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Productivity loss due to menstruation-related symptoms: a nationwide cross-sectional survey among 32,748 women

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4	2	continuel outwork of an 22 749 woman
5 6	2	sectional survey among 32,748 women
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Abstract **Objective** To evaluate age-dependent productivity loss caused by menstruation-related symptoms, measured in absenteeism (time away from work or school) and presenteeism (productivity loss while present at work/school). Methods from July to October 2017.

Design/Setting: Internet-based, cross-sectional survey conducted in The Netherlands

Participants: 32,748 women aged 15 to 45 years, recruited through social-media.

Outcome measures: Lost productivity in days, divided into absenteeism and presenteeism; impact of menstrual symptoms; reasons women give to their employer or school when calling in sick; and women's preferences regarding conditions at work and school.

Results

A total of 13.8% of all women reported absenteeism during their menstrual periods with 3.4% reporting absenteeism every or almost every menstrual cycle. The mean absenteeism related to a woman's period was 1.2 days per year. A total of 80.7% of the respondents reported presenteeism and decreased productivity a mean of 23.2 days per year. An average productivity loss of 33% resulted in a mean of 8.9 days of total lost productivity per year. Women under 21 were more likely to report absenteeism due to menstruation-related symptoms (OR 3.3, 95% confidence interval 3.1 to 3.6). When women called in sick due to their periods, only 20.1% told

57 their employer or school mentor that their absence was due to menstrual complaints.

Notably, 67.7% of the participants wished they had greater flexibility in their tasks and

59 working hours at work or school during their periods.

61 <u>Conclusions</u>

Menstruation-related symptoms cause a great deal of lost productivity, and presenteeism is a bigger contributor to this than absenteeism. There is an urgent need for more focus on the impact of these symptoms, especially in adolescents, for discussions of treatment options with women of all ages, and, ideally, more flexibility for women who work or go to school.

68 Strengths and limitations of this study

- This is the largest cohort study to analyse the impact of menstruation-related symptoms on work and school productivity.
- The survey was performed among the general female population and is consequently not per se related to one specific gynaecological condition.
 - Due to the way of recruitment of participants, there may have been some degree of selection bias.
 - The generalisability of the study may be limited to employment and school systems comparable to the Dutch.

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78 Background

Menstruation-related symptoms (MRSs) are diverse and widespread among women. Symptoms include dysmenorrhea, heavy menstrual bleeding and premenstrual mood disturbances with reported prevalence of 45% to 90%, 14% to 25%, and 20% to 29% respectively.¹⁻³ Studies show that women with MRSs have lower scores on several domains of quality of life such as general health and physical, mental, social, and occupational functioning during their periods.^{1 4-7} Furthermore, these symptoms may create considerable financial burdens on patients and their families as well as on society.^{5 6 8-12} Such financial burdens are related to the costs of visits to the doctor, over-the-counter drugs, and medical or surgical treatment. However, costs related to productivity loss could be the largest cost driver. Productivity costs are defined as costs associated with paid and unpaid production loss and the replacement of productive people due to illness or disability.¹³ Productivity costs can be divided into costs related to absenteeism and costs related to presenteeism. Absenteeism represents the total amount of time off work or away from school, and presenteeism represents the loss of productivity while present at a job or school.

Although the literature is scarce and the results are variable, studies on specific patient groups generally show that MRSs can cause absenteeism.¹⁴⁻¹⁶ Research on the association between MRSs and presenteeism is even more limited. It has been suggested that research into possible impairments in guality of life caused by menstrual symptoms should not focus on single symptoms but rather on a complex of symptoms that vary widely but that are all related to the menstrual cycle. This complex includes both standard symptoms, like heavy menstrual bleeding and abdominal cramps, and also less common symptoms, like nausea and cold sweats. 17 18

Taking all symptoms into account, it seems likely that the real impact of MRSs is underestimated in the general population. Despite being almost two decades into the 21st century, discussions about MRSs may still be rather taboo. This survey-based exploratory study aimed to quantify the burden of MRSs in the general female population, with burden defined as the number of lost days at work or school due to absenteeism and presenteeism. Furthermore, it was aimed to study the impact of specific symptoms on absenteeism and presenteeism.

111 Methods

This cross-sectional study consisted of an online survey that was distributed from July 12 to October 11, 2017. Approval for this study was obtained from the local medical ethics committee "Commissie Mensgebonden Onderzoek (CMO)" under number file number 2017-3387 on July 12th 2017. All data were anonymously collected and stored under the privacy rules of the Radboud University Medical Center. Patients gave informed consent when they initiated the questionnaire.

119 Patient and Public Involvement

A group of women, including several members of the Dutch Patient Endometriosis Foundation, was involved in the conduct of this study at several stages; i.e. in the development and dissemination of the questionnaire and in the analysis and interpretation of the results. One of the authors of this manuscript, BD, is the chair of the Dutch Patient Endometriosis Foundation. Additional contributions are noted in the Acknowledgements section.

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127 Questionnaire development

The questionnaire had several parts, and Appendix 1 provides details about the questions. Part 1 consisted of questions about each woman's basic characteristics. Part 2 had guestions about menstrual symptoms, and part 3 had guestions related to absenteeism and presenteeism. Adaptive questioning was used with a maximum of 6 questions per page. Additional questions about absenteeism and presenteeism were included that were based on the Productivity Cost Questionnaire from the Institute for Medical Technology Assessment (iMTA-PCQ).¹⁹ We modified the iMTA-PCQ-recommended recall period of four weeks so that it was in line with the relevant time period for this study and so we could include multiple menstrual periods. Visual analogue scales (VAS) were used to quantify the amount of pain, or the intensity of the symptom, and the impairment due to pain or the other symptom. Presenteeism was measured by asking women to what extent they were able to be as productive as possible compared to a day without MRSs. This was scored on a scale from 0 to 10, with 0 being totally unproductive and 10 fully productive.

143 Target population and recruitment

The study population comprised of women between 15 and 45 years old. The upper age limit was chosen to avoid interference from menopausal symptoms; the lower to have a time margin after the average menarche age, since it is known that the first periods are irregular and often accompanied with discomfort and uncertainty. A large number of women were approached with the aim of obtaining a cohort that was representative of the general female population in terms of level of education, medical history, and/or gynaecological diagnosis. Women were invited to complete a survey using an online questionnaire tool²⁰ through a campaign on social media

platforms Facebook and Twitter. Patient organisations, colleagues, and visitors of the Facebook page of one of the authors (TN) were asked to share the link to the questionnaire in order to reach the widest possible audience.

On July 12, 2017, a link to the questionnaire was posted on Facebook and Twitter through the account of one of the authors (TN). In the post both women with and without MRSs were encouraged to fill in the guestionnaire. Within 24 hours of the first posting on social media, over 6,000 respondents had filled in the questionnaire, and by July 18, there were 15,000 respondents, which was announced by a re-post of the link to the questionnaire. A third post was made on Facebook and Twitter on September 16, 2017, to reach women who may have been on holiday when the first P 101 posts were created.

Data analysis

The outcome measures were presented in a descriptive way; we used valid percentages in case of missing values where necessary. We distinguished between women who were mainly working or mainly studying. Therefore, we present these data for two groups i.e. for women who worked more than 5 hours per week ("working group") and for women who studied more than 5 hours per week ("studying group").

We used binary logistic regression to calculate odds ratios. Absenteeism and presenteeism were used as dependant variables. As independent variables we used the following parameters: women younger than 21 versus women aged 21 and older, smoking yes or no, reports of absenteeism not related to MRSs, educational level, the use of oral contraception and the use of an levonorgestrel-releasing intra uterine

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device. All independent variables were used in an univariate as well as a multivariate
analysis. We also studied the association between pain scores and both
absenteeism and presenteeism, given that the literature shows that pain scores of 0
to 4, 5 to 6, and 7 or higher have a different impact on activity, mood, and sleep.^{21 22}
Analyses were performed using IBM SPSS Statistics version 22.00.

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182 Assumptions and transformation of the original data

To present data on level of education in an international format we had to transform
the original data, which was based on the Dutch school system.²³

With regard to the evaluation of absenteeism and presenteeism, "the guideline for economic evaluations in healthcare in the Netherlands" was used.²⁴ A week of full time work or study accounted for 36 hours, one day was 8 hours, and in a year, women were able to work 1558 hours when they were working full time.

To calculate the percentages for absenteeism, one day of absenteeism accounted for 8 hours of lost productivity. When a woman reported to study or work more than 40 hours per week, we transformed these hours to 40 hours. We made a few other transformations for categorical data. For absenteeism related to MRSs, the category "more than three days per cycle" was considered to be 4 days per cycle. For absenteeism that was not related to MRSs, the category "more than ten days in the past six months" was considered to be 11 days in the past six months.

To present yearly data, we multiplied some of these data based on the original recall period. The number of days for absenteeism related to MRSs was based on days per cycle, which were therefore multiplied by 12.7 based on the reported average menstrual cycle of 28.8 days, see table 1. These values were then multiplied by one

> if the woman reported that she called in sick "every period", 0.75 if she reported "almost every period", 0.5 if she reported "half of all periods", and 0.25 if she reported calling in sick "only once in a while". Values for absenteeism that was not related to MRS were based on a recall period of 6 months and were therefore multiplied by two in order to obtain the number of days per year. The values for presenteeism were based on a recall period of three months and were therefore multiplied by four.

207 Results

A total of 44,173 women initiated the questionnaire. We excluded participants who did not report a date of birth or whose age did not fulfil the inclusion criteria (figure 1). There were no duplicates of IP addresses. Women who did not answer questions related to absenteeism and presenteeism were excluded. Furthermore, cases with extreme outliers (e.g. 10,000,000 days of presenteeism in three months or 140 changes of sanitary pads a day) were excluded. This resulted in a total of 32,748 women in the final analysis.

Table 1 summarizes the basic characteristics of the participants. We found that 45.4% (n=14,839) had visited a doctor for menstrual complaints in the past, with a total of 3017 (9.2%) women reporting a diagnosis of a menstrual disorder, such as endometriosis or fibroids.

The mean age of women in the working group was higher than the mean ager of women in the studying group. The mean number of working hours per week in the working group was 27.0 (SD 11.4), and the mean number of study hours in the studying group was 27.4 (SD 12.1). A total of 7,335 women (22.4%) reported both working and studying more than five hours per week. In this group, 3,001 women

were working more than 16 hours a week, and 5,284 women in the study group were

studying more than 16 hours a week.

