *et al* 2000). The FDs admitted also their insufficient skills in psychotherapy, lack of cooperation with psychiatrists and psychologists and low availability of psychotherapists. Psychiatric care and psychological care are concentrated into four major cities in Estonia. There is a shortage of psychologists and psychotherapists and most psychological service is not covered by health insurance.

An important evidence of the motivation of FDs to manage with patients with depression is the fact that the FDs considered continuous training still necessary, although most of them had passed advanced training in depression. For comparison, according to a study of Soykan, only the physicians who were interested in psychiatry had passed training in depression (Soykan *et al* 2003). It shows that our FDs recognize the need for dealing with the problem and are often engaged in it in their daily work.

The patients participating in the present survey visited their FD due to several acute or chronic problems, while depression itself happened seldom to be the reason for the visit. Yet in most depressive patients presenting the FD this problem is often masked by other complaints, or coexists with other diseases and conditions, as is noted also by other authors (Goodwin 2006, Saltman *et al* 2005). It has been shown that 81% of depressive patients visiting the FD have only somatic complaints and 56% also have a somatic disease (Lecubier 2001). In many cases neither the doctor nor the patient suspected depression and all attention was only given to the disease, which was the reason

for the visit. Studies of the FDs consultation carried out in Estonia have also indicated that in comparison with biomedical problems patients seldom bring out their psychological problems or expect them to be solved (Tähepõld *et al* 2003, Tähepõld *et al* 2006).

Up to a quarter of patients visiting their FD had depression during the past 6 months involving mostly severe or moderate depressive episode. Hence depression is even a more frequent problem than could be expected according to surveys performed elsewhere. Usually, in primary health care one-month prevalence of depression is estimated to be up to 5–10% (Wittchen *et al* 2002, Salokangas *et al* 1996, Paykel *et al* 1992). In comparison with the prevalence of depression among the patients of Estonian FDs, established with the same method as in other countries (Great Britain, the Netherlands, Slovenia, Spain and Portugal), higher prevalence was only found in Great Britain (Aluoja *et al* 2005). An interesting result of our study is the predominance of severe and moderate depressive episodes, while most previous surveys have found more of moderate and mild depression in primary health care (Hildebrandt *et al* 2003). A situation similar to ours seems to be in Germany where a recent survey found also predominantly heavy and moderate depression (Wittchen *et al* 2002).

Factors that promote depression are demographic indicators, among them marriage was associated with lower prevalence of depression only in the men, which was in concordance with other studies suggesting that marriage can be a protective factor against depression, coronary diseases, II type diabetes and risk behaviour (incurring accidents, smoking and abuse of alcohol) in men (Bebbington 1996, Ross et al 1990). Depression was more frequent in unemployed persons and in those who had hardships with financial subsistence. Thus the favourable processes taking place in the human development of the Estonian society should positively affect the prevalence of depression (http://hdr.undp.org/). At the same time, our study found association of depression with other factors, e.g. the number of negative life events in private life sphere. The same associations were stressed by other authors both among general population and in patients of family practitioners (Paykel et al 1982, Salokangas et al 1998). However, it is not clear if negative life events cause depression, or if persons who tend to be depressive boost their negative life events (Harkness et al 1999).

The Position of FDs in a health care system allows them to screen depression among their attendees. Early detection of depression via screening is the priority task in the whole management of depression Screening instruments are not the means of diagnosing but can be the first step in identifying depressive disorders, which is especially important in primary health care. The present study showed that many of our FDs use screening tests, preferring EST-Q2 in their daily work. The number of items in different screening measures varies, and it has been found that use of shorter screening measures may be as effective as use of longer ones (Robins *et al* 1988).

Our study is the first to compare CIDI and EST-Q2, allowing to evaluate the specificity and sensitivity of these screening instruments. We showed that EST-O2, developed on patients of the psychiatric ward, is applicable to primary care attendees. Our study proved that the sensitivity and specificity of the EST-Q2 depression subscale is good and comparable to the corresponding characteristics of other self-rate instruments (Henkel et al 2004, Dutton et al 2004, Arroll et al 2005, Robins et al 1988). The cut-off point used in EST-Q2 was tested on primary care users by raising it, but no significant improvement of the sensitivity or the specificity was achieved. It was found that although the sensitivity can be further improved by lowering the cut-off point, this brings about lowering of the specificity, yields too many false positive results and reduces the positive predictive value. Some patients with "false-positive" results on screening may have dystymia or some anxiety disorder with concomitant depressive symptoms instead of major depression (Robins et al 1988). Some authors propose, if the predictive value and the likelihood ratio are considered more important, a higher cut-off point can be used, which yields the highest positive predictive value (Peters et al 1995, Arroll et al 2005).

The efficacy of screening scales may depend on the included symptoms. Most self-rate depression screening scales attempt to assess all symptoms used in the diagnostic criteria. Nevertheless, the value of individual symptoms in screening of depression is not clear.

In our study we tried to identify the symptoms, which help FDs to discriminate between patients with and without depressive disorder and, as a result, we developed two new screening scales. The main known symptoms of depression are lowered mood, loss of interest and no enjoyment and reduced energy, accompanied by other symptoms like lower concentration and attention, reduced self-esteem, feeling of guilt and worthlessness, pessimism about the future, suicidality and disturbed sleep and appetite.

First, we tried to identify which combination of affective and cognitive symptoms of EST-Q2 depression subscales discriminate best CIDI-identified depression (WHO 1992). The typical symptoms of depression, sad mood and loss of interest appeared to be the most significant. The best identifier of depressive episode was loss of interest. This is supported by a study of screening depression using two questions, where loss of interest yielded the least number of false positive results and differentiated between depressive persons and non-depressive ones best (Zung 1965). The other indicator of anhedonia, impossibility to enjoy things, was also significantly related to depression. This finding stresses the importance of anhedonia in recognizing depression and supports the idea that while a high negative affect can be general to several negative mood states, lack of a positive affect is specific to depression (Clark et al 1991). Among the affective-cognitive symptoms of depression, also self-accusation and loneliness were significantly related to CIDI-identified depression. When we elaborated a new self-rate scale with 5

items using the symptoms significantly predicting depression (EST-Qnew1), it appeared to screen depression as well as the existing EST-Q2. Like other studies (Whooley *et al* 1997), this shows that reduction in the number of items in a questionnaire does not necessarily diminish its screening properties but a shorter version can be easier for the patient to complete.

Second, when we added somatic and behavioural self-rate symptoms to the model, the set of symptoms discriminating depression changed. In this combination, in addition to *sad mood, loss of interest* and *impossibility of enjoyment*, also *feeling of worthlessness, worrying* and *rest does not restore strength* became significant. Though excessive worrying is a typical symptom of generalized anxiety (GAD), our results suggest that it might be important in identifying primary care patients with depression. Lately the role of repetitive negative cognitions, like rumination and worry in maintaining mood and anxiety disorders have been highlighted (Papageoriou *et al* 2003). This is supported by other studies which show that the score of worrying is equally high in GAD and major depression, and that pathological worry is strongly related to depression (Starcevic *et al* 1995, Muris *et al* 2005).

The symptom of fatigue, which it is usually omitted from depression screening scales for primary care because it can be a sign of somatic illness (Beck *et al* 1997, Zigmond *et al* 1983), proved also important in distinguishing depressive patients. Our study showed that even in primary care patients' fatigue can be a significant identifier of depressive disorder and should be included in a self-report questionnaire, as is the case with PHQ-9 (Kroenke *et al* 2002). Some specific aspect of fatigue, like *fatigue not being relieved after rest*, acquires significance as a characteristic of depression.

On basis of the second combination of symptoms the new scale EST-Qnew2 was developed. It yielded a better result in screening depression than EST-Q2 or EST-Qnew1. Its specificity, FP, PPV and positive LR improved significantly, while the sensitivity and FN remained the same. The screening properties of EST-Qnew2 are equal or excel those of common self-administered scales (Henkel *et al* 2004).

Reporting preferences for drug use for treatment of depression, the FDs mentioned most often the same drugs that they actually prescribed. The reasons why they prescribed these drugs mostly coincided with the reasons presented in the literature (Cohen 1997, Zimmermann *et al* 2004, Gitlin 2002). In a handbook of depression the presented factors of a rational selection of a specific antidepressant were issues relating to complications, side effect profile, ease of administration, safety, history of past response, depressive subtype, neurotransmitter specificity, family history of response, blood level considerations and cost (Gitlin 2002). Different surveys indicate, in addition to presence of comorbid psychiatric disorders, also avoidance of specific side effects, drug-drug interactions and presence of specific clinical symptoms (Zimmermann *et al* 2004, Cohen 1997). In addition, in the present study FDs

considered the speed of achieving the effect, past experience with efficacy and ample information about the drug as important.

The FDs in our study preferred mainly antidepressants in management of depression while SSRIs were their first-line treatment preference. A similar result was obtained in a survey of psychiatrists' drug preferences (Dording *et al* 2002). The less mentioned drugs in preferences were TCA, NSRI and tranquillizers, while antipsychotics as RIMA, NaSSA formed a small part. Our FDs preferred citalopram, escitalopram, paroxetine and fluoxetine. A similar result was obtained from an opinion survey performed on a sample of French psychiatrists in 1999 among which 63.5% mentioned paroxetine as the first-line drug and followed by fluoxetine with 54.5% (Depont *et al* 2003). Escitalopram noted as the second-line drug by our FDs was not used at that time. Evidence from randomized clinical trials suggests that escitalopram is superior to placebo in short-term treatment of depression (Burke *et al* 2002), with efficacy and tolerability comparable to those of other antidepressants including venlafaxine (Bielski *et al* 2004, Montgomery *et al* 2004) and citalopram (Burke *et al* 2002, Lepola *et al* 2004).

Regarding psychopharmacon drug visits made by patients to their FDs or psychiatrists, antidepressant visits accounted for the largest share (89%). Antipsychotics and mood stabilizers were used seldom as, according to general consensus, they are largely prescribed in specific situations when dealing with melancholic, atypical or psychotic depression, and it is not important to use these drugs as first aid in depression (Isometsä 2000, Pincus *et al* 1998). There were no significant differences in prescribing antipsychotics among the FDs and the psychiatrists. Similar trends in use of drugs have been revealed in other surveys as well, where depression was treated with antidepressants in 60% of cases and with antipsychotics and mood stabilizers in a fewer cases (Pincus *et al* 1998, Ernst *et al* 2006).

Of all antidepressant visits 65% were made to FDs, which shows that patients trust their FDs in treatment of their depression. Surveys on patient satisfaction with the services provided by their FD, carried out in Estonia show that patients appreciate highly the professionalism and effectiveness of their FDs (Põlluste *et al* 2004). Similar findings have been reported from other surveys where 65% of persons with depression appreciate highly or very highly FDs skills and knowledge to treat depression (Schwenk *et al* 2004). It is therefore not surprising that a large number of drug visits are made to FDs and the number has is increasing from year to year (Pincus *et al* 1998, Harman *et al* 2003).

Of the antidepressants 65% were prescribed by FDs, which indicated that a large number of patients with depression are treated by FDs. Our results are consistent with those of a survey from Australia where FDs prescribed 86% of the drugs to treat depression (McManus *et al* 2003). FDs mostly used SSRIs to treat depression, followed by TCA-s, while the other antidepressant drugs

where altogether prescribed in less than 3.5% of cases. Similar results have been obtained from other surveys carried out among FDs and psychiatrists (Wilson et al 2003, Petersen et al 2002, Depont et al 2003). Use of mainly the SSRI group of drugs by FDs to treat depression is in accordance with depression guidelines (Depression Guideline Panel 1993). The advantage of the SSRIs over other drugs is ease and safety of use; the therapeutic dose is often achieved with administrating only one pill, the profile of the side effects is lower and there is no need to take blood tests to evaluate the concentration of the drug in the blood (Gitlin 2002). A survey on the use of drugs clearly showed that FDs prescribed the SSRI group of drugs always in therapeutic daily doses, which they did not do in the case of drugs from the TCA and other groups. Psychiatrists, on the other hand, used the TCA group of drugs in bigger doses compared with FDs (McManus et al 2003). It has also been shown that patients tend to discontinue treatment with TCAs more often than treatment with SSRIs (Katon et al 1992), which may indicate that taking SSRIs has more often a positive effect in treatment of depression.

Regarding antidepressants, both the FDs and the psychiatrists prescribed fluoxetine and escitalopram in most cases. However, there were significant differences in the prescribed medicines regarding active agents. Regarding less used drugs the psychiatrists prescribed them more often than the FDs. The drugs preferred by the FDs belong to the SSRI group except for one drug from the TCA group. This demonstrates once more that the FDs used the SSRI group of drugs more often and more liberally. The FDs preferred drug from the TCA group to treat geriatric patients. Similar finding from other studies show that management of depression in the elderly may be conservative owing to which older antidepressants may be over-prescribed (Dearman *et al* 2006).

The most frequently prescribed antidepressants were preferred for several good reasons: good tolerance, good effect and few side effects. A similar finding was obtained in an opinion-survey among psychiatrists where tolerability of the drug appeared important in the choice of the antidepressant (Depont *et al* 2003). The above drugs occupied the first place regarding preference, as reported also by FDs.

### The limitation of the study

The limitation of the study was the fact that only 41% of the practicing FDs agreed to take part. However, as sex and employment characteristics of the respondents corresponded to those of the FDs in general, our results should reflect the real situation regarding the studied problem.

### 9. CONCLUSIONS

- 1. Family doctors are ready and feel motivation to manage patients with depressive complaints, as depression is a frequent important problem in primary care in Estonia. Motivation for dealing with the patients' psychological problems is supported by the family doctors' wish to improve the patients' health as well as doctor-patient relationship. Family doctors acknowledge the great need for treatment of patients with depression. Problematic issues for family doctors are high prevalence of depression among family doctors' attendees, high cost of management of depression, patients' compliance with diagnosis and treatment and physicians' inadequate time resources. Family doctors would like to receive additional training in specific skills for management of depressive patients. Also they consider good cooperation with the psychiatrist and the psychologist important in management of patients with depression.
- 2. One-month depressive episode occurred in every sixth family doctor's attendee and six-month depressive episode occurred in every fourth family doctor's attendee. Moderate depressive episode and severe depressive episode were more prevalent compared with mild depression. Depression is more prevalent among Estonian primary care users than expected.
- 3. Depression was more related to female gender, unemployment, poor financial coping, and with having more than two negative incidents during the past six months. Also, depression was more prevalent among the patients who reported having some long-term illness or disability as well as among those who estimated their general health as poor.
- 4. The EST-Q2 is the most commonly used screening instrument among Estonian family doctors for screening of depression. Testing of different cut-off points for its sensitivity and specificity shows that the cut-off point >11 was the most appropriate. However, although EST-Q2 has good sensitivity and specificity, should be taken into account that it yields false positive results among family doctors' attendees. The screening instrument EST-Q2, created on the basis of population and psychiatric patients, is suitable for use among primary care users.
- 5. Reduction in the number of items in the EST-Q2 depression scale and adding of some somatic and behavioural symptoms plus one anxiety symptom improved the screening properties of the instrument, which makes its completion easier and quicker. The best combination of symptoms allowing family doctors to distinguish patients with depression from patients with other biomedical or psychosocial problems are *loss of interest, sad mood, strength not restored by resting, feeling of worthlessness, excessive worry and incapability for enjoyment.* On the basis of these symptoms, a new screening instrument, EST-Qnew2, vas developed for

- primary health care compared with EST-Q2 has the same sensitivity but higher specificity.
- 6. The drug preference and the actual drug prescription of the family doctors overlapped. The preferences in drug prescription by the family doctors are assessed as rational drug selection in dealing with depression.
- 7. The family doctors prescribed mainly antidepressants for treatment of depression. Among the antidepressants, the family doctors prescribed mostly SSRIs. Estonian family doctors treated depression in accordance with internationally acknowledged treatment guidelines.

### 10. REFERENCES

- Aluoja A, Leinsalu M, Shlik J, Vasar V, Luuk K. Symptoms of depression in the Estonian population: prevalence, sociodemographic correlates and social adjustment. J Affect Disord 2004; 78:27–35.
- Aluoja A, Leinsalu M, Shlik J, et al. Symptoms of depression in the Estonian population: prevalence, sociodemographic correlates and social adjustment. J Affect Disord 2002; 78:27–35.
- Aluoja A, Shlik J, Vasar V, Luuk K, Leinsalu M. Development and psychometric properties of the Emotional State Questionnaire, a self-report questionnaire for depression and anxiety. Nord J Psychiatry 1999; 53: 443-449.
- Aluoja A, Kalda R, Ööpik P jt. Depressioon esmatasandi meditsiinis: PREDICT uuringu esimese etapi tulemused. Eesti Arst 2005; 84(9):652.
- Anderson J, Huppert F, Rose G. Normality, deviance and minor psychiatric morbidity in the community. A population-based approach to General Health Questionnaire data in the Health and Lifestyle Survey. Psychol Med. 1993; 23:475–485.
- Andrade L, Caraveo-Anduaga JJ, Berglund P, Bijl RV, De Graaf R, Vollebergh W, Dragomirecka E, Kohn R, Keller M, Kessler RC, Kawakami N, Kilic C, Offord D, Ustun TB, Wittchen HU. The epidemiology of major depressive episodes: results from the International Consortium of Psychiatric Epidemiology (ICPE) Surveys. Int J Methods Psyhiatr Res 2003; 12(1):3–21.
- Andrews G, Peters L. The psychometric properties of the Composite International Diagnostic Interview. Soc Psychiatry Psychiatr Epidemiol 1998; 33:80–88.
- Arroll B, Goodyear-Smith F, Kerse N, Fishman T, Gunn J. Effect of the addition of a "help" question to two screening questions on specificity for diagnosis of depression in general practice: diagnostic validity study. BMJ 2005; 331: 884.
- Arroll B, Khin N, Kerse N. Screening for depression in primary care with two verbally asked questions: cross sectional study. BMJ 2003; 327:1144–1146.
- Ayuso-Mateos JL, Vazquez-Barquero JL, Dowrick C, Lehtinen V, Dalgard OS, Casey P, Wilkinson C, Lasa L, Page H, Dunn G, Wilkinson G; ODIN Group. Depressive disorders in Europe: prevalence figures from the ODIN study. Br J Psychiatry. 2001; 179:308–316.
- Beck AT. Beck depression inventory. Philadelphia, Pa.:Center for Cognitive Therapy, 1961.
- Beck AT, Guth D, Steer RA, Ball L. Screening for major depression in medical inpatients with the Beck Depression Inventory for Primary Care. Behav Res Thera 1997; 35:785–791.
- Beck AT, Steer RA, Brown GK. BDI-II, Beck Depression inventory: manual. 2nd ed. Boston: Harcourt, Brance and Company; 1996.
- Bebbington P. The origins of sex differences in depressive disorder: bridging the gap. Int Rev Psychiatry 1996; 8:295–332.
- Bielski RJ, Ventura D, Chang CC. A Double-blind Comparison of escitalopram and Venlafaxine extended release in the treatment of major depressive disorders. J Clin Psychiatry 2004; 65:1190–1196.
- Biggerstaff BJ. Comparing diagnostic tests: a simple graphic using likelihood ratios. Statist Med 2000; 19:649–663.

- Blazer DG, Kessler RC, McGognale KA, Swartz MS. The Prevalence and Distribution of Major Depression in a National Comorbidity Sample: The National Comorbidity Survey. Am J Psychiatry 1994; 151:979–986.
- Blazer DG, Kessler RC, Swartz MS. Epidemiology of recurrent major and minor depression with a seasonal pattern. The National Comorbidity Survey. Br J Psychiatry 1998; 172:164–167.
- Boyatzis, R. Transforming qualitative information: Thematic analysis and code development. Thousand Oaks, CA: Sage; 1998.
- Burke WJ, Gergel I, Bose A. Fixed-dose trial of the single isomer SSRI escitaloprpm in depressed outpatients. J Clin Psychiatry 2002; 63:331–336.
- Bruce ML, Takeuchi DT, Leaf PJ. Poverty and psychiatric status. Longitudinal evidence from the New Haven Epidemiologic Catchments Area study. Arch Gen Psychiatry 1991; 48:470–474.
- Carmody DP. Psychometric characteristics of the Beck Depression Inventory-II with college students of diverse ethnicity. J of Psych in Clin Pract 2005; 9(1):22–28.
- Chilvers C, Dewey M, Fielding K, Gretton V, Miller P, Palmer B, Weller D, Churschill R, Williams I, Bedi N, Duggan C, Lee A, Harruson G. Antidepressant drugs and generic counselling for treatment of major depression in primary care: randomised trial with patient preference arms. BMJ 2001; 322:772
- Chisholm D, Diehr P, Knapp M, Patrick D, Treglia M, Simon G. Depression status, medical comorbidity and resource costs. Br J Psychiatry 2003; 183:121–131.
- Clinical Practice Guideline Number 5: Depression in Primary Care, vol 2: treatment of Major Depression. Rockville, Md: US Dept Health Human Services, Agency for Health Care Policy and Research; 1993. AHCPR publication 93–0550.
- Cohen LJ. Rational drug use in the treatment of depression. Pharmacotherapy 1997; 17:45–61.
- Composite International Diagnostic Interview. Version 2.1. Geneva, World Health Organization. 1997.
- Davidson JR, Meltzer-Brody SE. The under recognition and under treatment of depression: what is the breadth and depth of the problem? J Clin Psychiatry 1999; 60(7):4–9.
- Dearman SP, Waheed W, Nathoo V, Baldwin RC. Management strategies in geriatric depression by primary care physicians and factors associated with the use of psychiatric services: a naturalistic study. Aging Ment Health 2006; 10(5):521–524.
- Denzin NK, Lincoln YS. (Eds.) (1994). Handbook of qualitative research. Thousland Oaks a.o.: Sage Publications. 643 p.
- Depont F, Rambelomanana S, Le Puil S, Begaud B, Verdoux H, Moore N. Anti-depressants: psychiatrists' opinions and clinical practice. Acta Psychiatr Scand 2003; 108:24–31.
- Depression Guideline Panel. Clinical practice Guideline: Depression in Primary Care, 2: Treatment of Major Depression. Rockville, Md: US Dept of Health and Human Services, Agency for Health Care Policy and research; 1993. AHCRP publication 93–0551.
- Dording CM, Mischoulon D, Petersen TJ, Kronbluh R, Gordon J, Nierenberg AA, Rosenbaum JE, Fava M. The pharmacologic management of SSRI-induced side effects: a survey of psychiatrists. Ann Clin Psychiatry. 2002; 14(3):143–147.