	Number (percentage)	Mean ± SD	Media
Age, years		28.6±8.6	28
15-19	6,141 (18.8%)		
20-24	6,118 (18.7%)		
25-29	5,825 (17.8%)		
30-34	5,483 (16.7%)		
35-40	4,687 (14.3%)		
41-45	4,494 (13.7%)		
Level of education	1,101 (1011 /0)		
Low	4,020 (12.3%)		
Medium	12,335 (37.9%)		
High	16,229 (49.8%)		
Hours/week	10,220 (40.070)		
Paid work		21.7±14.7	24
Study		7.4±13.6	0
Voluntary work		0.8±3.1	0
Menstrual cycle		0.0±3.1	0
Regular cycle	25,717 (78.5%)		
Duration	25,717 (70.570)	28.8±3.0	28
Amenorrhoea due to LG-IUD/OC	2 675 (11 20/)	20.0±3.0	20
	3,675 (11.2%)		
Irregular, variation >10 days per cycle	2,495 (7.6%)		
Do not know	861 (2.6%)	- 4.4.0	
Days with blood loss per cycle		5.4±1.6	5
Visited a doctor for MRSs			
No	17,873 (54.6%)		
Yes, general practitioner	10,141 (31.0%)		
Yes, gynaecologist	4,698 (14.4%)		
Diagnosis for MRSs*		4	
Νο	29,731 (90.8%)		
Yes	3,017 (9.2%) 🚿		
Endometriosis	1,120 (3.4%)		
PCOS	588 (1.8%)		
Adenomyosis	103 (0.3%)		
Fibroids	275 (0.8%)		
Other	1,901 (5.8%)		
Contraception*			
Hormonal contraception	11,993 (36.6%)		
OC	8,650 (26.4%)		
LG-IUD	2,752 (8.4%)		
Other hormonal: injection, transdermal etc.	882 (2.7%)		
No hormonal contraception	20,755 (63.4%)		
Cu-IUD	771 (2.4%)		
Female sterilisation	423 (1.3%)		
No female contraception	19,639 (60.0%)		
			1

Paid work >5 hr a week	26,104 (79.7%)		
Age		29.7±8.3	29
Hours of paid work/week		27.0±11.4	28
Hours of study/week		7.5±13.4	0
Study >5 hr a week	8,764 (26.8%)		
Age		22.0±6.2	20
Hours of paid work/week		15.5±11.3	12
Hours spent on studying/week		27.4±12.1	30

 Table 1 | Basic characteristics of study participants (n=32,748) Mean duration of cycle based on women with a regular cycle. SD = standard deviation, MRSs = menstruation-related symptoms, PCOS = polycystic ovary syndrome, OC = oral contraceptive, LG-IUD = levonorgestrel-releasing intrauterine device, Cu-IUD = copper intrauterine device. *More than one possible answer.

233 Absenteeism

Table 2 shows the results on absenteeism due to MRSs. Although 13.8% of the women (n=4,514) reported absenteeism due to MRSs, only 1,108 women (3.4%) reported absenteeism every cycle or almost every cycle. The percentage of absenteeism in every cycle or almost every cycle was 2.4% in the working group and 4.5% in the studying group. The mean absenteeism due to MRSs was 0.9 days per year for the working group and 1.6 day per year for the study group.

We also calculated the mean total absenteeism that was not related to MRSs. For the entire group, this was 3.3 days per year; for the working group, it was 3.5 days, and for the studying group, it was 4.3 days. The mean percentage of absenteeism that was not related to MRSs was 3.5% in the working group and 3.7% in the studying group. Consequently, absenteeism due to MRSs in our cohort accounted for, on average, 22% of the total absenteeism in the working group and 24% in the studying group.

	Number (percentage)	Mean ± SE
All (n=32,748)		
Absenteeism	4,514 (13.8)	
≤0.5 day	538 (1.6%)	
1 day	2,259 (6.9%)	
2 days	1,171 (3.6%)	
3 days	349 (1.1%)	
>3 days	184 (0.6%)	
Total days of absenteeism per year	· · ·	1.3±5.9
Work (n=26,104)		
Absenteeism	2,926 (11.2%)	
≤0.5 day	374 (1.4%)	
1 day	1,476 (5.7%)	
2 days	757 (2.9%	
3 days	211 (0.8%)	
>3 days	98 (0.4%)	
Total days of absenteeism per year		0.9±3.9
Study (n=8,764)		
Absenteeism	1,715 (19.6%)	
≤0.5 day	234 (2.7%)	
1 day	921 (10.5%)	
2 days	423 (4.8%)	
3 days	92 (1.0%)	
>3 days	41 (0.5%)	
Total days of absenteeism per year		1.6±5.0

Table 2 | Reported absenteeism caused by menstruation-related symptoms. Women
were asked to report the amount of days on which they were absent due to
menstruation-related symptoms during the last 3 months. The total days of
absenteeism per year was calculated. The added numbers of women in the work and
study group exceed the total amount of participants, since 2,120 women reported to
both study and work more than 5 hours/week. SD = standard deviation

256 Presenteeism

Table 3 shows the numbers reported for presenteeism. Over 80% of all women reported presenteeism during their periods. The differences between the working group and the study group were not large in terms of prevalence and lost productivity. The mean number of lost productive days per year due to presenteeism was more than seven-fold greater than the mean number of lost productive days due to absenteeism.

2 3 4			Number (percentage)	Mean ± SD	Median
5		All (n=32,748)			
6 7		Presenteeism	26,438 (80.7%)		
8		Number of days in the past 3 months	, , , , , , , , , , , , , , , , ,	5.8±5.3	5.0
9		Percentage of productivity loss per day		33.0±24.8	30.0
10		Days/year of lost productivity		8.9±11.0	5.6
11		Work (n=26,104)			
12 13		Presenteeism	21,252 (81.4%)		
14 15		Number of days in the past 3 months		5.7±5.2	5.0
16		Percentage of productivity loss per day		31.7±24.7	30.0
17		Days/year of lost productivity		8.4±10.6	4.8
18		Study (n=8,764)			
19		Presenteeism	7,385 (84.3%)		
20		Number of days in the past 3 months		6.3±5.3	5.0
21		Percentage of productivity loss per day		36.8±24.2	40.0
22		Days/year of lost productivity		10.5±11.8	7.2
23	263				
24 25	264	Table 3 Reported presenteeism caused by	menetruation_rela	ted symptoms	Women
26					
27	265	were asked to report the amount of days on	•	•	
28	266	what extent. The total days of lost productivit	••••		added
29	267	numbers of women in the work and study gro	•		
30	268	participants, since 2,120 women reported to	both study and w	ork more than	5
31 32	269	hours/week. SD = standard deviation			
32 33 34	270				
35 36 37	271	Factors associated with absenteeism and pr	esenteeism		
38 39	272	Figure 2 shows the association between rep	oorted pain or disc	comfort scores	and both
40 41	273	absenteeism and presenteeism. As seen in	detail in table 4,	high VAS sc	ores were
42 43 44	274	significantly associated with higher levels	of absenteeism	and presentee	eism. The
45 46	275	strongest relationship was found for abdomin	nal pain scores th	at were 7 or h	igher on a
47 48	276	scale from 0 to ten. Odds ratios were 5.6 fo	or absenteeism (9	5% confidenc	e interval:
49 50	277	5.0 to 6.2) and 8.8 for presenteeism (95%	confidence interv	al: 8.1 to 9.5)	. Figure 3
51 52	278	shows the association between age and b	ooth presenteeisn	n and absente	eeism. As
53 54	279	shown in both figure 3 and table 4, we	e found that yo	unger womer	n showed
55 56	280	significantly higher rates of absenteeism	and presenteei	sm. A levon	orgestrel–
57 58 59	281	releasing intrauterine device is associated w	ith especially less	s presenteeism	
59 60		For peer review only - http://bmjopen.b	mj.com/site/about/g	uidelines.xhtml	14

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		OR (95% CI)	OR after correction (95% CI)	OR (95% CI)	OR aft correcti (95% C
Age<21 years ^a		3.7 (3.4-3.9)	3.3 (3.1-3.6)	1.4 (1.3-1.5)	1.3 (1.2-
Smoking ^b		1.3 (1.2-1.5)	1.3 (1.2-1.4)	1.5 (1.3-1.6)	1.4 (1.3-
Absenteeism not related to MRSs in the past six months ^c Level of education ^d		2.2 (2.1-2.4)	1.7 (1.6-1.9)	1.4 (1.3-1.5)	1.3 (1.2-
	Low	4.5 (4.1-4.9)	2.7 (2.4-3.0)	1.3 (1.2-1.4)	1.1 (1.0-1
	Medium	2.2 (2.1-2.4)	1.7 (1.5-1.8)	1.3 (1.2-1.4)	1.2 (1.1-
	High	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
Oral contraception ^e					、 、
	No	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	Yes	1.2 (1.1-1.3)	1.0 (0.9- 1.1)**	0.9 (0.9-1.0)	0.9 (0.8-0
LG-IUD ^e					
	No	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	Yes	0.7 (0.6-0.8)	0.9 (0.8-1.0)	0.5 (0.5-0.6)	0.5 (0.5-
Abdominal pain score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	5-6	2.6 (2.3-2.9)	2.2 (1.9-2.4)	5.2 (4.8-5.7)	5.3 (4.9-
	>7	7.0 (6.4-7.8)	5.6 (5.0-6.2)	8.7 (8.0-9.4)	8.8 (8.1-9
Headache pain score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	5-6	1.5 (1.3-1.6)	1.5 (1.4-1.6)	3.0 (2.7-3.3)	3.1 (2.8-
	>7	2.0 (1.8-2.1)	2.3 (2.1-2.5)	3.5 (3.2-3.9)	3.7 (3.4-4
Backache pain score ^e	0.4	1.0 (p/p)	1.0 (p/p)	1.0 (n/n)	1.0 (p/
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	5-6 >7	1.6 (1.5-1.7)	1.4 (1.3-1.5)	3.5 (3.2-3.9)	3.5 (3.2-3
Tirodnoco intercity	>(2.7 (2.5-2.9)	2.2 (2.1-2.4)	4.7 (4.2-5.2)	4.5 (4.0-
Tiredness intensity score ^e	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	0-4 5-6	1.8 (1.7-2.0)	1.8 (1.6-2.0)	3.3 (3.1-3.6)	3.3 (3.1-3
	>7	3.0 (2.8-3.2)	2.8 (2.6-3.1)	5.1 (4.7-5.6)	5.2 (4.7-
Psychological complaints intensity score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a
	5-6	1.6 (1.5-1.7)	1.5 (1.4-1.7)	2.7 (2.5-2.9)	2.6 (2,5-2
	>7	2.2 (2.0-2.4)	2.1 (2.0-2.3)	4.4 (4.0-4.7)	4.3 (4.0-4