- Dowrick C, Gask L, Perry R, Dixon C, Usherwood T. Do general doctors' attitudes towards depression predict their clinical behaviour. Psychol Med 2000; 30:413–419.
- Dowrick C, Dunn G, Ayuso-Mateos JL, Dalgard OS, Page H, Lehtinen V, Casey P, Wilkinson C, Vazquez-Braquero JL, Wilkinson G. Problem solving treatment and group psychoeducation for depression: multicentre randomized controlled trial. Outcomes of Depression International Network (ODIN) Group. BMJ 2000; 321:1450–1454.
- Dutton GR, Grothe KB, Jones GN, Whitehead D, Kendra K, Brantley PJ. Use of the Beck Depression Inventory-II with African American primary care patients. Gen Hosp Psychiatry 2004; 26:437–442.
- Goodwin GM. Depression and associated physical diseases and symptoms. Dialogues Clin Neurosci 2006; 8:259–265.
- Hildebrandt MG, Stage KB, Kragh-Soerensen P. Gender and depression: a study of severity and symptomatology of depressive disorders (ICD-10) in general practice. Acta Psychiatr Scand 2003; 107(3):197–202.
- Endler NS, Rutherford A, Denisoff E. Beck Depression Inventory: Exploring its dimensionality in a nonclinical population. J Clin Psychol 1999; 55.1307–1312.
- Ernst CL, Bird SA, Goldberg JF, Ghaemi SN. The prescription of psychotropic medications for patients discharged from a psychiatric emergency service. J Clin Psychiatry 2006; 67(5):720–726.
- EU Predict Depression Study. [http://www.rfc.ucl.ac.uk/departments/EUPredict/index.htm]
- Fernandez A, Haro JM, Codony M, Vilagut G, Martinez-Alonso M, Autonell J, Salvador-Carulla L, Ayuso-Mateos JL, Fullana MA, Alonso J. Treatment adequacy of anxiety and depressive disorders: primary versus specialised care in Spain. J Affect Disord 2006; 96(1–2):9–20.
- Frefuson JM. Depression: Diagnosis and Managment for the Primary Care Physician. Primary Care Companjon J Clin Psychiatry 2000; 2:173–178.
- Frenrich M, Weissman MM, Warner V. Screening for depressive disorder in children and adolescents: validating the Centre for Epidemiologic Studies Depression Scale for Children. Am J Epidemiol 1990; 131:538–551.
- Gitlin MJ. Pharmacological treatment of depression. In: Gotlib IH, Hammen CL, eds. Handbook of Depression. New York, London. The Guilford Press. 2002. p 360–82.
- Goldberg D. The detection of psychiatric illness by questionnaires. London, Oxford University Press, 1972.
- Goldberg DP. Huxley P. Common Mental Disorders: a bio-social model. London, New York: Travistock/Routledge; 1992.
- Goldberg DP, Williams P. The user's guide to the general health questionnaire. Windsor: NFER-Nelson, 1988.
- Hall DJR. Current Clinical Strategies. Handbook of Psychiatric Drugs. Laguna Hills, Calif: Current Clinical Strategies Publisching; 1997.
- Harman J, Crystal S, Walkup J, Olfson M. Trends in Elderly Patients' Office Visits for the Treatment of Depression According to Physician Speciality: 1985–1999. The Journal of Behavioral Health Services & Research 2003; 30:332–341.

- Harman JS, Schulberg HC, Muslant BH, Reynolds CF. The Effect of Patient and Visit Characteristics on Diagnosis of Depression in Primary Care. J Fam Pract 2001, 50:1068.
- Harkness KL, Monroe SM, Simons AD, et al. The generation of life events in recurrent and non-recurrent depression. Psychol Med 1999; 29:135–144.
- Henkel V, Mergel R, Kohnen R, Allgaier A-K, Möller H-J, Hegerl U. Use of brief depression screening tool in primary care: consideration of heterogeneity in different patient groups. Gen H Psychiatry 2004; 26:190–198.
- Herman H, Patrick DL, Diehr P, Martin ML, Fleck M, Simon GE, Buesching DP. Longitudinal investigation of depression outcomes in primary care in six countries: the LIDO Study. Functional status, health service use and treatment of people with depressive symptoms. Psych Medicine 2002; 32:889–902.
- Human Development Report 2006 [http://hdr.undp.org/].
- Isometsä E. Depressioon. Lönnqvist J, Heikkinen M, Hendriksson M, Marttunen M, Partonen T. Psühhiaatria. Tallinn. AS Medicina 2000. p. 111–30
- Katon W, Von Korff M, Lin E, Buch T, Ormel J. Adequacy and duration of antidepressant in primary care. Med Care. 1992; 30:67–76.
- Katon W, Schulberg H. Epidemiology of depression in primary care. Gen Hosp Psychiatry 1992; 14:237–247.
- Keller MB, McCullough JP, Klein DN, Arnow B, Dunner DL, Gelenberg AJ, Markowitz JC, Nemerhoff CB, Russell JM, Thase ME, Trivedi MH, Zajetcka J. A comparison of nefazodine, the cognitive behavioural-analysis system of psychotherapy, and their combination for the treatment of chronic depression. N Engl J Med 2000; 342:1462–1470.
- Kessler RC, McGonagle KA, Zhao S, et al. Lifetime and 12-month prevalence of DSM-III-R psychiatric disorders in the United States. Results from the National Comorbidity Survey. Arch Gen Psychiatry 1994; 51:8–19.
- Kessler RC, Berglund P, Demler O, Jin R, Koretz D, Merikangas KR, Rush AJ, Walters EE, Wang PS. The epidemiology of major depressive disorder: results from the National Comorbidity Survey Replication (NCS-R). JAMA 2003 Jun 18; 289(23):3095–3105.
- Kessler RC, Demler O, Frank RG, Olfson M, Pincus HA, Walters EE, Wang P, Wells KB, Zaslavsky AM. Prevalence and Treatment of Mental Disorders, 1990 to 2003. N Engl J Med 2005; 352:2515–2523.
- King M, Weich S, Torres F, Svab I, Maaroos H, et all. Prediction of depression in European general practice attendees: The PREDICT study. BMC Public Health 2006 Jan 12; 6(1):6.
- Kornbluh R, Papakostas GI, Petersen T, Neault NB, Nierenberg AA, Rosenbaum JF, Fava M. A survey of prescribing preferences in the treatment of refractory depression: recent trends. Psychopharmacol Bull. 2001; 35:150–6.
- Kroenke K, Spitzer RL. The PHQ-9: a new depression diagnostic and severity measure. Psychiatr Annals 2002; 32:509–521.
- Lecrubier Y. The Burden of depression and anxiety in general medicine. J Clin sychiatry 2001; 62:3–12.
- Lecrubier Y, Boyer P, Lepine J-P, et al. The identification of psychiatric disorders in primary care. Eur Psychiatry 1996; 11(4):178–179.

- Lepine JP, Gastpar M, Mendlewicz J, Tylee A. Depression in the community: the first pan-European study DEPRES (Depression Research in European Society). Int Clin Psychopharmacol. 1997; 12(1):19–29.
- Lepola U, Wade A, Andersen HF. Do equivalent doses of escitalopram and citalopram have similar efficacy? A pooled analysis of two positive placebo-controlled studies in major depressive disorder. Int Clin Psychopharmacol 2004; 19:149–155.
- Maaroos HI. Family Medicine as a Model of Transition from Academic Medicine to Academic Health Care: Estonian's Experience. CMJ 2004, 45(5):563–566.
- Maaroos HI, Meiesaar K. Does equal availability of geographical and human resources guarantee access to family doctors in Estonia. Croatian Medical Journal 2004; 45:567–72.
- Malt UF, Robac OH, Madsbu H-P, Bakke O, Loeb M. The Norwegian naturalistic treatment study of depression in general practice (NORDEP)-I: randomised double blind study. BMJ, 1999; 318(7192):1180–1184.
- Marks, D. & Yardley, L., Research methods for clinical and health psychology. London: Sage; 2004.
- McManus P, Mant A, Mitcell P, Britt H, Dudley J. Use of antidepressants by general practitioners and psychiatrist in Australia. Aust N Z J Psychiatry. 2003; 37:184–189.
- Meltzer H, Gill B, Petticrew M, Hinds K. OPCS Surveys of Psychiatric Morbidity in Great Britain. Report No 1. The prevalence of psychiatric morbidity among adults aged 16–64 living in private households in Great Britain. HMSO: London; 1995.
- Montgomery SA, Huusom AK, Bothmer J. A randomised study comparing escitalopram with vanlafaxine XR in primary care patients with major depressive disorder. Neuropsychology 2004; 50:57–64.
- Murray CJL, Lopez AD. Global mortality, disability, and the contribution of risk factors: Global Burden of Disease Study. Lancet 1997, 349:1436–1442.
- Murray CJL, Lopez AD. Alternative projections of mortality and disability by cause 1990–2020: Global Burden of Disease Study. Lancet 1997, 349:1498–1504.
- Mynors-Wallis LM, Gath DH, Day A, Baker F. Randomised Controlled Trial of problem solving treatment, Antidepressant medication, and combined treatment for major depression in primary care. BMJ 2000; 320:26–30.
- Patten SB, Wang JL, Williams JV, Currie S, Beck CA, Maxwell CJ, El-Guebaly N. Descriptive epidemiology of major depression in Canada. Can J Psychiatry. 2006; 51(2):84–90.
- Paykel ES, Priest GR. Recognition and management of depression in general practice: consensus statement. Br Med J 1992; 305:1198–1202.
- Paykel ES, Cooper Z. Life events and social stress. In: Paykel S, ed. Handbook of affective disorders. Singapore: Churchill Livingstone; 1982. p. 149–170.
- Peters, L. & Andrews, G. A procedural validity study of the computerized version of the Composite International Diagnostic Interview. Psychological Medicine 1995; 25:1269–1280.
- Petersen T, Dording C, Neault NB, Kornbluh R, Alpert JE, Nierenberg AA, Rosenbaum JF, Fava M. A survey of prescribing practices in the treatment of depression. Prog Neuropsychopharmacol Biol Psychiatry. 2002; 26:177–187.
- Physician's Desk Reference. Montvale, NJ. Medical Economics Co Inc;1997.

- Pincus HA, Tanielian TL, Marcus SC, Olafson M, Zirin DA, Thomson J, et al. Prescribing Trends in Psychotropic Medications: Primary Care, Psychiatry and Other Medical Specialties. JAMA 1998; 279:526–531.
- Poutanen O. Depression in primary health care patient; prevalence, recognition, definition, and the consulting situation. Psychiatria Fennica 1996; 27:115–128.
- Põlluste K, Kalda R, Lember M. Evaluation of Primary Health Care Reform in Estonia from Patients' Perspective: Acceptability and Satisfaction. CMJ 2004; 45:582–587.
- Radloff LS. The CES-D scale: a self-report depression scale for research in the general population. Applied Psychological Measurement 1977; 1:385–401.
- Ralition S, Mowat H, Bain J. Optimizing the care of patients depression in primary care: the views of general doctors. Health Soc Care Community 2000, 8:119–128.
- Riffenburgh RH. Statistics in medicine. Clinical Investigation Department. Naval Medical Center San Diego, San Diego, California. 1999 Academic Press
- Robins L N, Wing J K, Wittchen H-U, Helzer J E, Babor T, Burke J, Framer A E, Jablensky A, Pickens R, & Regier D. The Composite International Diagnostic Interview. An epidemiologic instrument for use in conjunction with different diagnostic systems and in different cultures. Archives of General Psychiatry 1988; 45:1069–1077.
- Ross CE, Mirowsky J, Goldsteen K. The impact of the family on health: the decade in review. JMF (Journal of Marriage and the Family) 1990; 52:1059–1078.
- Rutz W. Preventing suicide and premature death by education and treatment. J Affect Disord 2001: 62:123–129.
- Salokangas RK, Poutanen O. Risk factors for depression in primary care. Findings of the TADEP project. J Affect Disord 1998; 48(2–3):171–80.
- Salokangas RKR, Poutanen O, Stengard E, et al. Prevalence of depression among patients seen in community health centres and community mental health centres. Acta Psychiatr Scand 1996; 93:427–433.
- Saltman DC, Sayer GP, Whicker SD. Co-morbidity in general practices. Postgrad Med J 2005; 81:474–480.
- Satcher D. Global mental health: its time has come. JAMA 2001; 285:1697.
- Sheikh JI, Yesavage JA. Geriatric depression scale (GDS): recent evidence and development of a shorter version. In: Brink TL, ed. Clinical gerontology: a guide to assessment and intervention. New York: Haworth, 1986:165–73.
- Schulberg HC, Katon W, Simon GE, Rush AJ. Treating major depression in primary care practice: an update of the Agency for Health Care Policy and Research practice guidelines. Arch Gen Psychiatry 1998; 55:1121–1127.
- Schwenk TL, Evans DL, Laden SK, Lewis L. Treatment Outcome and Physician-Patients Communication in Primary Care Patients With Chronic, recurrent Depression. Am J psychiatry 2004; 161:1892–1901.
- Sharp LK, Lipsky M. Screening for Depression across the Lifespan: A review of Measures for Use in Primary Care Settings. Am Fam Physican 2002; 66:1001–1008.
- Simon GE, Fleck M, Lucas R, Bushnell DM, LIDO Group. Prevalence and Predictors of Depression Treatment in an International Primary Care Study. Am J psychiatry 2004; 161:1626–1634.

- Simson GE, VonKorff M, Piccinelli M, Fullerton C, Ormel J. An International Study of the Relation between Somatic Symptoms and Depression. N Engl J Med 1999; 341:1329–1335.
- Simon GE, Fleck M, Lucas R, Bushnell DM, LIDO Group. Prevalence and Predictors of Depression Treatment in an International Primary Care Study. Am J Psychiatry 2004; 161:1626–1634.
- Sleath B, Rubin RH. Gender, ethnicity, and physician-patient communication about depression and anxiety in primary care. Patient Educ Couns 2002; 48:243–252.
- Soykan A, Oncu B. Which GP deals better with depressed patients in primary care in Kastamonu, Turkey: the impacts of 'interest in psychiatry' and 'continuous medical education'. Fam Pract 2003; 20:558–562.
- Spitzer RL, Kroenke K, Williams JBW, and the patient ahealth Questionnaire Primary Care Study Group. Validation and Utility of a Self-report Version of PRIME-MD: The PHQ Primary Care Study. JAMA 1999; 282(18):1737–1744.
- Sutherland JE, Sutherland SJ, Hoehns JD. Achieving the best outcome in treatment of depression. J Fam Pract 2003; 52:201–209.
- Tardieu S, Bottero A, Blin P, Bohbot M, Goni S, Gerard A, Gasquet I. Roles and practices of general practitioners and psychiatrists in management of depression in the community. BMC Fam Pract. 2006; 30:7–15.
- Telford R, Hutchinson A, Jones R, Rix S, Howe A. Obstacles to effective treatment of depression: a general practice perspective. Fam Pract 2002; 19:45–52.
- The ICD-10 Classification of Mental and Behavioral Disorders: Clinical descriptions and diagnostic guidelines. World Health Organization, 1992.
- Thase M. How should efficacy be evaluated in randomized clinical trials of treatments for depression? J Clin Psychiatry 1999; 60:23–31.
- Tylee A. Depression in Europe: experience from the DEPRESS II survey. Eur Neuropsyhoph 2000; 10:445–448.
- Tylee A, Gaspar M, Lepine JP, Mendlewicz J. Identification of depressed patient types in the community and their treatment needs: finding from the DEPRESS II survey. DEPRES Steering Committee. Int Clin Psychopharmacol 1999; 14(3):153–165.
- Tähepõld H, Maaroos HI, Kalda R, van den Brink-Muinen A. Structure and duration of consultations in Estonian family practice. Scand J Prim Health Care 2003; 21:167–170.
- Tähepõld H, van den Brink-Muinen A, Maaroos HI. Patient expectations from consultation with family physician. Croat Med J 2006; 47:148–154.
- U. S. Preventive Services Task Force. Screening for depression: Recommendations and Rationale [Clinical Guidelines]. Ann Intern Med 2002; 136:760–764
- Üstün TB, Sartorius N. Mental Illness in General Health Care. An International Study. Chichester: John Wiley & Sons; 1995.
- Yesavage JA, Brink TL, Rose TL, Lum O, Huang V, Adey M, et al. Development and Validation of a geriatric depression screening scale: a preliminary report. J Psychiatr Res 1983; 17:37–49.
- Watson L, Pignone MP. Screening accuracy for late-life depression in primary care: A systematic review. J Fam Pract 2003; 52(12):956–964.

- Ward E, King M, Loyd M, Bower P, Sibbald B, Farrelly S, Gabbay M, Tarrier N, Addington-Hall J. Randomised controlled trial of non-directive counselling, cocnitive-behaviour therapy, and usual general practitioner care for patients with depression. I: Clinical effectiveness. BMJ 2000; 321:1383–1388.
- Weich S, Lewis G, Churchill R, Mann A. Strategies for the prevention of psychiatric disorder in primary care in south London. J Epidemiol Community Health. 1997; 51:304–309.
- Weich S, Lewis G. Poverty, unemployment, and common mental disorders: population based cohort study. BMJ 1998; 317:115–119.
- Wells KB, Strum R, Sherbourne CD, Meredith LS. Caring for depression. Cambridge, Mass.: Harvard University Press, 1996.
- Wilhelm K, Mitchell P, Slade T, Brownhill S, Andrews G. Prevalence and correlates of DSM-IV major depression in an Australian national survey. J Affect Disord. 2003; 75(2):155–162.
- Williams DR, Gonzalez HM, Neighbors H, Nesse R, Abelson JM, Sweetman J, Jackson JS. Prevalence and distribution of major depressive disorder in african americans, Caribbean blacks, and non-Hispanic whites: results from the national survey of american life. Arch Gen Psychiatry 2007; 64(3):305–115.
- Wilson I, Duszynski K, Mant A. A 5-year follow-up of general practice patients experiencing depression. Fam Practice 2003; 20:685–689.
- Wittchen HU. Reliability and validity studies of the WHO-Composite International Diagnostic Interview (CIDI): a critical review. J Psychiatry Res 1994; 28:57–84.
- Wittchen HU, Pittrow D. Prevalence, recognition and management of depression in primary care in Germany: the Depression 2000 study. Hum Psychopharmacol 2002; 17(1):1–11.
- Whooley MA, Simon GE. Managing Depression in Medical Outpatients. Eng J Medicine 2000; 343:1942–1950.
- World Health Organization. Composite International Diagnostic Instrument (CIDI). Version 2.1. Geneva: WHO; 1997.
- Zimmermann M, Posternak M, Friedman M, Attiullah N, Baymiller S, Boland R, et al. Which factors influence psychiatrists' selection of antidepressants? Am J Psyciatry 2004: 161:1285–1289.
- Zung WW. A self rating depression screening scale. Arch Gen Psychiatry 1965; 12:63–70.

# 11. APPENDICES

# 11.1. Questionnaire for family doctors

# Appendix 1

1.	Personal data:				
	Age				
	Sex				
	Length of service as a physician				
	Length of service as a family doctor				
	Type of practice: Solo Group				
	Location: Rural Urban				
	What problems do you meet in your everyday work with depressive				
	patients?				
5.	Is management of depression your daily work? Yes No				
	. Are you ready to deal with depressive patients? Yes No				
7.	. If yes, what is you motivation to deal with depressive patients?				
8.	Do you deal with depression? Yes No				
9.	Do you have sufficient knowledge to deal with depression?				
10	.Do you need further training to deal with depression? Yes No				
11	.Do you use any screening instruments for screening of depression? Yes				
	No				
12	.If yes, what kind of screening instruments?				
13	.What medicines do you preferred for treatment of depression according to				
	name?				
14	Why do you prefer these medicines?				