Table 4 | Odd ratios (ORs) and 95% confidence intervals (95% CIs) for factors related to absenteeism and presenteeism. ORs >1 correlate with more absenteeism or presenteeism. ORs <1 correlate with less absenteeism or presenteeism. LG-IUD = Levonorgestrel-releasing intra uterine device ^aCorrection for smoking and absenteeism that was not related to menstruation-related symptoms (MRSs); ^bCorrection for age, absenteeism that was not related to MRSs, and level of education; ^cCorrection for age, smoking, and level of education; ^aCorrection for age, smoking, and absenteeism that was not related to MRSs; ^eCorrection for age, smoking, absenteeism that was not related to MRSs, and level of education. *p=0.26, **p=0.73 For all other ORs, p<0.05

294 Menstruation and suggested implications for schools and workplaces

From the respondent who had been calling in sick due to MRSs, 20.1% (n=908) told their employer or school menstrual symptoms were the reason, 46.4% (n=2092) only mentioned the presenting symptom. No reason was given by 27.7% (n=1250), while 5.8% (n=260) made up another reason. Women were asked to report suggestions on how work places and conditions could be changed in order for them to function better during their menstrual periods. There were 32,708 responses to this multiple-choice question, to which each woman could give more than one answer. The majority of women (67.7%, n=22,154) preferred more flexibility during their periods, such as the possibility of doing less physical work (32.1%, n=10,499), the ability to work from home (39.5%, n=12,917), more time for personal care (28.3%, n=9,241), or the ability to take a day off and make up for it later (11.5%, n=3,756). In addition, 32.9% wished they could take a complete day off without any consequences. 27.2% (n=8.890) did not wish for any changes in policy. Many women (79.7%, n=26,072) were open to discussing MRSs with their company doctor, and 56.7% (n=18,579) thought that doing so would draw more attention to MRS-related matters.

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Discussion

This survey-based study showed that menstruation-related absenteeism and, to a greater extent, presenteeism are widespread in the general female population. In our cohort, MRSs accounted for up to 24% of total absenteeism for women who were working and studying. The annual productivity loss due to presenteeism was seven-fold times more than the annual productivity loss due to absenteeism and women younger than 21 years experience the largest burden. Symptom severity scores showed significant and strong associations with both absenteeism and presenteeism. When women called in sick due to MRSs, only one in five stated openly that menstrual symptoms were the main reason. A majority of women prefers more flexibility during their periods when it comes to work or school.

There have been few studies on absenteeism and presenteeism related to MRSs in the general female population. To our knowledge, Tanaka's study ²⁵ is the only other published study on absenteeism and presenteeism due to MRSs in the general female population. In a cohort of 19,254 Japanese women, a total of 3,311 (17.2%) reported work productivity lost in the prior 3 months, mostly in the form of decreased efficiency (62.0%, n=2,052). Of these 2,052 subjects, the mean number of workdays lost due to decreased efficiency was 5.7 days in 3 months. After recalculation, this accounts for 2.4 days per year for the entire population. This is fewer days than the 8.9 days per year in our cohort. On the other hand, the numbers for absenteeism were more similar, with a mean of 1.0 day of absenteeism per year in the entire Japanese cohort compared to 1.2 days in our cohort. Cultural diversity might explain these differences, although it has been suggested in research on musculoskeletal symptoms that rates of absenteeism might be lower in Japan compared to European

countries and the United States. Consequently, presenteeism might therefore be a
 more representative variable.^{26 27}

More data are available regarding the impact of dysmenorrhoea on guality of life and absenteeism. De Sanctis et al. reviewed studies on dysmenorrhea in multiple countries, some of which included menstruation-related absenteeism data. They found that the prevalence of school absences in adolescents that was due to dysmenorrhea varied between 7.7% and 57.8%. Since the review included 41,140 women in 27 countries, and there was a high degree of heterogeneity in the outcome measures, no firm conclusions could be drawn. Hailemeskel et al. evaluated 440 female university students in Ethiopia.²⁸ Among students with dysmenorrhea, 66.8% reported a loss of concentration in class, and 56.3% reported class absences during the last month. In a questionnaire-based study of 706 Hispanic female adolescents, 38% reported missing school due to dysmenorrhea during the 3 months prior to the survey, and 59% reported a decrease in concentration in class due to dysmenorrhea.

Absenteeism and presenteeism due to endometriosis in other studies was greater than in our study, which was not surprising.^{9 14 30} However, some interesting parallels can be drawn to findings from a recent study by Soliman et al.¹⁴ They found that the average number of hours of presenteeism, 5.3 hours per week, was far greater than the number of hours of absenteeism, which was 1.1 hours per week. Furthermore, younger women had significantly higher levels of lost productivity than their older counterparts, and more severe symptoms were associated with more absenteeism and presenteeism.

357 Our finding that only 20.1% of women were open about their menstrual symptoms as 358 a reason for calling in sick may confirm the general idea that women tend not to

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speak openly about MRSs. Wong et al. found that in a cohort of schoolgirls in Malaysia, 76.1% considered dysmenorrhoea a normal part of the menstrual cycle.¹⁵ In the context of the findings noted above, our study also suggests there is a taboo for women in terms of discussing menstrual problems with their employers. The latter may therefore conclude that the impact of MRSs on their employees is negligible. Considering the fact that we also found that 68% of women wish that they had greater flexibility during their periods, either at school or at work, more openness about MRSs in the employment setting seems desirable. The reasons underlying this taboo are likely to have a historical basis; indeed, since ancient times, menstruation has been surrounded with mythical stories and has not been well understood. However, in recent years, the lay literature in developed countries has focused more attention on MRSs. ³¹⁻³³ The prevalence and the impact of MRSs on the general population and the number of women who are asking for a different approach all reflect the need to change the view of menstrual symptoms and the way they are addressed in society.

This study consisted of a large cohort, and it reached a large number of women within the age range that was aimed for. The questionnaire was developed in collaboration with patient representatives to make it understandable by and relevant to most women. The cohort appeared to be a representative sample of the general female population based on the number of working hours. ³³ When we compare our data with the national registries the total amount of absenteeism is found to be comparable, regardless of whether it was related to MRSs.^{34 35} It is difficult to compare our numbers on women with a diagnosis explaining their MRSs with numbers found in other studies. We found that only 9% of the participants had such a diagnosis, which seems about as expected or even somewhat low.^{3 36-38} In contrast, 45% of the women in the study reported consulting a physician for their MRSs. This

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number was relatively high compared to other studies in which, for example, the 385 386 percentage of women with dysmenorrhoea who sought medical advice was approximately 15%.¹⁵ ¹⁶ An important factor might be the Dutch health system in 387 which general practitioners are available free of charge. Women with a low level of 388 education were relatively underrepresented.³⁹ As our results show, especially 389 390 absenteeism related to MRSs is associated with a low level of education, and this might have biased our results. We expect women with lower educational levels to do 391 392 more physical jobs or jobs with less flexibility. Therefore our findings on work productivity loss might be underestimated. On the other hand, our finding could be 393 394 overestimated due to the possibility that women with more MRSs might be more likely to respond to a questionnaire, as it may seem more relevant to them. Thus, 395 these results must be interpreted with caution. Due to the way that the questionnaire 396 397 was distributed through social media, there may have been some selection bias. However, a recent review stated that Facebook is a useful recruitment tool for 398 healthcare research.⁴⁰ Although we did not use a validated guestionnaire, our most 399 important outcomes were based on questions derived from the PCQ, which itself is 400 based on validated questions and which is recommended by guidelines for cost 401 research.²⁴ Self-reported absenteeism generally shows a good correlation with 402 official records, although accuracy decreases with increasing recall period.⁴¹ This 403 404 might have initiated a recall bias in our cohort. It is unknown to what extend recall bias affects reports on presenteeism.⁴² In general, although results vary among 405 studies on premenstrual complaints, a prospective collection of data on symptoms is 406 advisable.^{43 44} Finally, these results may not be generalized internationally due to 407 408 variability in the regulation of social services in different countries, and this is also a limitation of our study. In The Netherlands, wages are paid during sick leave that has 409 duration of less than 1 year, but women in other countries may not have this benefit. 410

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Since we know that many factors influence menstrual symptoms, including biological, cultural, and environmental factors, these differences might well influence both absenteeism and presenteeism.^{6 14 45}

In conclusion, we have found that the impact of MRSs on work and school productivity is considerable and that presenteeism contributes significantly more to the matter than absenteeism. Future research should identify how women affected by MRSs could be helped best and how their productivity can be improved in order to reduce the societal impact regarding absenteeism and presenteeism. Ster

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568 Competing interest statement

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CZ.

575 Contributorship statement

576 TN, BD and JA conceived the study. MS wrote the first and successive drafts of the 577 manuscript. MS, TN and EA modelled and analysed the data. TN, EA, JM, BD and 578 JA contributed to study conception and design. MS and TN collected the data. All 579 authors revised the manuscript for important intellectual content. MS and TN had full 580 access to the data and take responsibility for the integrity of the data and the 581 accuracy of the data analysis. TN is the guarantor.

583 Data sharing statement

584 No additional data from this study are available from a repository. Data are available 585 on request from the corresponding author.

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The relationship between pain and intensity scores, related to absenteeism and

The relationship between age and average absenteeism and presenteeism.

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Legends to figures

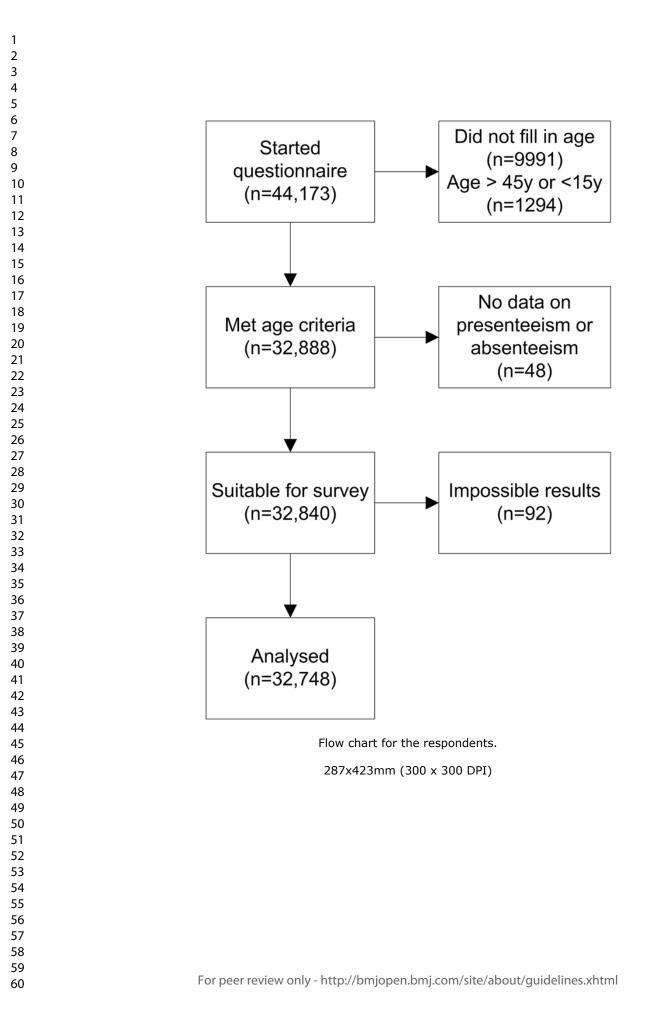
Flow chart for the respondents.

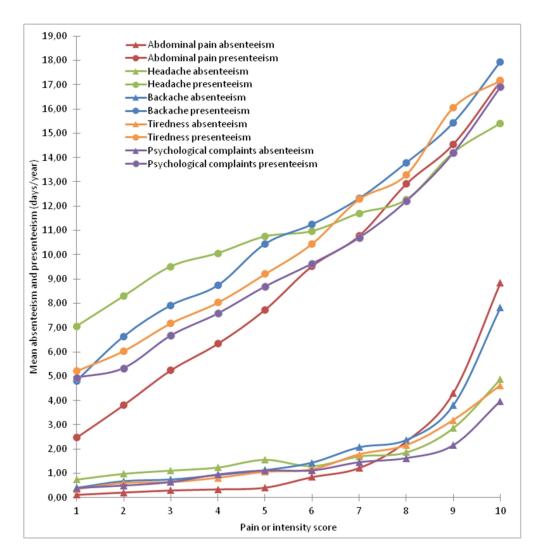
presenteeism, in lost days per year.

Figure 1

Figure 2

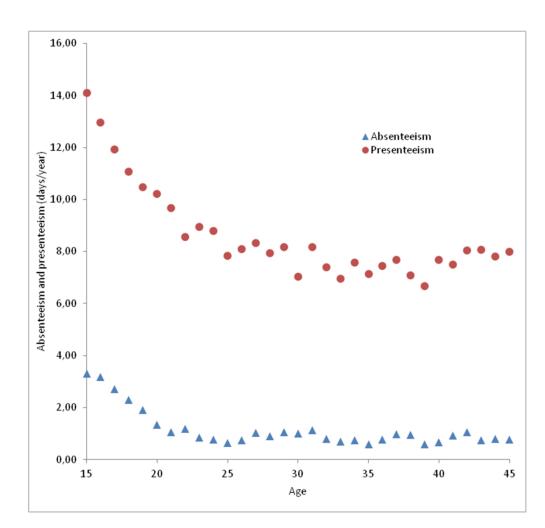
Figure 3





The relationship between pain and intensity scores, related to absenteeism and presenteeism, in lost days per year.

202x209mm (300 x 300 DPI)



The relationship between age and average absenteeism and presenteeism.

206x199mm (300 x 300 DPI)

Appendix

Summary of the questionnaire

Part 1: Questions about basic characteristics

- Age, marital status, nationality, level of education
- Work, voluntary work and study: hours per week, sector,
- Smoking, stress
- Menstrual cycle: length, number of days with blood loss, number of times needing to change pad or tampon
- Medical and obstetric history
- Contraception use

Part 2: General questions about menstruation-related symptoms

- Symptoms: abdominal pain, heavy menstrual blood loss, headache, fatigue,
 backache, nausea and vomiting, tender breasts, problems with stool, psychological
 complaints
- Symptoms: number of days the symptom was present; number of days that the symptom influenced daily functioning; the magnitude of the symptom's influence on daily functioning; pain or intensity score (not for heavy menstrual bleeding, nausea and vomiting); continuation of daily activities despite symptoms; and the extent to which women forced themselves to continue their daily activities.
- Usage of over-the-counter drugs, narcotics, painkillers, alternative medicine

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3	Part 3: Questions related to absenteeism and presenteeism
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6	 Number of days of absenteeism due to menstruation-related symptoms
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8	 Number of days of presenteeism due to menstruation-related symptoms
9	· · · · · / · · · · · · · · · · · · · ·
10	- Mean amount of productivity loss due to menstruation-related symptoms during
11	- Mean amount of productivity loss due to mensil dation-related symptoms during
12	
13	these days
14	
15	- Number of days of absenteeism not related to menstruation-related symptoms
16	
17	- Reasons given for absenteeism due to menstruation-related symptoms
18	neusons given of ussenteelsin due to mensi dution related symptoms
19	Currentians for the second the supplies of the selection of the time hotten.
20	 Suggestions for changes at the workplace or schools in order to function better
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22	during menstrual periods
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STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of cross-sectional studies

Section/Topic	ltem #	Recommendation	Reported on page #	
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	2	
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2	
Introduction				
Background/rationale 2 Explain the scientific background and rationale for the investigation being reported				
Objectives	3	State specific objectives, including any prespecified hypotheses	5	
Methods				
Study design	4	Present key elements of study design early in the paper	5	
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	5-7	
Participants 6 (<i>a</i>) Give the eligibility criteria, and the sources and methods of selection of participants		6		
Variables	7 Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable		8, appendix	
Data sources/ 8* For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe measurement comparability of assessment methods if there is more than one group			7-8	
Bias	9	Describe any efforts to address potential sources of bias	7-9	
Study size	10	Explain how the study size was arrived at	7	
Quantitative variables 11 Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why		7-9		
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	7-8	
		(b) Describe any methods used to examine subgroups and interactions	7-8	
		(c) Explain how missing data were addressed	7,9	
		(d) If applicable, describe analytical methods taking account of sampling strategy	n/a	
		(e) Describe any sensitivity analyses	n/a	
Results				

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Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility,	9
		confirmed eligible, included in the study, completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	9/Figure 1
		(c) Consider use of a flow diagram	Figure 1
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	9/Table 1
		(b) Indicate number of participants with missing data for each variable of interest	Not reported, can be reported if desirable.
Outcome data	15*	Report numbers of outcome events or summary measures	9-11/Table 2,3
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence	Table 4
		interval). Make clear which confounders were adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	Table 1,4
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	n/a
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	n/a
Discussion		6	
Key results	18	Summarise key results with reference to study objectives	11-12
Limitations 19 Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias		14-15	
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	12-15
Generalisability	21	Discuss the generalisability (external validity) of the study results	14-15
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	n/a

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

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Productivity loss due to menstruation-related symptoms: a nationwide cross-sectional survey among 32,748 women

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Keywords:	Menstruation, Community gynaecology < GYNAECOLOGY, Menstrual cycle, Presenteeism, Absenteeism



1 2		
2 3 4	1	Productivity loss due to menstruation-related symptoms: a nationwide cross-
5 6 7	2	sectional survey among 32,748 women
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19 20 21	28	
22 23 24 25	29	Word count: 4,056
26 27 28	30	
29 30	31	Key Words: Absenteeism, Presenteeism, Menstrual cycle, Menstruation, Community
31 32 33	32	Gynaecology
34 35 36		
37 38		
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2 3 4 5	33	Abstract
6 7	34	Objective
8 9	35	To evaluate age-dependent productivity loss caused by menstruation-related
10 11 12	36	symptoms, measured in absenteeism (time away from work or school) and
13 14	37	presenteeism (productivity loss while present at work/school).
15 16 17 18	38	
19 20	39	Methods
21 22 23	40	Design/Setting: Internet-based, cross-sectional survey conducted in The Netherlands
23 24 25	41	from July to October 2017.
26 27 28 29	42	Participants: 32,748 women aged 15 to 45 years, recruited through social-media.
30 31	43	Outcome measures: Lost productivity in days, divided into absenteeism and
32 33 34	44	presenteeism; impact of menstrual symptoms; reasons women give to their employer
35 36	45	or school when calling in sick; and women's preferences regarding the implications of
37 38	46	menstruation related symptoms for schools and workplaces.
39 40 41	47	Results
42 43 44	48	A total of 13.8% of all women reported absenteeism during their menstrual periods
45 46	49	with 3.4% reporting absenteeism every or almost every menstrual cycle. The mean
47 48	50	absenteeism related to a woman's period was 1.2 days per year. A total of 80.7% of
49 50	51	the respondents reported presenteeism and decreased productivity a mean of 23.2
51 52 53	52	days per year. An average productivity loss of 33% resulted in a mean of 8.9 days of
54 55	53	total lost productivity per year. Women under 21 were more likely to report
56 57	54	absenteeism due to menstruation-related symptoms (OR 3.3, 95% confidence
58 59 60	55	interval 3.1 to 3.6). When women called in sick due to their periods, only 20.1% told
	56	their employer or school mentor that their absence was due to menstrual complaints.

Notably, 67.7% of the participants wished they had greater flexibility in their tasks and
working hours at work or school during their periods.

60 <u>Conclusions</u>

Menstruation-related symptoms cause a great deal of lost productivity, and presenteeism is a bigger contributor to this than absenteeism. There is an urgent need for more focus on the impact of these symptoms, especially in adolescents, for discussions of treatment options with women of all ages, and, ideally, more flexibility for women who work or go to school.

67 Strengths and limitations of this study

- This is the largest cohort study to analyse the impact of menstruation-related symptoms on work and school productivity.
- The survey was performed among the general female population and is consequently not per se related to one specific gynaecological condition.
 - Due to the way of recruitment of participants, there may have been some degree of selection bias.
 - The generalisability of the study may be limited to employment and school systems comparable to the Dutch.

77 Background

Menstruation-related symptoms (MRSs) are diverse and widespread among women. Symptoms include dysmenorrhea, heavy menstrual bleeding and premenstrual mood disturbances with reported prevalence of 45% to 90%, 14% to 25%, and 20% to 29% respectively.¹⁻³ Studies show that women with MRSs have lower scores on several domains of guality of life such as general health and physical, mental, social, and occupational functioning during their periods.^{1 4-7} Furthermore, these symptoms may create considerable financial burdens on patients and their families as well as on society.^{5 6 8-12} Such financial burdens are related to the costs of visits to the doctor, over-the-counter drugs, and medical or surgical treatment. However, costs related to productivity loss could be the largest cost driver. Productivity costs are defined as costs associated with paid and unpaid production loss and the replacement of productive people due to illness or disability.¹³ Productivity costs can be divided into costs related to absenteeism and costs related to presenteeism. Absenteeism represents the total amount of time off work or away from school, and presenteeism represents the loss of productivity while present at a job or school.