# 11.2. Questionnaire for patients

# Appendix 2

Emotional State Questionnaire (EST-Q2)

Below are given some problems that people may have. Please indicate how often each problem has bothered you during the past month and mark one of the

boxes to the right that best corresponds to your problems.

boxes to the right that best corresponds to	Not at	Sel-	Some-	Often	All the
	all	dom	times	Onten	time
Feelings of sadness	0	1	2	3	4
2. Feeling no interest or pleasure in things	0	1	2	3	4
3. Feelings of worthlessness	0	1	2	3	4
4. Self-accusations	0	1	2	3	4
5. Recurrent thoughts of death or suicide	0	1	2	3	4
6. Feeling lonely	0	1	2	3	4
7. Hopelessness about the future	0	1	2	3	4
8. Impossible to enjoy things	0	1	2	3	4
9. Feeling easily irritated or annoyed	0	1	2	3	4
10. Feeling anxious or fearful	0	1	2	3	4
11. Tension or inability to relax	0	1	2	3	4
12. Excessive worry about several different things	0	1	2	3	4
13. Feeling so restless that it is hard to sit still	0	1	2	3	4
14. Easily startled	0	1	2	3	4
15. Sudden attacks of panic with palpitations, shortness of breath, faintness or other frightening bodily sensations	0	1	2	3	4
16. Fear of being outside home alone	0	1	2	3	4
17. Feeling afraid in streets or open places	0	1	2	3	4
18. Fear of fainting in public	0	1	2	3	4
19. Feeling afraid of travelling by bus, train or car	0	1	2	3	4
20. Afraid to be the centre of attention	0	1	2	3	4
21. Fear of interaction with strangers	0	1	2	3	4
22. Fatigue or loss of energy	0	1	2	3	4
23. Diminished ability to think or concentrate	0	1	2	3	4
24. Rest does not restore strength	0	1	2	3	4
25. Being easily fatigued	0	1	2	3	4
26. Difficulty falling asleep	0	1	2	3	4
27. Restless or disturbed sleep	0	1	2	3	4
28. Waking up too early	0	1	2	3	4

### **SUMMARY IN ESTONIAN**

### Depressiooni käsitlus peremeditsiinis

Depressioon on üks sagedaseim psühhiaatriline häire esmatasandi arstiabi kasutajate seas. Euroopas esineb depressiooni kuni 10%-l elanikkonnast ja 25%-l esmatasandi arstiabi kasutajatest esineb depressiooni sümptomeid. Levimusnäitajad erinevad paikkonniti kaks kuni kolm korda sarnase metoodika kasutamisel. Eestis läbi viidud uuringu alusel kannatab 11.1% täiskasvanud elanikkonnast depressiivsete sümptomite all. Krooniliste haiguste seas esmatasand arstiabis on ta oma sageduselt teisel kohal hüpertensiooni järel ja enamus patsiente (73%) kellel on diagnoositud depressioon, kogevad rohkem kui ühe depressiivse episoodi oma eluea jooksul. Töövõimetuse põhjustajana prognoositakse depressiooni maailmas aastaks 2020 teiseks südame-veresoonkonna haiguste järel. Depressioon tekib bioloogiliste, sotsiaalsete ja psühholoogiliste tegurite koosmõjus. Universaalseteks riskiteguriteks peetakse naissugu ja madalat sotsiaal-majanduslikku staatust. Veel seostatakse depressiooni vähese sissetuleku, madala haridustaseme, töötuse, negatiivsete elusündmuste ja kehalise tervise probleemidega. Eestis on leitud, et depressioon seostub oluliselt halvema subjektiivse sotsiaalse funktsioneerimisega. Vaatamata sagedasele esinemisele, jääb uuringute alusel enamik depressiivsete häiretega isikutest endiselt õige diagnoosita ja adekvaatse ravita. Esmatasandi arstid diagnoosivad ligikaudu ainult 25–50%-l depressiooni all kannatajatest depressiivset häiret. Samas on need patsiendid perearstidele suuremaks probleemiks, kui somaatilisi haigusi põdevad patsiendid. Depressiooni diagnoosi püstitamine osutub sageli keerukaks nõudes eelnevalt palju kliinilisi uuringuid ja aega. Depressiooni diagnoosimiseks olevad struktureeritud psühhiaatrilised intervjuud on liiga palju aega nõudvad ja ei ole kasutatavad perearsti igapäevases töös. Kasutusel on küll mitmeid enesehinnangulisi skriining instrumente, nagu Becki Depressiooni Skaala, Zungi Enesehinnanguline Depressiooni skaala ning Üldine Tervise Küsimustik, mis erinevad küsimuste arvu ja erinevate sümptomite poolest, kuid need ei diagnoosi depressiooni. Eestis on välja töötatud on populatsiooni ja psühhiaatriliste patsientide uuringu alusel Emotsionaalse Enesehinnangu Küsimustik. Depressiooni ravi kestab kaua ja suur osa patsientidest katkestab selle. Enam kui 60 aastat kasutatakse depressiooni raviks antidepressante ning suure muudatuse tõi uue ravimi rühma — selektiivsete serotoniini tagasihaarde inhibiitorite kasutusele võtmine. Perearstide võime diagnoosida ja ravivida depressiooni on seotud tema teadmistega ja sooviga oma teadmisi täiendada. Vähe on uuritud perearstide hoiakuid ja arvamust depressiooniga seonduvate probleemide kohta.

Eestis on patsiendi esmase kontakti isik perearst, samas on võimalik pöörduda psühhiaatri poole ilma saatekirjata. Depressiooni ravi määravad patsientidele Eestis valdavalt psühhiaatrid ja perearstid. Samas ei ole Eestis uuritud

perearstide arvamust depressiooni probleemidega seonduva kohta ja ei ole tehtud ühtegi struktureeritud diagnostilisel intervjuul põhinevat meeleoluhäirete ning depressiooni uuringut. Pole teada, depressiivsete häirete esinemissagedust perearsti poole pöörduvate patsientide seas. Ei ole teada, millised sümptomid aitavad kõige paremini perearstidel eristada depressiooni põdevaid patsiente teiste psüühiliste ja tervise probleemidega patsientidest. Nende teadmiste omamine aitaks planeerida diagnostilisi ja raviressursse, juhtida perearstide tähelepanu depressiivse häirega patsiendile, tundes riskirühmi, hoida mõnel juhul ära depressiooni vallandumise ning ennetada depressiooniepisoodide kordumist ja häire krooniliseks muutumist.

Käesolev uurimitöö on läbi viidud koostöös üle-eurooroopalise depressiooni uurimisprojekiga PREDICT, mille tausta uuringuks viidi läbi perearstide arvamusuuring. PREDICT uuring võimaldas läbi viia ulatusliku perearsti patsientide intervjueerimise ravusvahelise diagnostilise intervjuu (*Composite International Diagnostic Interview*, CIDI) depressiooni alaosaga, uurida depressiooni riskidegurite seost ja esinemist selleks uuringuks välja töötatud riskitegurite küsimustikuga. Sellele uuringule lisasime Emotsionaalse Enesetunde Küsimustiku testimaks viimase kasutatavust esmatasandi meditsiinis.

### Uurimistöö eesmärgid

- 1. Uurida perearstide valmisolekut, seda motiveerivaid tegureid, probleeme ja vajadusi tegelemaks depressiooni sümptome omavate patsientidega.
- 2. Hinnata depressiooni esinemissagedus perearsti poole pöörduvate patsientide seas.
- 3. Analüüsida depressiooni seoseid sotsiaaldemograafilise tausta, elusündmuste ja tervisehinnanguga perearsti poole pöörduvatel patsientidel.
- 4. Testida emotsionaalse enesehinnangu küsimustiku EEK-2 kasutatavust perearsti patsientide seas depressiooni skriinimiseks.
- 5. Leida kombinatsioon sümptomitest, mis kõige paremini aitab perearstidel eristada depressiooni põdevaid patsiente teiste probleemidega patsientidest.
- 6. Välja selgitada perearstide ravimi eelistus ja selle põhjused depressiooni raviks.
- 7. Teada saada milliseid ravimeid perearstid patsientidele määrasid depressiooni raviks.

### Uuritavad ja meetodid

Teada saamaks perearstide valmisolekut tegelemaks depressiivseid häireid omavate patsientidega ja sellega seonduvat motivatsiooni, probleeme ja vajadusi viidi läbi 2002 aasta sügisel arvamusuuring, milles osales 205 praksist omavat perearsti kutsega perearsti Eesti erinevatest paikadest. Perearstide poolt

täitdetud küsimustikku analüüsiti sagedusjaotus tabelitega ja kvalitatiivse temaatilise analüüsiga.

Uuringusse hõlmati 1370 järjestikust 18–75 aastat patsienti 23 perearsti (15 maal, 8 linnas) praksisest, sõltumata pöördumise põhjusest järgneva 2–3 kuu jooksul. Neist osales 1100 patsienti. Uuringusse ei võetud piiratud liikumis ja teovõimega, raske somaatilise või psüühilise haigusega patsiente, uuringus osalemine eeldas eesti keele oskust.

Depressiooni esinemissageduse hindamiseks ja seoste uurimiseks sotsiaaldemograafilise tausta, elusündmuste ning tervisehinnanguga, viidi läbi prospektiivne uuring, milles osales korrektselt CIDI ja riskiküsimustiku täitnud 1094 patsienti. Depressiooni ja taustategurite seose olulisust hinnati  $\chi^2$ -testiga, seose iseloomu täpsustati logistilise regressioonanalüüsi abil ja rvutati šansside suhe.

EEK-2 kasutatavuste hindamiseks perearsti patsientidel ja kõige parema sümptomite kombinatsiooni leidmiseks, mis aitaksid perearstil välja sõeluda depressiivseid patsiente, viidi läbi prospektiivne uuring 1058 parsieniga, kellel oli korrektselt täidetud CIDI ja EEK-2. Võrdluseks kasutati neliktabelid, arvutati välja EEK-2 tundlikkus, spetsiifilisus, ennustatav prognoosiväärtus ja haiguse esinemise tõenäosus. Kõige parema sümptomite kombinatsiooni välja selgitamiseks kasutati astmelist logistilist regressiooni.

Perearstide ravimite eelistuse ja selle põhjuste välja selgitamiseks depressiooni ravis, saadi andmed arvamusuuringust, mis sisaldas vastavaid küsimusi. Eelistuse hindamiseks liideti ravimi nimetamise korrad, eelistuse põhjuseid analüüsiti sisuanalüüsi meetodit.

Teada saamaks, milliseid ravimeid määrasid perearstid ja psühhiaatrid depressiooni raviks, küsiti andmed Eesti Haigekassast vastavalt rahvusvahelise haiguste klassifikatsiooni-10 diagnoosi koodide F23 ja F33 järgi 2003 aastal. Määratud ravimite tulemuste analüüsimisel kasutati sagedusjaotuse tabeleid. Seoste statistilist olulisust hinnati hii-ruut-testiga.

### Uurimistöö peamised tulemused

Küsimustikule vastanud perearstidest 185(90%) pidasid depressiivseid häireid omavate patsientidega tegelemist oma kompetentsi kuuluvaks, 180(88%) on valmis tegelema selliste patsientidega. Perearstide motiveerivad tegurid tegelemaks depressiivsete patsientidega, jagasime viide suurde rühma. Nendeks olid: depressiooni sage esinemine esmatasandi meditsiinis; perearstide missjoonitunne – soov ja lootus aidata oma patsiente; positiivne ravi tulemus – paraneb koostöö patsiendiga; perearstide eelised – perearsti konsultatsioonle pääseb kergemini, ta teab omapatsiendi terviseprobleeme paremini võrreldes psühhiaatriga; patsientide mugavus – sageli patsiendid ei soovi konsulteerida psühhiaatriga, voodihaigel patsiendil ei ole võimelik saada psühhiaatri konsultatsiooni. Depressiivse patsiendiga seonduvad probleemid jagasime nelja rühma. Pea-

miseks probleemiks, nagu ka motivatsiooniks, oli depressiooni sage esinemine. Teisteks probleemiks olid depressiooni ravimite kõrge hind, patsientide soovimatus depressiooni diagnoosi omaks võtta ja järgida talle määratud pikaajalist ravi ning perearstide ebapiisavad võimalused ja psühhiaatria alased oskused. Perearstid soovisid paremat koostööd psühhiaatrite ja psühholoogidega, planeerida rohkem aega depressiooni probleemidega patsientidele ning 181(88%) perearstidest soovis täiendkoolitust.