Although the literature is scarce and the results are variable, studies on specific patient groups generally show that MRSs can cause absenteeism.¹⁴⁻¹⁶ Research on the association between MRSs and presenteeism is even more limited. It has been suggested that research into possible impairments in quality of life caused by menstrual symptoms should not focus on single symptoms but rather on a complex of symptoms that vary widely but that are all related to the menstrual cycle. This complex includes both standard symptoms, like heavy menstrual bleeding and abdominal cramps, and also less common symptoms, like nausea and cold sweats. 17 18

Taking all symptoms into account, it seems likely that the real impact of MRSs is underestimated in the general population. Despite being almost two decades into the 21st century, discussions about MRSs may still be rather taboo. This survey-based exploratory study aimed to quantify the burden of MRSs in the general female population, with burden defined as the number of lost days at work or school due to absenteeism and presenteeism. Furthermore, it was aimed to study the impact of specific symptoms on absenteeism and presenteeism.

110 Methods

This cross-sectional study consisted of an online survey that was distributed from July 12 to October 11, 2017. Approval for this study was obtained from the local medical ethics committee "Commissie Mensgebonden Onderzoek (CMO)" under number file number 2017-3387 on July 12th 2017. All data were anonymously collected and stored under the privacy rules of the Radboud University Medical Center. Patients gave informed consent when they initiated the questionnaire.

118 Patient and Public Involvement

A group of women, among which were several members of the Dutch Patient Endometriosis Foundation, women with a linguistic education and women with a medical origin, was involved in the conduct of this study at several stages; i.e. in the development and dissemination of the questionnaire and in the analysis and interpretation of the results. One of the authors of this manuscript, BD, is the chair of the Dutch Patient Endometriosis Foundation. Additional contributions are noted in the Acknowledgements section.

1 2		
3 4	126	
5 6 7 8	127	Questionnaire development
9 10	128	The questionnaire had several parts, and Appendix 1 provides details about the
11 12	129	questions. Part 1 consisted of questions about each woman's basic characteristics.
13 14 15	130	Part 2 had questions about menstrual symptoms, and part 3 had questions related to
16 17	131	absenteeism and presenteeism. Adaptive questioning was used with a maximum of 6
18 19	132	questions per page. Participants were asked in a lay manner how long their
20 21 22	133	menstrual cycle was and what the exact meaning of a menstrual cycle was. The
23 24	134	duration of the cycle was divided in 5 categories (25 days or less, 26-30 days, 31-35
25 26	135	days, 36-40 days and 41 days or more). Furthermore participants could indicate if
27 28 29	136	they had an irregular cycle, meaning more than 10 days difference per cycle, if they
30 31	137	were amenorrheic due to the use of an Intra-Uterine Device (IUD) or the continuous
32 33	138	use of oral contraceptives, or the option "I do not know". Additional questions about
34 35	139	absenteeism and presenteeism were included that were based on the Productivity
36 37 38	140	Cost Questionnaire from the Institute for Medical Technology Assessment (iMTA-
39 40	141	PCQ). ¹⁹ We modified the iMTA-PCQ-recommended recall period of four weeks to
41 42	142	three months so that it was in line with the relevant time period for this study and so
43 44 45	143	we could include multiple menstrual periods. We assumed the amount of
46 47	144	presenteeism to be larger than the amount of absenteeism. Therefore, the recall
48 49	145	period for absenteeism was extended to six months to maintain accuracy. Visual
50 51 52	146	analogue scales (VAS) were used to quantify the amount of pain, or the intensity of
52 53 54	147	the symptom, and the impairment due to pain or the other symptom. Presenteeism
55 56	148	was measured by asking women to what extent they were able to be as productive
57 58	149	as possible compared to a day without MRSs. This was scored on a scale from 0 to
59 60	150	10, with 0 being totally unproductive and 10 fully productive. In separate questions,

participants were asked to quantify their absenteeism that was related to MRSs and
absenteeism for any other reason than MRSs. For the latter, we did not specifically
ask the underlying reason.

155 Target population and recruitment

The study population comprised of women between 15 and 45 years old. The upper age limit was chosen to avoid interference from menopausal symptoms; the lower to have a time margin after the average menarche age, since it is known that the first periods are irregular and often accompanied with discomfort and uncertainty. A large number of women were approached with the aim of obtaining a cohort that was representative of the general female population in terms of level of education, medical history, and/or gynaecological diagnosis. Women were invited to complete a survey using an online questionnaire tool²⁰ through a campaign on social media platforms Facebook and Twitter. Patient organisations, colleagues, and visitors of the Facebook page of one of the authors (TN) were asked to share the link to the questionnaire in order to reach the widest possible audience.

On July 12, 2017, a link to the questionnaire was posted on Facebook and Twitter through the account of one of the authors (TN). In the post both women with and without MRSs were encouraged to fill in the questionnaire. Within 24 hours of the first posting on social media, over 6,000 respondents had filled in the questionnaire, and by July 18, there were 15,000 respondents, which was announced by a re-post of the link to the guestionnaire. A third post was made on Facebook and Twitter on September 16, 2017, to reach women who may have been on holiday when the first posts were created.

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2 3 4	175	
5 6 7	176	Data analysis
8 9 10	177	The outcome measures were presented in a descriptive way; we used valid
11 12	178	percentages in case of missing values where necessary. We distinguished between
13 14 15	179	women who were mainly working or mainly studying. Therefore, we present these
16 17	180	data for two groups i.e. for women who worked more than 5 hours per week
18 19	181	("working group") and for women who studied more than 5 hours per week ("studying
20 21 22	182	group").
23 24 25	183	We used binary logistic regression to calculate odds ratios. Absenteeism and
26 27	184	presenteeism were used as dependant variables. As independent variables we used
28 29	185	the following parameters: women younger than 21 versus women aged 21 and older,
30 31 32	186	smoking yes or no, reports of absenteeism not related to MRSs, educational level,
33 34	187	the use of oral contraception and the use of an levonorgestrel-releasing intra uterine
35 36	188	device. All independent variables were used in an univariate as well as a multivariate
37 38 39	189	analysis. We also studied the association between pain scores and both
39 40 41	190	absenteeism and presenteeism, given that the literature shows that pain scores of 0
42 43	191	to 4, 5 to 6, and 7 or higher have a different impact on activity, mood, and sleep. ^{21 22}
44 45 46	192	Analyses were performed using IBM SPSS Statistics version 22.00.
47 48 49	193	
50 51 52	194	Assumptions and transformation of the original data
53 54 55	195	To present data on level of education in an international format we had to transform
56 57	196	the original data, which was based on the Dutch school system.23 The categorical
58 59 60	197	data of participants' length of menstrual cycle were transformed into averages.

With regard to the evaluation of absenteeism and presenteeism, "the guideline for economic evaluations in healthcare in the Netherlands" was used.²⁴ A work-day accounts for 8 hours. For most sectors in the Netherlands, a full-time work-week is 36 hours. The maximum amount of working hours per year was set at 1558 when they were working full time. We asked women to report their absenteeism due to MRSs per cycle and used a recall period of six months.

To calculate the percentages for absenteeism, one day of absenteeism accounted for 8 hours of lost productivity. When a woman reported to study or work more than 40 hours per week, we transformed these hours to 40 for reasons of clarity in the calculations and comparability with the data of the Dutch Central Bureau of Statistics, CBS. We made a few other transformations for categorical data. For absenteeism related to MRSs, the category "more than three days per cycle" was considered to be 4 days per cycle. For absenteeism that was not related to MRSs, the category "more than ten days in the past six months" was considered to be 11 days in the past six months.

To present yearly data, we multiplied some of these data based on the original recall period. The number of days for absenteeism related to MRSs was based on days per cycle, which were therefore multiplied by 12.7 based on the reported average menstrual cycle of 28.8 days, see table 1. These values were then multiplied by one if the woman reported that she called in sick "every period", 0.75 if she reported "almost every period", 0.5 if she reported "half of all periods", and 0.25 if she reported calling in sick "only once in a while". Values for absenteeism that was not related to MRS were based on a recall period of 6 months and were therefore multiplied by two in order to obtain the number of days per year. The values for presenteeism were based on a recall period of three months and were therefore multiplied by four.

1 2 3	223					
4 5	223					
5 6 7 8	224	Results				
9 10	225	A total of 44,173 women initiated the questionnaire. W	e excluded parti	cipants who		
11 12	226	did not report a date of birth or whose age did not fulfil th	e inclusion criter	ia (figure 1).		
13 14 15	227	There were no duplicates of IP addresses. Women whether the second s	no did not answe	er questions		
15 16 17	228	related to absenteeism and presenteeism were exclud	ed. Furthermore,	cases with		
18 19	229	impossible results (e.g. 10,000,000 days of presentee	ism in three mo	nths or 140		
20 21	230	changes of sanitary pads a day) were excluded. This	resulted in a tota	al of 32,748		
22 23 24	231	women in the final analysis.				
25 26 27	232	Table 1 summarizes the basic characteristics of the	participants. We	found that		
28 29	233	45.4% (n=14,839) had visited a doctor for menstrual complaints in the past, with a				
30 31 32	234	total of 3017 (9.2%) women reporting a diagnosis of a menstrual disorder, such as				
32 33 34	235	endometriosis or fibroids.				
35 36 37	236	The mean age of women in the working group was higher than the mean ager of				
38 39	237	women in the studying group. The mean number of working hours per week in the				
40 41 42	238	working group was 27.0 (SD 11.4), and the mean number of study hours in the				
43 44	239	studying group was 27.4 (SD 12.1). A total of 7,335 women (22.4%) reported both				
45 46	240	working and studying more than five hours per week. In this group, 3,001 women				
47 48 49 50 51 52 53 54 55 56 57	241	were working more than 16 hours a week, and 5,284 women in the study group were				
	242	studying more than 16 hours a week.				
		Number (percentaç		Median		
		Age, years	28.6±8.6	28		
58 59		15-19 6,141 (18.8	%)			
60		20-24 6,118 (18.7	'%)			

$\begin{array}{c}1\\2\\3\\4\\5\\6\\7\\8\\9\\10\\11\\2\\13\\14\\15\\16\\17\\8\\19\\20\\22\\23\\24\\25\\27\\28\\29\\30\\1\\23\\34\\35\\36\\37\\89\\0\\41\\42\\44\\45\\6\\7\\8\end{array}$	
43 44 45 46 47	
55 56 57 58 59 60	

25-29	5,825 (17.8%)		
30-34	5,483 (16.7%)		
35-40	4,687 (14.3%)		
41-45	4,494 (13.7%)		
Level of education			
Low	4,020 (12.3%)		
Medium	12,335 (37.9%)		
High	16,229 (49.8%)		
Hours/week			
Paid work		21.7±14.7	24
Study		7.4±13.6	0
Voluntary work		0.8±3.1	0
Menstrual cycle			
Regular cycle	25,717 (78.5%)		
Duration		28.8±3.0	28
Amenorrhoea due to LG-IUD/OC	3,675 (11.2%)		
Irregular, variation >10 days per cycle	2,495 (7.6%)		
Do not know	861 (2.6%)		
Days with blood loss per cycle	0	5.4±1.6	5
Visited a doctor for MRSs			
Νο	17,873 (54.6%)		
Yes, general practitioner	10,141 (31.0%)		
Yes, gynaecologist	4,698 (14.4%)		
Diagnosis for MRSs*			
Νο	29,731 (90.8%)		
Yes	3,017 (9.2%)		
Endometriosis	1,120 (3.4%)		
PCOS	588 (1.8%)		
Adenomyosis	103 (0.3%)		

	Fibroids	275 (0.8%)	
-	Other	1,901 (5.8%)	
-	Contraception*		
-	Hormonal contraception	11,993 (36.6%)	
-	OC	8,650 (26.4%)	
-	LG-IUD	2,752 (8.4%)	
_	Other hormonal: injection, transdermal etc.	882 (2.7%)	
-	No hormonal contraception	20,755 (63.4%)	
-	Cu-IUD	771 (2.4%)	
-	Female sterilisation	423 (1.3%)	
_	No female contraception	19,639 (60.0%)	
-	Nulliparous	21,585 (66.0%)	
-	Paid work >5 hr a week	26,104 (79.7%)	
-	Age	0,	29.7±8.3
_	Hours of paid work/week	2	27.0±11.4
_	Hours spent on study/week	(C)	7.5±13.4
-	>40 hours of paid work/week	1,047 (3.2%)	
-	Study >5 hr a week	8,764 (26.8%)	
-	Age		22.0±6.2
_	Hours spent on study/week		27.4±12.1
-	Hours of paid work/week		15.5±11.3
-	>40 hours spent on study/week	322 (1.0%)	

Table 1 | Basic characteristics of study participants (n=32,748) Mean duration of cycle based on women with a regular cycle. SD = standard deviation, MRSs = menstruation-related symptoms, PCOS = polycystic ovary syndrome, OC = oral contraceptive, LG-IUD = levonorgestrel-releasing intrauterine device, Cu-IUD = copper intrauterine device. *More than one answer possible.

250 Absenteeism

Table 2 shows the results on absenteeism due to MRSs. Although 13.8% of the women (n=4,514) reported absenteeism due to MRSs, only 1,108 women (3.4%) reported absenteeism every cycle or almost every cycle. The percentage of absenteeism in every cycle or almost every cycle was 2.4% in the working group and 4.5% in the studying group. The mean absenteeism due to MRSs was 0.9 days per year for the working group and 1.6 day per year for the study group.

We also calculated the mean total absenteeism that was not related to MRSs. For the entire group, this was 3.3 days per year; for the working group, it was 3.5 days, and for the studying group, it was 4.3 days. The mean percentage of absenteeism that was not related to MRSs was 3.5% in the working group and 3.7% in the studying group. Consequently, absenteeism due to MRSs in our cohort accounted for, on average, 22% of the total absenteeism in the working group and 24% in the studying group.

59

60

	Number (percentage)	Mean ± SE
All (n=32,748)		
Absenteeism	4,514 (13.8)	
≤0.5 day	538 (1.6%)	
1 day	2,259 (6.9%)	
2 days	1,171 (3.6%)	
3 days	349 (1.1%)	
>3 days	184 (0.6%)	
Total days of absenteeism per year	•	1.3±5.9
Work (n=26,104)		
Absenteeism	2,926 (11.2%)	
≤0.5 day	374 (1.4%)	
1 day	1,476 (5.7%)	
2 days	757 (2.9%	
3 days	211 (0.8%)	
>3 days	98 (0.4%)	
Total days of absenteeism per year	2	0.9±3.9
Study (n=8,764)	0	
Absenteeism	1,715 (19.6%)	
≤0.5 day	234 (2.7%)	
1 day	921 (10.5%)	
2 days	423 (4.8%)	
3 days	92 (1.0%)	
>3 days	41 (0.5%)	
Total days of absenteeism per year	•	1.6±5.0

Table 2 | Reported absenteeism caused by menstruation-related symptoms. Women
were asked to report the average amount of days on which they were absent due to
menstruation-related symptoms per cycle. The total days of absenteeism per year

3 4 5 6	269 270 271	was calculated. The added numbers of women in the work and study group exceed the total amount of participants, since 2,120 women reported to both study and work more than 5 hours/week. SD = standard deviation						
7 8	272							
9 10 11	273	Presenteeism						
12 13 14	274	Table 3 shows the numbers reported for	presenteeism. O	ver 80% of a	ll women			
15 16	275	reported presenteeism during their periods	. The difference	s between the	e working			
17 18 19	276	group and the study group were not large in t	terms of prevalen	ice and lost pro	oductivity.			
20 21	277	The mean number of lost productive days p	per year due to p	presenteeism v	was more			
22 23	278	than seven-fold greater than the mean n	umber of lost p	roductive day	s due to			
24 25 26	279	absenteeism.						
20 27 28 29 30			Number (percentage)	Mean ± SD	Median			
31 32		All (n=32,748)						
33 34 35		Presenteeism	26,438 (80.7%)					
36 37		Number of days in the past 3 months	4	5.8±5.3	5.0			
38 39		Percentage of productivity loss per day		33.0±24.8	30.0			
40 41		Days/year of lost productivity		8.9±11.0	5.6			
42 43		Work (n=26,104)						
44 45 46 47		Presenteeism	21,252 ~ (81.4%)					
47 48 49		Number of days in the past 3 months		5.7±5.2	5.0			
50 51		Percentage of productivity loss per day		31.7±24.7	30.0			
52 53		Days/year of lost productivity		8.4±10.6	4.8			
54 55		Study (n=8,764)						
56 57		Presenteeism	7,385 (84.3%)					
58 59		Number of days in the past 3 months		6.3±5.3	5.0			
60								

1 2

54 55

56

57

58

59 60 **Smoking**^b

Absenteeism not

related to MRSs in

the past six months^c

Level of education^d

	Days/year of lost	productivity		10.5±11.8	7.2
280					
281	Table 3 Reported	presenteeism caused	by menstruation	-related symptom	s. Women
282	were asked to repo	rt the amount of days of	on which they we	ere less productiv	e and to
283	what extent. The to	tal days of lost produc	tivity per year wa	as calculated. The	added
284	numbers of women	in the work and study	group exceed th	e total amount of	
285	participants, since 2	2,120 women reported	to both study an	id work more thar	า 5
286	hours/week. SD = s	standard deviation			
287					
288	Factors associated	with absenteeism and	l presenteeism		
289	Figure 2 shows the	association between	reported pain or	discomfort score	s and both
290	absenteeism and p	presenteeism. As seen	n in detail in tabl	e 4, high VAS so	cores were
291	significantly associ	ated with higher leve	els of absenteeis	sm and presente	eism. The
 strongest relationship was found for abdominal pain scores that were 7 or hi scale from 0 to ten. Odds ratios were 5.6 for absenteeism (95% confidence) 			nigher on a		
			n (95% confidend	ce interval:	
294	5.0 to 6.2) and 8.8	for presenteeism (95	% confidence in	terval: 8.1 to 9.5). Figure 3
295	shows the associa	tion between age and	d both presente	eism and absent	eeism. As
296	shown in both fig	jure 3 and table 4,	we found that	younger wome	n showed
297	significantly higher	rates of absenteeis	sm and preser	teeism. A levor	orgestrel-
298	releasing intrauterir	ne device is associated	d with especially	less presenteeisr	n.
		Absenteeis	sm	Presenteeism	
		OR (95% (CI) OR after correction (95% CI)	OR (95% CI)	OR after correction (95% CI)
					(

1.3 (1.2-1.5)

2.2 (2.1-2.4)

1.3 (1.2-1.4)

1.7 (1.6-1.9)

1.5 (1.3-1.6)

1.4 (1.3-1.5)

1.4 (1.3-1.6)

1.3 (1.2-1.4)

	Low	4.5 (4.1-4.9)	2.7 (2.4-3.0)	1.3 (1.2-1.4)	1.1 (1.0-1.2
	Medium	2.2 (2.1-2.4)	1.7 (1.5-1.8)	1.3 (1.2-1.4)	1.2 (1.1-1.2
	High	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
Oral contraception ^e					
	No	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	Yes	1.2 (1.1-1.3)	1.0 (0.9- 1.1)**	0.9 (0.9-1.0)	0.9 (0.8-0.9
LG-IUD ^e					
	No	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	Yes	0.7 (0.6-0.8)	0.9 (0.8-1.0)	0.5 (0.5-0.6)	0.5 (0.5-0.6
Abdominal pain score ^e	~				
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	2.6 (2.3-2.9)	2.2 (1.9-2.4)	5.2 (4.8-5.7)	5.3 (4.9-5.7
	>7	7.0 (6.4-7.8)	5.6 (5.0-6.2)	8.7 (8.0-9.4)	8.8 (8.1-9.5
Headache pain score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	1.5 (1.3-1.6)	1.5 (1.4-1.6)	3.0 (2.7-3.3)	3.1 (2.8-3.4
	>7	2.0 (1.8-2.1)	2.3 (2.1-2.5)	3.5 (3.2-3.9)	3.7 (3.4-4.1
Backache pain score ^e				1	
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	1.6 (1.5-1.7)	1.4 (1.3-1.5)	3.5 (3.2-3.9)	3.5 (3.2-3.8
	>7	2.7 (2.5-2.9)	2.2 (2.1-2.4)	4.7 (4.2-5.2)	4.5 (4.0-5.0
Tiredness intensity score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	1.8 (1.7-2.0)	1.8 (1.6-2.0)	3.3 (3.1-3.6)	3.3 (3.1-3.6
	>7	3.0 (2.8-3.2)	2.8 (2.6-3.1)	5.1 (4.7-5.6)	5.2 (4.7-5.7
Psychological complaints intensity					