Rahvusvahelise haiguste klassifikatsiooni-10 diagnostilistele kriteeriumitele vastavalt esines depressiooni episood 6 kuu jooksul 258(23.6%) ja viimasel kuul 169(15.4%) perearsti külastanud patsiendil. Kuue kuu depressiooni episood jagunes vastavalt raskusastmetele võrdselt 43% raskeks ja keskmiseks ning kergeks 14%. Kerget depressiooni esines meestel võrreldes naistega rohkem ( $\chi^2 = 12.13$ ; p<0.001).

Depressioon esines sagedamini naistel, madalama haridustasemega isikutel, töötutel, madalama majandusliku toimetulekuga isikutel, madalama enesehinnanguga ja puudega isikutel ja neil, kellel eelneva 6 kuu jooksul oli läbi elatud mõni negatiivne elusündmus. Ei olnud seost vanuse ja depressiooni vahel ja abielulise staatuse vahel. Võrreldes meeste ja naiste abielulisust ja depressiooni, selgus, et naistel ei esine mingit seost, küll aga meestel, kes ei olnud abielus, esines depressiooni rohkem.

CIDI diagnoosis viimase kuu jooksul depressiooni episoodi 162(15.3%), EEK-2 liigitas depressiivseks 300(28.4%) uuringus osalejat. Rasket depressiooni episoodi esines 52%-l, keskmist 38%-l ja kerget 10%-l. Võrreldes CIDIga liigitas EEK-2 vale negatiivseks 2.8% ja vale positiivseks 15.9% uuringus osalenud patsientidest. EEK-2 tundlikkus ja spetsiifilisus oli 0.81 äralõikepunkti >11 juures. Äralõikepunkti langetamine 1 võrra parandas tundlikkust, halvenes aga spetsiifilisus ja äralõikepunkti tõstmisel oli vastupidine efekt.

Sümptomite olulisuse hindamisel osutusid EEK-2 depressiooni skaala sümptomitest oluliseks kurvameelsus, huvi kadumine, enesesüüdistused, üksildustunne ja võimetus rõõmu tunda. Selle sümptomite kombinatsiooni tundlikkus oli 0.81 ja spetsiivilisus 0.82 kõige optimaalsema äralõikepunkti >8 juures ja me nimetasime ta EEK-uus1. EEK-2 depressiooni skaala sümptomitele somaatiliste, käitumulike ja ärevuse sümptomite lisamisel osustusis kõige olulisemateks sümptomid: kurvemeelsus, huvi kadumine, alaväärsustunne, võimetus rõõmu tunda, liigne muretsemine paljude asjade pärast ja puhkamine ei taasta jõudu, mille kombinatsiooni nimetasime EEK-uus2. Uue kombinatsiooni tundlikkus kõige optimaalsema äralõikepunkti >11 juures ei muutunud võrreldes EEK-2-ga, paranes spetsiifilisus 0.85-le ning teised testi statistilised näitajad, mille tulemusena vähenes valepositiivsete arv.

Perearstid nimetasid depressiooni ravis oma esmase valiku preparaadiks SSRI rühm kuuluvaid ravimeid. Eelistuse moodustasid SSRI (69%), TCA (15%), SNRI (7%), RIMA (1%), NaSSA (1%), trankvilisaatorid (5%) ja antipsühhootikumid (2%). Perearstide tegelik ravimite määramine ühtis nende

eelistustega. Eelistuste põhjustena nimetati ravimi efektiivsust, kõrvaltoimete vähesust, toime aega, manustamisviisi ja lihtsust, kaasuvaid kaebusi, ravimi hinda ja teadmisi ravimi omaduste kohta.

Perearstide poolt diagnoositi depressiooni 19 521 isikul 37 029-st 2003 aastal. Depressiooni raviks määratud ravimitest 88% moodustasid antidepressandid, milledest 65% oli määratud perearstide poolt. Antipsühhootikume määrati 1% ravimitest ja meeleolu stabilisaatoreid veel vähem. Perearstid määrasid antidepressantidest SSRI rühma ravimeid 76.8%, TCA rühma ravimeid 19.3% ja teisi ravimrühmi kokku vähem kui 3.5%.

### Järeldused

- 1. Perearstid on valmis tegelema depressiivseid häireid omavate patsientidega ja nad on selleks piisavalt motiveeritud kuna depressioon on oluline ja sage probleem Eesti esmatasandi meditsiinis. Perearste motiveerib soov aidata oma patsiente teades nende teisi haigusi ja probleeme, ning positiivse ravi tulemusena suureneb patsientide usaldus oma arsti vastu veelgi. Peamisteks probleemideks perearstidele seoses depressiooniga on selle sage esinemine patsientide seas, ravi kallidus, patsientide soovimatus diagnoosi omaks võtta ning nende ravi soostumus ja piiratud ajalimiit patsiendi konsultatsiooniks. Selleks, et paremini ravida depressiooni põdevaid patsiente, soovivad perearstid rohkem sptesiifilist täiendõpet ja paremini planeerida aega depressiooni probleemidega patsiendi vastuvõtuks. Oluliseks peavad perearstid psühhiaatrite ja psühholoogidega koostöö paranemist.
- 2. Rahvusvahelise Haiguste Klassifikatsiooni-10 diagnostilistele kriteeriumitele vastav depressiooniepisood esines viimasel kuul igal kuuendal ja eelneva 6 kuu jooksul igal neljandal perearsti külastanud patsiendil. Keskmist ja rasket depressiooni episoodi esines rohkem kui kerget depressiooni episoodi. Eestis esines depressiooni perearsti külastavate patsientide seas rohkem, kui oli oodata, tuginedes varasematele uuringutele.
- 3. Depressiooni esinemine seondus sagedamini naissooga, töötusega, halvema majandusliku olukorraga ja suurema hulga negatiivsete elusündmuste esinemisega viimase kuue kuu jooksul. Samuti oli depressioon sagedasem kroonilisi haigusi põdevata patsientide seas, invaliididel ja neil, kes pidasid oma tervislikku seisundit halvaks.
- 4. Emotsionaalse enesehinnangu küsimustik EEK-2 on kõige sagedamini kasutatav depressiooni skriining instrument Eesti perearstide seas. Kuigi EEK-2 omab head tundlikkust ja spetsiifilisust vastavalt tunnustatud skriinig instrumentide parameetritele, liigitas ta suure hulga tervetest patsientidest haigeteks e. valepositiivseteks. Testides tulemuse parandamiseks erinevaid äralõikepunkte, leidsime, et praegu kasutatav äralõikepunkt >11 annab kõige parema tulemuse testi erinevate näitajate seoses. Skriining instrument EEK-

- 2, mis on välja töötatud psühhiaatriliste patsientide ja populetsiooni baasil, on kasutatav esmatasandi arstiabis.
- 5. Kõige parema sümptomite kombinatsiooni eristamaks depressiivseid patsiente teiste haiguslike probleemidega patsientidest, saime vähendades EEK-2 depressiooni skaala sümptome ja lisades somaatilisi ja käitumuslikke sümptome ning ärevusega seotud sümptomi, muutes sellega küsimustiku lühemaks ja lihtsamaks. Selleks sümptomite kombinatsiooniks osutus: huvi kadumine, kurvameelsus, puhkamine ei taasta jõudu, alaväärsustunne, liigne muretsemine paljude asjade pärast ja võimetus rõõmu tunda. Nende sümptomite baasil tegime uue skriining instrumendi esmatasandi arstiabi jaoks, EEK-uus2, mille tundlikkus võrreldes EEK-2-ga jäi samaks, paranes aga pstesiifilisus.
- 6. Perearstide ravimite eelistus ja ravimite tegelik valik depressiooni raviks ühtisid. Perearstide ravimite valiku põhjused olid samad, mis on üldtunnustatud ravimite selektsiooni faktorid depressiooni ravis.
- 7. Perearstid määrasid depressiooni raviks peamiselt antidepressante ning nende hulgast kõige sagedamini selektiivsete serotoniini tagasihaarde inhibiitorite rühma kuuluvaid ravimeid. Eesti perearstid ravivad depressiooni kooskõlas depressiooni rahvusvaheliste ravijuhistega.

### 8. ACKNOWLEDGMENTS

I wish to express my deepest graduate to everyone who has contributed to the accomplishment of thesis. In particular, I would like to thank:

- Professor Heidi-Ingrid Maaroos, my supervisor and co-worker, for offering me the opportunity to perform the studies, and for providing with the necessary resources and for creating an excellent working atmosphere. Her help and advice throughout the study has allowed me this study. Her brilliant mind, enthusiastic attitude towards research as well as her generous personality, have contributed much to this work.
- Docent Anu Aluoja, the best co-worker and colleague, for help and advice, for sharing her vast and knowledge and experience in psychology.
- All co-authors, for fruitful partnership.
- All my colleagues from the Department of Polyclinic and Family Medicine, for creating a friendly atmosphere, and for their constant support and enjoyable discussions. Special thanks to my colleagues Ruth Kalda and Marje Oona.
- Mrs. Maie Thetloff for advice in statistics.
- Mrs. Ester Jaigma for the profound and competent technical revision of the manuscript.
- All Estonian work team in PREDICT study.
- All my colleagues from the Ädala Family Doctors Centre for good cooperation.
- To my family for their patient and love during the preparation of this thesis.
- The present study was supported by the Estonian Science Foundation grant No 5696 and by Estonian Target Funding topic No 0821.
- I am also grateful for support from the European Commission's Fifth Framework, grant number Predict-QL4-CT2002-00683 and we thank all PREDICT group members.

# **PUBLICATIONS**

# **CURRICULUM VITAE**

# Pille Ööpik

Citizenship: Estonian Born: Nov 26, 1960 in Tallinn, Estonia 1 child

Home address: Põllu tee 20, 76901 Tabasalu, Harjumaa, Estonia Office address: Dep of Polyclinic and Family Medicine, University of Tartu

Puusepa 1a, 50406 Tartu, Estonia Phone: +372 731 9215 Fax: +372 731 9213 E-mail: pille.oopik@ut.ee

### **Education**

1972–1979	Kose High School
1979–1985	University of Tartu Faculty of Medicine,
	Department of Pediatrics
1986–1987	Tartu Children's Hospital, internship
1990-1993	University of Tartu, vocational training in Family Medicine
2001-	University of Tartu, Department of Polyclinic and Family
	Medicine, doctoral student

### **Professional employment**

1987–1994	Tartu Children's Hospital, pediatrician
1994–1998	Estonian Health Insurance Fund, counsellor
1998-2000	Department of Social and Healthcare, Tallinn City Government,
	specialist
1995–	Ädala Family Centre, family doctor
2005-	University of Tartu, assistant

### Scientific work

Resears topics: Depression in primary care, prevalence of disease, symptoms of depression, screening of depression, risk factors of depression, prevention and treatment of depression.

A total of 14 publications in peer reviewed international journals and in others journals.

Distinctions: II–III price for presentation on the Conference Faculty of Medicine of University of Tartu, 2005

# **Practipation in professional organizations**

Estonian Society of Family Doctors Society of Family Doctors in Tallinn

### **ELULOOKIRJELDUS**

# Pille Ööpik

Kodakondsus: Eesti Sünniaeg: 26, november 1960

1 laps

Kodune aadress: Põllu tee 20, 76901 Tabasalu, Harjumaa, Essti Aadress tööl: Tartu Ülikooli polikliiniku ja perearstiteaduse õppetool,

Puusepa 1a, 50406 Tartu, Eesti Telefon: +372 731 9215 Fax: +372 731 9213 Meiliaadress: pille.oopik@ut.ee

### Haridus

1972–1979	Kose Keskkool
1979–1985	Tartu Ülikool arstiteaduskond, pediaatria osakond
1986–1987	Tartu Lastekliinik, intern
1990-1993	Tartu Ülikool arstiteaduskond, spetsiaaliseerumiskursus perearsti
	erialal
2001-	Tartu Ülikool polikliiniku ja perearstiteaduse õppetool,
	doktorantuur

### Erialane teenistuskäik

1987–1994	Tartu Lastekliinik, pediaater
1994–1998	Eesti Keskhaigekassa, arst-spetsialist
1998-2000	Tallinna Sotsiaal- ja Tervishoiuamet, peaspetsialist
1995–	Ädala Perearstikeskus, perearst
2005-	Tartu Ülikool polikliiniku ja perearstiteaduse õppetool, assistent

### **Teadustegevus**

Peamised uurimisvaldkonnad: Depressioon esmatasandi meditsiinis: haiguse levik, avaldumise sümptomid, depressiooni riskifaktorid, depressiooni skriinimine, profülaktika ja ravi, depressiooniga seodnuvad probleemid perearstide igapäevases töös.

Ilmunud 14 publikatsiooni rahvusvahelise levikuga eelretsenseeritavates ajakirjades ja teistes ajakirjades.

Uurimistöö on pälvinud auhindu: Tartu Ülikool, Arstiteaduskond; preemia: Arstiteaduskonna aastakonverents 2005, diplom suuliste ettekannete eest, II–III koht; tunnustatud töö nimetus: Depressioon esmatasandi meditsiinis: PREDICT uuringu esimese etapi tulemused

# Kuuluvus erialaorganisatsioonidesse

Eesti perearstide Selts Tallinna Perearstide Selts

# DISSERTATIONES MEDICINAE UNIVERSITATIS TARTUENSIS

- 1. **Heidi-Ingrid Maaroos.** The natural course of gastric ulcer in connection with chronic gastritis and *Helicobacter pylori*. Tartu, 1991.
- 2. **Mihkel Zilmer.** Na-pump in normal and tumorous brain tissues: Structural functional a. tumorigenesis aspects. Tartu, 1991.
- 3. **Eero Vasar.** Role of cholecystokinin receptors in the regulation of behaviour and in the action of haloperidol and diazepam. Tartu, 1992.
- 4. **Tiina Talvik.** Hypoxic-ischaemic brain damage in neonates (clinical, biochemical and brain computed tomographical investigation). Tartu, 1992.
- 5. **Ants Peetsalu.** Vagotomy in duodenal ulcer disease: A study of gastric acidity, serum pepsinogen I, gastric mucosal histology and *Helicobacter pylori*. Tartu, 1992.
- 6. **Marika Mikelsaar.** Evaluation of the gastrointestinal microbial ecosystem in health and disease. Tartu, 1992.
- 7. **Hele Everaus.** Immuno-hormonal interactions in chronic lymphocytic leukaemia and multiple myeloma. Tartu, 1993.
- 8. **Ruth Mikelsaar.** Etiological factors of diseases in genetically consulted children and newborn screening: dissertation for the commencement of the degree of doctor of medical sciences. Tartu, 1993.
- 9. **Agu Tamm.** On metabolic action of intestinal microflora: clinical aspects. Tartu, 1993.
- 10. **Katrin Gross.** Multiple sclerosis in South-Estonia (epidemiological and computed tomographical investigations). Tartu, 1993.
- 11. **Oivi Uibo.** Childhood coeliac disease in Estonia: occurrence, screening, diagnosis and clinical characterization. Tartu, 1994.
- 12. **Viiu Tuulik.** The functional disorders of central nervous system of chemistry workers. Tartu, 1994.
- 13. **Margus Viigimaa.** Primary haemostasis, antiaggregative and anticoagulant treatment of acute myocardial infarction. Tartu, 1994.
- 14. **Rein Kolk.** Atrial versus ventricular pacing in patients with sick sinus syndrome. Tartu, 1994.
- 15. **Toomas Podar.** Incidence of childhood onset type 1 diabetes mellitus in Estonia. Tartu. 1994.
- 16. **Kiira Subi.** The laboratory surveillance of the acute respiratory viral infections in Estonia. Tartu, 1995.
- 17. **Irja Lutsar.** Infections of the central nervous system in children (epidemiologic, diagnostic and therapeutic aspects, long term outcome). Tartu, 1995.
- 18. **Aavo Lang.** The role of dopamine, 5-hydroxytryptamine, sigma and NMDA receptors in the action of antipsychotic drugs. Tartu, 1995.

- 19. **Andrus Arak.** Factors influencing the survival of patients after radical surgery for gastric cancer. Tartu, 1996.
- 20. **Tõnis Karki.** Quantitative composition of the human lactoflora and method for its examination. Tartu, 1996.
- 21. **Reet Mändar.** Vaginal microflora during pregnancy and its transmission to newborn. Tartu, 1996.
- 22. **Triin Remmel.** Primary biliary cirrhosis in Estonia: epidemiology, clinical characterization and prognostication of the course of the disease. Tartu, 1996.
- 23. **Toomas Kivastik.** Mechanisms of drug addiction: focus on positive reinforcing properties of morphine. Tartu, 1996.
- 24. **Paavo Pokk.** Stress due to sleep deprivation: focus on GABA<sub>A</sub> receptor-chloride ionophore complex. Tartu, 1996.
- 25. **Kristina Allikmets.** Renin system activity in essential hypertension. Associations with atherothrombogenic cardiovascular risk factors and with the efficacy of calcium antagonist treatment. Tartu, 1996.
- 26. **Triin Parik.** Oxidative stress in essential hypertension: Associations with metabolic disturbances and the effects of calcium antagonist treatment. Tartu, 1996.
- 27. **Svetlana Päi.** Factors promoting heterogeneity of the course of rheumatoid arthritis. Tartu, 1997.
- 28. **Maarike Sallo.** Studies on habitual physical activity and aerobic fitness in 4 to 10 years old children. Tartu, 1997.
- 29. **Paul Naaber.** Clostridium difficile infection and intestinal microbial ecology. Tartu, 1997.
- 30. **Rein Pähkla.** Studies in pinoline pharmacology. Tartu, 1997.
- 31. Andrus Juhan Voitk. Outpatient laparoscopic cholecystectomy. Tartu, 1997.
- 32. **Joel Starkopf.** Oxidative stress and ischaemia-reperfusion of the heart. Tartu, 1997.
- 33. **Janika Kõrv.** Incidence, case-fatality and outcome of stroke. Tartu, 1998.
- 34. Ülla Linnamägi. Changes in local cerebral blood flow and lipid peroxidation following lead exposure in experiment. Tartu, 1998.
- 35. **Ave Minajeva.** Sarcoplasmic reticulum function: comparison of atrial and ventricular myocardium. Tartu, 1998.
- 36. **Oleg Milenin.** Reconstruction of cervical part of esophagus by revascularised ileal autografts in dogs. A new complex multistage method. Tartu, 1998.
- 37. **Sergei Pakriev.** Prevalence of depression, harmful use of alcohol and alcohol dependence among rural population in Udmurtia. Tartu, 1998.
- 38. **Allen Kaasik.** Thyroid hormone control over β-adrenergic signalling system in rat atria. Tartu, 1998.
- 39. **Vallo Matto.** Pharmacological studies on anxiogenic and antiaggressive properties of antidepressants. Tartu, 1998.

- 40. **Maire Vasar.** Allergic diseases and bronchial hyperreactivity in Estonian children in relation to environmental influences. Tartu. 1998.
- 41. **Kaja Julge.** Humoral immune responses to allergens in early childhood. Tartu, 1998.
- 42. **Heli Grünberg.** The cardiovascular risk of Estonian schoolchildren. A cross-sectional study of 9-, 12- and 15-year-old children. Tartu, 1998.
- 43. **Epp Sepp.** Formation of intestinal microbial ecosystem in children. Tartu, 1998.
- 44. **Mai Ots.** Characteristics of the progression of human and experimental glomerulopathies. Tartu, 1998.
- 45. **Tiina Ristimäe.** Heart rate variability in patients with coronary artery disease. Tartu, 1998.
- 46. **Leho Kõiv.** Reaction of the sympatho-adrenal and hypothalamo-pituitary-adrenocortical system in the acute stage of head injury. Tartu, 1998.
- 47. **Bela Adojaan.** Immune and genetic factors of childhood onset IDDM in Estonia. An epidemiological study. Tartu, 1999.
- 48. **Jakov Shlik.** Psychophysiological effects of cholecystokinin in humans. Tartu. 1999.
- 49. **Kai Kisand.** Autoantibodies against dehydrogenases of  $\alpha$ -ketoacids. Tartu, 1999.
- 50. **Toomas Marandi.** Drug treatment of depression in Estonia. Tartu, 1999.
- 51. Ants Kask. Behavioural studies on neuropeptide Y. Tartu, 1999.
- 52. **Ello-Rahel Karelson.** Modulation of adenylate cyclase activity in the rat hippocampus by neuropeptide galanin and its chimeric analogs. Tartu, 1999.
- 53. **Tanel Laisaar.** Treatment of pleural empyema special reference to intrapleural therapy with streptokinase and surgical treatment modalities. Tartu, 1999.
- 54. **Eve Pihl.** Cardiovascular risk factors in middle-aged former athletes. Tartu, 1999.
- 55. **Katrin Õunap.** Phenylketonuria in Estonia: incidence, newborn screening, diagnosis, clinical characterization and genotype/phenotype correlation. Tartu, 1999.
- 56. **Siiri Kõljalg.** *Acinetobacter* an important nosocomial pathogen. Tartu, 1999.
- 57. **Helle Karro.** Reproductive health and pregnancy outcome in Estonia: association with different factors. Tartu, 1999.
- 58. **Heili Varendi.** Behavioral effects observed in human newborns during exposure to naturally occurring odors. Tartu, 1999.
- 59. **Anneli Beilmann.** Epidemiology of epilepsy in children and adolescents in Estonia. Prevalence, incidence, and clinical characteristics. Tartu, 1999.
- 60. **Vallo Volke.** Pharmacological and biochemical studies on nitric oxide in the regulation of behaviour. Tartu, 1999.