S	core ^e						
		0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	
		5-6	1.6 (1.5-1.7)	1.5 (1.4-1.7)	2.7 (2.5-2.9)	2.6 (2,5-2.9)	
		>7	2.2 (2.0-2.4)	2.1 (2.0-2.3)	4.4 (4.0-4.7)	4.3 (4.0-4.7)	
299							
300	Table 4 Odd ratio	os (ORs) an	d 95% confider	ice intervals (9	5% CIs) for fac	tors	
301	related to absente	eism and pr	esenteeism. O	Rs >1 correlate	e with a higher p	orevalence	
302	of absenteeism or	presenteeis	sm. ORs <1 cor	relate with a lo	ower prevalence	e of	
303	absenteeism or pr	esenteeism					
304	LG-IUD = Levonor	gestrel-rele	asing intra uter	ine device			
305	^a Correction for sm	oking and a	bsenteeism tha	at was not relat	ed to menstrua	tion-	
306	related symptoms (MRSs); ^b Correction for age, absenteeism that was not related to						
307	MRSs, and level o	f education;	; ^c Correction for	⁻ age, smoking	, and level of e	ducation;	
 ^dCorrection for age, smoking, and absenteeism that was not related to MRSs; ^eCorrection for age, smoking, absenteeism that was not related to MRSs, and let 			Ss;				
			and level of				
310	education. *p=0.26	6, **p=0,73	For all other O	Rs, p<0.05			
311							
312	Menstruation and	suggested i	mplications for	schools and w	orkplaces		
13	From the respond	ent who had	d been calling i	n sick due to I	MRSs, 20.1% (I	n=908) told	
14	their employer or	school men	strual symptom	s were the rea	son, 46.4% (n=	=2092) only	
315	mentioned the pre	esenting syn	nptom. No reas	son was given	by 27.7% (n=1	250), while	
316	5.8% (n=260) ma	ide up ano	ther reason. C	comparing wo	men aged belo	ow 21 with	
317	women aged 21 a	and above,	we found that	younger wom	ien were less d	open about	
318	their MRSs being	the reason	for calling in si	ck (12.0%) ve	rsus women old	der than 21	
319	(27.0%). Women	were aske	ed to report s	suggestions o	n how work p	places and	
320	conditions could	be change	d in order fo	r them to fur	nction better d	luring their	
321	menstrual periods	. There wer	e 32,708 respo	onses to this m	nultiple-choice c	question, to	
322	which each wom	an could g	ive more than	one answer.	The majority	of women	

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(67.7%, n=22,154) preferred more flexibility during their periods, such as the possibility of doing less physical work (32.1%, n=10,499), the ability to work from home (39.5%, n=12,917), more time for personal care (28.3%, n=9,241), or the ability to take a day off and make up for it later (11.5%, n=3,756). In addition, 32.9% wished they could take a complete day off without any consequences. 27.2% (n=8,890) did not wish for any changes in policy. Many women (79.7%, n=26.072) were open to c, .ttention to MF. discussing MRSs with their company doctor, and 56.7% (n=18,579) thought that doing so would draw more attention to MRS-related matters.

331 Discussion

This survey-based study showed that menstruation-related absenteeism and, to a greater extent, presenteeism are widespread in the general female population. In our cohort, MRSs accounted for up to 24% of total absenteeism for women who were working and studying. The annual productivity loss due to presenteeism was seven-fold times more than the annual productivity loss due to absenteeism and women younger than 21 years experience the largest burden. Symptom severity scores showed significant and strong associations with both absenteeism and presenteeism. When women called in sick due to MRSs, only one in five stated openly that menstrual symptoms were the main reason. A majority of women prefers more flexibility during their periods when it comes to work or school.

There have been few studies on absenteeism and presenteeism related to MRSs in the general female population. To our knowledge, Tanaka's study ²⁵ is the only other published study on absenteeism and presenteeism due to MRSs in the general female population. In a cohort of 19,254 Japanese women, a total of 3,311 (17.2%) reported work productivity lost in the prior 3 months, mostly in the form of decreased efficiency (62.0%, n=2,052). Of these 2,052 subjects, the mean number of workdays lost due to decreased efficiency was 5.7 days in 3 months. After recalculation, this accounts for 2.4 days per year for the entire population. This is fewer days than the 8.9 days per year in our cohort. On the other hand, the numbers for absenteeism were more similar, with a mean of 1.0 day of absenteeism per year in the entire Japanese cohort compared to 1.2 days in our cohort. Differences in regulations of social services, a difference in attitude towards sick leave and a different method of data collection might explain these differences. It has been suggested in research on musculoskeletal symptoms that rates of absenteeism might be lower in Japan

compared to European countries and the United States. Consequently, presenteeism
 might therefore be a more representative variable.^{26 27}

More data are available regarding the impact of dysmenorrhoea on guality of life and absenteeism. De Sanctis et al. reviewed studies on dysmenorrhea in multiple countries, some of which included menstruation-related absenteeism data. They found that the prevalence of school absences in adolescents that was due to dysmenorrhea varied between 7.7% and 57.8%. Since the review included 41,140 women in 27 countries, and there was a high degree of heterogeneity in the outcome measures, no firm conclusions could be drawn. Hailemeskel et al. evaluated 440 female university students in Ethiopia.²⁸ Among students with dysmenorrhea, 66.8% reported a loss of concentration in class, and 56.3% reported class absences during the last month. In a questionnaire-based study of 706 Hispanic female adolescents, 38% reported missing school due to dysmenorrhea during the 3 months prior to the survey, and 59% reported a decrease in concentration in class due to dysmenorrhea.

Absenteeism and presenteeism due to endometriosis in other studies was greater than in our study, which was not surprising.^{9 14 30} However, some interesting parallels can be drawn to findings from a recent study by Soliman et al.¹⁴ They found that the average number of hours of presenteeism, 5.3 hours per week, was far greater than the number of hours of absenteeism, which was 1.1 hours per week. Furthermore, younger women had significantly higher levels of lost productivity than their older counterparts, and more severe symptoms were associated with more absenteeism and presenteeism. This was in line with our findings, since we also found higher rates of both absenteeism and presenteeism in younger women. A taboo on talking openly about MRS, under-treatment and less flexibility at school might be possible

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explanations for these differences. Comparing our outcomes with studies on other non-gynaecological conditions is hard due to differences in methods and presentation of findings and the cyclic character of MRSs. However, the incidence of presenteeism seems to be as high as it is in patients with inflammatory bowel disease.³¹ Moreover, the amount of work impairment is comparable with severe gastroesophageal reflux (31.9%), moderate irritable bowel syndrome (36.6%) and allergic rhinitis (33.4%-39.8%).³²

Our finding that only 20.1% of women were open about their menstrual symptoms as a reason for calling in sick may confirm the general idea that women tend not to speak openly about MRSs. Wong et al. found that in a cohort of schoolgirls in Malaysia, 76.1% considered dysmenorrhoea a normal part of the menstrual cycle.¹⁵ In the context of the findings noted above, our study also suggests there is a taboo for women in terms of discussing menstrual problems with their employers. The latter may therefore conclude that the impact of MRSs on their employees is negligible. Considering the fact that we also found that 68% of women wish that they had greater flexibility during their periods, either at school or at work, more openness about MRSs in the employment setting seems desirable. The reasons underlying this taboo are likely to have a historical basis; indeed, since ancient times, menstruation has been surrounded with mythical stories and has not been well understood. However, in recent years, the lay literature in developed countries has focused more attention on MRSs. ³³⁻³⁵ The prevalence and the impact of MRSs on the general population and the number of women who are asking for a different approach all reflect the need to change the view of menstrual symptoms and the way they are addressed in society.

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This study consisted of a large cohort, and it reached a large number of women within the age range that was aimed for. The questionnaire was developed in collaboration with patient representatives to make it understandable by and relevant to most women. The cohort appeared to be a representative sample of the general female population based on the number of working hours. ³³ When we compare our data with the national registries the total amount of absenteeism is found to be comparable, regardless of whether it was related to MRSs.^{36 37} It is difficult to compare our numbers on women with a diagnosis explaining their MRSs with numbers found in other studies. We found that only 9% of the participants had such a diagnosis, which seems about as expected or even somewhat low.³ ³⁸⁻⁴⁰ In contrast, 45% of the women in the study reported consulting a physician for their MRSs. This number was relatively high compared to other studies in which, for example, the percentage of women with dysmenorrhoea who sought medical advice was approximately 15%.¹⁵ ¹⁶ An important factor might be the Dutch health system in which general practitioners are available free of charge. Women with a low level of education were relatively underrepresented.⁴¹ As our results show, especially absenteeism related to MRSs is associated with a low level of education, and this might have biased our results. We expect women with lower educational levels to do more physical jobs or jobs with less flexibility. Therefore, our findings on work productivity loss might be underestimated. On the other hand, our finding could be overestimated due to the possibility that women with more MRSs might be more likely to respond to a questionnaire, as it may seem more relevant to them. Moreover, we were not able to provide data on presenteeism not related to MRSs nor were we able to correct for comorbid health conditions. Thus, these results must be interpreted with caution. Due to the way that the questionnaire was distributed through social media, there may have been some selection bias. However, a recent

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review stated that Facebook is a useful recruitment tool for healthcare research.42 Although we did not use a validated guestionnaire, our most important outcomes were based on questions derived from the PCQ, which itself is based on validated questions and which is recommended by guidelines for cost research.²⁴ Self-reported absenteeism generally shows a good correlation with official records, although accuracy decreases with increasing recall period.⁴³ This might have initiated a recall bias in our cohort. It is unknown to what extend recall bias affects reports on presenteeism.⁴⁴ In general, although results vary among studies on premenstrual complaints, a prospective collection of data on symptoms is advisable.^{45 46} Furthermore, an extrapolation of a three and six months timeframe to a yearly basis may intrinsically imply some degree of uncertainty, for example when the influenza season is not included in the original analysis. Finally, these results may not be generalized internationally due to variability in the regulation of social services in different countries, and this is also a limitation of our study. In The Netherlands, wages are paid during sick leave that has duration of less than 1 year, but women in other countries may not have this benefit. Since we know that many factors influence menstrual symptoms, including biological, cultural, and environmental factors, these differences might well influence both absenteeism and presenteeism.⁶¹⁴⁴⁷

In conclusion, we have found that the impact of MRSs on work and school productivity is considerable and that presenteeism contributes significantly more to the matter than absenteeism. Future research should identify how women affected by MRSs could be helped best and how their productivity can be improved in order to reduce the societal impact regarding absenteeism and presenteeism.