- 61. **Pilvi Ilves.** Hypoxic-ischaemic encephalopathy in asphyxiated term infants. A prospective clinical, biochemical, ultrasonographical study. Tartu, 1999.
- 62. **Anti Kalda.** Oxygen-glucose deprivation-induced neuronal death and its pharmacological prevention in cerebellar granule cells. Tartu, 1999.
- 63. **Eve-Irene Lepist.** Oral peptide prodrugs studies on stability and absorption. Tartu, 2000.
- 64. **Jana Kivastik.** Lung function in Estonian schoolchildren: relationship with anthropometric indices and respiratory symptomas, reference values for dynamic spirometry. Tartu, 2000.
- 65. **Karin Kull.** Inflammatory bowel disease: an immunogenetic study. Tartu, 2000.
- 66. **Kaire Innos.** Epidemiological resources in Estonia: data sources, their quality and feasibility of cohort studies. Tartu, 2000.
- 67. **Tamara Vorobjova.** Immune response to *Helicobacter pylori* and its association with dynamics of chronic gastritis and epithelial cell turnover in antrum and corpus. Tartu, 2001.
- 68. **Ruth Kalda.** Structure and outcome of family practice quality in the changing health care system of Estonia. Tartu, 2001.
- 69. **Annika Krüüner.** *Mycobacterium tuberculosis* spread and drug resistance in Estonia. Tartu, 2001.
- 70. **Marlit Veldi.** Obstructive Sleep Apnoea: Computerized Endopharyngeal Myotonometry of the Soft Palate and Lingual Musculature. Tartu, 2001.
- 71. **Anneli Uusküla.** Epidemiology of sexually transmitted diseases in Estonia in 1990–2000. Tartu, 2001.
- 72. **Ade Kallas.** Characterization of antibodies to coagulation factor VIII. Tartu, 2002.
- 73. **Heidi Annuk.** Selection of medicinal plants and intestinal lactobacilli as antimicrobil components for functional foods. Tartu, 2002.
- 74. **Aet Lukmann**. Early rehabilitation of patients with ischaemic heart disease after surgical revascularization of the myocardium: assessment of health-related quality of life, cardiopulmonary reserve and oxidative stress. A clinical study. Tartu, 2002.
- 75. **Maigi Eisen.** Pathogenesis of Contact Dermatitis: participation of Oxidative Stress. A clinical biochemical study. Tartu, 2002.
- 76. **Piret Hussar.** Histology of the post-traumatic bone repair in rats. Elaboration and use of a new standardized experimental model bicortical perforation of tibia compared to internal fracture and resection osteotomy. Tartu, 2002.
- 77. **Tõnu Rätsep.** Aneurysmal subarachnoid haemorrhage: Noninvasive monitoring of cerebral haemodynamics. Tartu, 2002.
- 78. **Marju Herodes.** Quality of life of people with epilepsy in Estonia. Tartu, 2003.

- 79. **Katre Maasalu.** Changes in bone quality due to age and genetic disorders and their clinical expressions in Estonia. Tartu, 2003.
- 80. **Toomas Sillakivi.** Perforated peptic ulcer in Estonia: epidemiology, risk factors and relations with *Helicobacter pylori*. Tartu, 2003.
- 81. **Leena Puksa.** Late responses in motor nerve conduction studies. F and A waves in normal subjects and patients with neuropathies. Tartu, 2003.
- 82. **Krista Lõivukene**. *Helicobacter pylori* in gastric microbial ecology and its antimicrobial susceptibility pattern. Tartu, 2003.
- 83. **Helgi Kolk.** Dyspepsia and *Helicobacter pylori* infection: the diagnostic value of symptoms, treatment and follow-up of patients referred for upper gastrointestinal endoscopy by family physicians. Tartu, 2003.
- 84. **Helena Soomer.** Validation of identification and age estimation methods in forensic odontology. Tartu, 2003.
- 85. **Kersti Oselin.** Studies on the human MDR1, MRP1, and MRP2 ABC transporters: functional relevance of the genetic polymorphisms in the *MDR1* and *MRP1* gene. Tartu, 2003.
- 86. **Jaan Soplepmann.** Peptic ulcer haemorrhage in Estonia: epidemiology, prognostic factors, treatment and outcome. Tartu, 2003.
- 87. **Margot Peetsalu.** Long-term follow-up after vagotomy in duodenal ulcer disease: recurrent ulcer, changes in the function, morphology and *Helico-bacter pylori* colonisation of the gastric mucosa. Tartu, 2003.
- 88. **Kersti Klaamas.** Humoral immune response to *Helicobacter pylori* a study of host-dependent and microbial factors. Tartu, 2003.
- 89. **Pille Taba.** Epidemiology of Parkinson's disease in Tartu, Estonia. Prevalence, incidence, clinical characteristics, and pharmacoepidemiology. Tartu, 2003.
- 90. **Alar Veraksitš**. Characterization of behavioural and biochemical phenotype of cholecystokinin-2 receptor deficient mice: changes in the function of the dopamine and endopioidergic system. Tartu, 2003.
- 91. **Ingrid Kalev.** CC-chemokine receptor 5 (CCR5) gene polymorphism in Estonians and in patients with Type I and Type II diabetes mellitus. Tartu, 2003.
- 92. **Lumme Kadaja.** Molecular approach to the regulation of mitochondrial function in oxidative muscle cells. Tartu, 2003.
- 93. **Aive Liigant**. Epidemiology of primary central nervous system tumours in Estonia from 1986 to 1996. Clinical characteristics, incidence, survival and prognostic factors. Tartu, 2004.
- 94. **Andres, Kulla.** Molecular characteristics of mesenchymal stroma in human astrocytic gliomas. Tartu, 2004.
- 95. **Mari Järvelaid.** Health damaging risk behaviours in adolescence. Tartu, 2004.
- 96. **Ülle Pechter.** Progression prevention strategies in chronic renal failure and hypertension. An experimental and clinical study. Tartu, 2004.

- 97. **Gunnar Tasa.** Polymorphic glutathione S-transferases biology and role in modifying genetic susceptibility to senile cataract and primary open angle glaucoma. Tartu, 2004.
- 98. **Tuuli Käämbre.** Intracellular energetic unit: structural and functional aspects. Tartu, 2004.
- 99. **Vitali Vassiljev.** Influence of nitric oxide syntase inhibitors on the effects of ethanol after acute and chronic ethanol administration and withdrawal. Tartu, 2004.
- 100. **Aune Rehema.** Assessment of nonhaem ferrous iron and glutathione redox ratio as markers of pathogeneticity of oxidative stress in different clinical groups. Tartu, 2004.
- 101. **Evelin Seppet.** Interaction of mitochondria and ATPases in oxidative muscle cells in normal and pathological conditions. Tartu, 2004.
- 102. **Eduard Maron.** Serotonin function in panic disorder: from clinical experiments to brain imaging and genetics. Tartu, 2004.
- 103. **Marje Oona.** *Helicobacter pylori* infection in children: epidemiological and therapeutic aspects. Tartu, 2004.
- 104. **Kersti Kokk.** Regulation of active and passive molecular transport in the testis. Tartu, 2005.
- 105. **Vladimir Järv.** Cross-sectional imaging for pretreatment evaluation and follow-up of pelvic malignant tumours. Tartu, 2005.
- 106. **Andre Õun.** Epidemiology of adult epilepsy in Tartu, Estonia. Incidence, prevalence and medical treatment. Tartu, 2005.
- 107. **Piibe Muda.** Homocysteine and hypertension: associations between homocysteine and essential hypertension in treated and untreated hypertensive patients with and without coronary artery disease. Tartu, 2005.
- 108. **Külli Kingo.** The interleukin-10 family cytokines gene polymorphisms in plaque psoriasis. Tartu, 2005.
- 109. **Mati Merila.** Anatomy and clinical relevance of the glenohumeral joint capsule and ligaments. Tartu, 2005.
- 110. **Epp Songisepp**. Evaluation of technological and functional properties of the new probiotic *Lactobacillus fermentum* ME-3. Tartu, 2005.
- 111. **Tiia Ainla.** Acute myocardial infarction in Estonia: clinical characteristics, management and outcome. Tartu, 2005.
- 112. **Andres Sell.** Determining the minimum local anaesthetic requirements for hip replacement surgery under spinal anaesthesia a study employing a spinal catheter. Tartu, 2005.
- 113. **Tiia Tamme.** Epidemiology of odontogenic tumours in Estonia. Pathogenesis and clinical behaviour of ameloblastoma. Tartu, 2005.
- 114. **Triine Annus**. Allergy in Estonian schoolchildren: time trends and characteristics. Tartu, 2005.
- 115. **Tiia Voor.** Microorganisms in infancy and development of allergy: comparison of Estonian and Swedish children. Tartu, 2005.

- 116. **Priit Kasenõmm.** Indicators for tonsillectomy in adults with recurrent tonsillitis clinical, microbiological and pathomorphological investigations. Tartu, 2005.
- 117. **Eva Zusinaite.** Hepatitis C virus: genotype identification and interactions between viral proteases. Tartu, 2005.
- 118. **Piret Kõll.** Oral lactoflora in chronic periodontitis and periodontal health. Tartu, 2006.
- 119. **Tiina Stelmach.** Epidemiology of cerebral palsy and unfavourable neurodevelopmental outcome in child population of Tartu city and county, Estonia Prevalence, clinical features and risk factors. Tartu, 2006.
- 120. **Katrin Pudersell.** Tropane alkaloid production and riboflavine excretion in the field and tissue cultures of henbane (*Hyoscyamus niger* L.). Tartu, 2006.
- 121. **Külli Jaako.** Studies on the role of neurogenesis in brain plasticity. Tartu, 2006
- 122. **Aare Märtson.** Lower limb lengthening: experimental studies of bone regeneration and long-term clinical results. Tartu, 2006.
- 123. **Heli Tähepõld.** Patient consultation in family medicine. Tartu, 2006.
- 124. **Stanislav Liskmann.** Peri-implant disease: pathogenesis, diagnosis and treatment in view of both inflammation and oxidative stress profiling. Tartu, 2006.
- 125. **Ruth Rudissaar.** Neuropharmacology of atypical antipsychotics and an animal model of psychosis. Tartu, 2006.
- 126. **Helena Andreson.** Diversity of *Helicobacter pylori* genotypes in Estonian patients with chronic inflammatory gastric diseases. Tartu, 2006.
- 127. **Katrin Pruus.** Mechanism of action of antidepressants: aspects of serotoninergic system and its interaction with glutamate. Tartu, 2006.
- 128. **Priit Põder.** Clinical and experimental investigation: relationship of ischaemia/reperfusion injury with oxidative stress in abdominal aortic aneurysm repair and in extracranial brain artery endarterectomy and possibilities of protection against ischaemia using a glutathione analogue in a rat model of global brain ischaemia. Tartu, 2006.
- 129. **Marika Tammaru.** Patient-reported outcome measurement in rheumatoid arthritis. Tartu, 2006.
- 130. **Tiia Reimand.** Down syndrome in Estonia. Tartu, 2006.
- 131. **Diva Eensoo.** Risk-taking in traffic and Markers of Risk-Taking Behaviour in Schoolchildren and Car Drivers. Tartu, 2007.
- 132. **Riina Vibo.** The third stroke registry in Tartu, Estonia from 2001 to 2003: incidence, case-fatality, risk factors and long-term outcome. Tartu, 2007.
- 133. **Chris Pruunsild.** Juvenile idiopathic arthritis in children in Estonia. Tartu, 2007.
- 134. **Eve Õiglane-Šlik.** Angelman and Prader-Willi syndromes in Estonia. Tartu, 2007.

135. <b>Kadri Haller.</b> Antibodies to follicle stimulating hormone. Significance in female infertility. Tartu, 2007.	