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594 **Competing interest statement**

595 All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi disclosure.pdf and declare: no support from any organisation for 596 597 the submitted work; no financial relationships with any organisations that might have an interest in the submitted work in the previous three years; no other relationships or 598 activities that could appear to have influenced the submitted work. 599

600

601 Contributorship statement

TN, BD and JA conceived the study. MS wrote the first and successive drafts of the manuscript. MS, TN and EA modelled and analysed the data. TN, EA, JM, BD and JA contributed to study conception and design. MS and TN collected the data. All authors revised the manuscript for important intellectual content. MS and TN had full access to the data and take responsibility for the integrity of the data and the accuracy of the data analysis. TN is the guarantor.

608

609 Data sharing statement

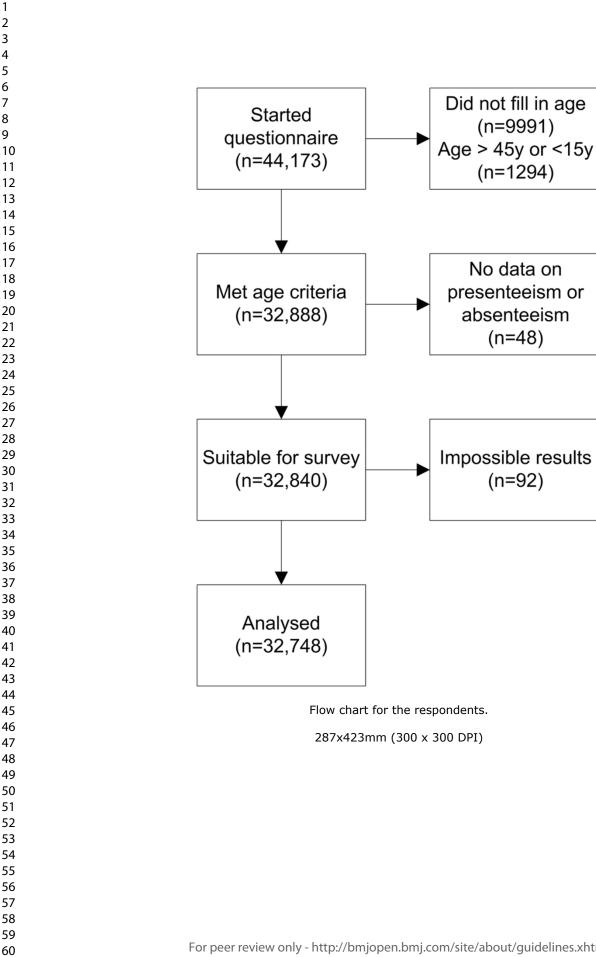
 $\binom{7}{8}$ 610 No additional data from this study are available from a repository. Data are available $\binom{9}{0}$ 611 on request from the corresponding author.

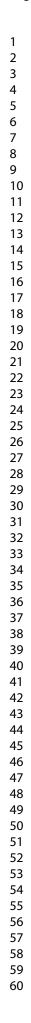
612 Acknowledgements

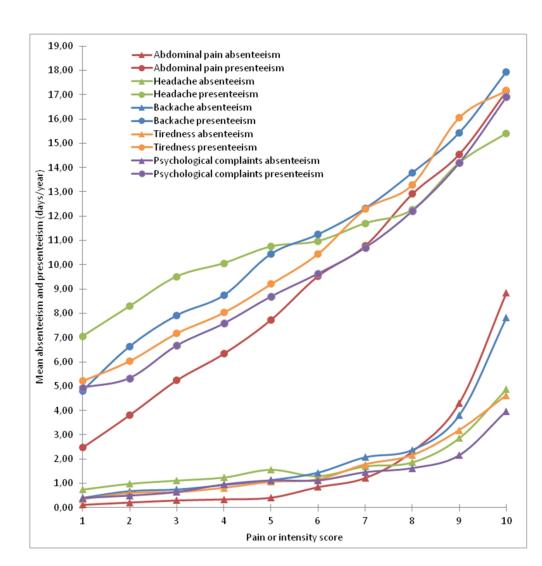
The authors would like to thank Reinoud Oomen, Peter de Vroed, Steven Giesbers,
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contributions in the development and dispersion of the questionnaire.

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2 3	617	Legends to figures
4 5		
6 7	618	
8 9		
10	619	Figure 1
11 12		
13 14	620	Flow chart for the respondents.
15	621	
16 17	021	
18 19	622	Figure 2
20 21		
22 23	623	The relationship between pain and intensity scores, related to absenteeism and
24	624	presenteeism, in lost days per year.
25 26		
27 28	625	
29		
30 31	626	Figure 3
32 33		
34 35	627	The relationship between age and average absenteeism and presenteeism.
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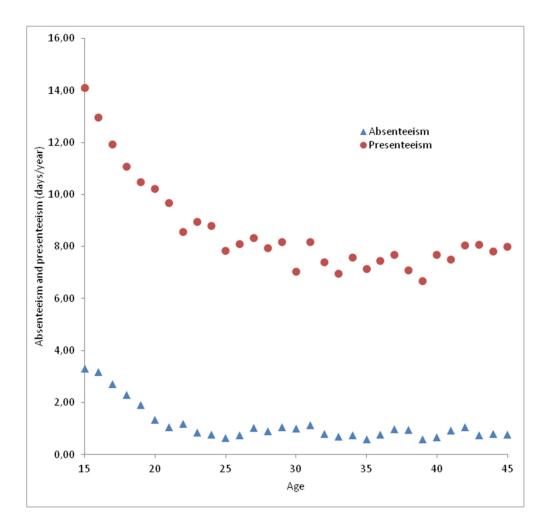






The relationship between pain and intensity scores, related to absenteeism and presenteeism, in lost days per year.

202x209mm (300 x 300 DPI)



The relationship between age and average absenteeism and presenteeism.

206x199mm (300 x 300 DPI)

Work, voluntary work and study: hours per week, sector, experienced stress

Menstrual cycle: total length, number of days with blood loss, number of times

Symptoms: abdominal pain, heavy menstrual blood loss, headache, fatigue,

backache, nausea and vomiting, tender breasts, problems with stool, psychological

Per symptom: number of days the symptom was present; number of days that the

symptom influenced daily functioning; the magnitude of the symptom's influence on

daily functioning; pain or intensity score (not for heavy menstrual bleeding, nausea

and vomiting); continuation of daily activities despite symptoms; and the extent to

3	Appendix
4 5	
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10	Summary of the questionnaire
11	
12	
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14 15	
16	
17	Part 1: Questions about basic characteristics
18	
19	
20	 Age, marital status, nationality, level of education
21	
22	 Work, voluntary work and study: hours per week, sector,
23	
24 25	- Smoking
26	
27	- Menstrual cycle: total length, number of days with blood
28	,
29	needing to change pad or tampon 🛹
30	
31	- Medical and obstetric history
32	- Wieulcal and Obstetric history
33 34	
35	- Contraception use
36	
37	Part 2. Constal questions about monstruction related sumstams
38	Part 2: General questions about menstruation-related symptoms
39	
40	- Symptoms: abdominal pain, heavy menstrual blood loss,
41	- Symptoms, abdominal pain, neavy menstrual blood loss,
42 43	
44	backache, nausea and vomiting, tender breasts, problems
45	
46	complaints
47	
48	 Per symptom: number of days the symptom was present;
49	
50 51	symptom influenced daily functioning; the magnitude of t
52	
53	daily functioning; pain or intensity score (not for heavy m
54	
55	and vomiting); continuation of daily activities despite sym
56	and volume, continuation of daily activities despite sym
57	
58 50	which women forced themselves to continue their daily a
59 60	
00	 Usage of over-the-counter drugs, narcotics, painkillers, al

Usage of over-the-counter drugs, narcotics, painkillers, alternative medicine

which women forced themselves to continue their daily activities

Part 3: Questions related to absenteeism and presenteeism (full text)

- How often did you call in sick due to menstrual complaints in the past six months? _
 - Every menstrual cycle
 - Almost every menstrual cycle
 - Half of all menstrual cycles
 - Sometimes
 - Never Ο
- What is the mean number of days you called in sick due to menstruation-related _

symptoms per cycle in the past six months?

- More than three days
- Three days Ο
- Two days Ο
- One day Ο
- ė Lieu Half a day or a few hours
- What is the number of days you were present at work while being hindered by menstrual complaints?
- How was your performance during these days?
 - o A Visual Analog Scale by means of sliding a bar was used to score the amount of productivity loss, where the left end said "I could not do anything" and the right end said "I could do just as much as I normally do".

2	
3 _	What was the number of days you called in sick because of non-menstruation-related
4	
5	symptoms in the past six months?
6	symptoms in the past six months:
7	
8	
9	
10	What reason do you give when you call in sick due to menstruation-related
11 -	
12	
13	symptoms?
14	
15	 Menstrual complaints
16	
17	 Only the presenting symptom, for example abdominal pain or headache
18 19	• Only the presenting symptom, for example abdominal pair of neaddene
20	
20	 You do not give a reason
22	
23	 You think up another reason
24	
25	
26	
27	
28 -	What measures would be desirable for women on the workplace or at school during
29	
30	their menstrual period? It is possible to give more than one answer.
31	
32	
33	 A day off without any consequences
34	
35	\circ A day off to catch up later \sim
36	
37	 Flexibility to perform less physical work
38	o meximity to perform less physical work
39	
40	 More time for personal care
41	
42	• The possibility to work from home
43	
44	• None
45	 None
46	
47	 Not applicable
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STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of cross-sectional studies

Section/Topic	ltem #	Recommendation	Reported on page #
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	2
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	4
Objectives	3	State specific objectives, including any prespecified hypotheses	5
Methods			
Study design	4	Present key elements of study design early in the paper	5
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	5-7
Participants 6 (a) Give the eligibility criteria, and the sources and methods of selection of participants		6	
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	8, appendix
Data sources/ 8* For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe measurement comparability of assessment methods if there is more than one group		7-8	
Bias	9	Describe any efforts to address potential sources of bias	7-9
Study size	10	Explain how the study size was arrived at	7
Quantitative variables 11 Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why		7-9	
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	7-8
		(b) Describe any methods used to examine subgroups and interactions	7-8
		(c) Explain how missing data were addressed	7,9
		(d) If applicable, describe analytical methods taking account of sampling strategy	n/a
		(e) Describe any sensitivity analyses	n/a
Results			

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Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility,	9
		confirmed eligible, included in the study, completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	9/Figure 1
		(c) Consider use of a flow diagram	Figure 1
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	9/Table 1
		(b) Indicate number of participants with missing data for each variable of interest	Not reported, can be reported if desirable.
Outcome data	15*	Report numbers of outcome events or summary measures	9-11/Table 2,3
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence	Table 4
		interval). Make clear which confounders were adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	Table 1,4
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	n/a
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	n/a
Discussion			
Key results	18	Summarise key results with reference to study objectives	11-12
Limitations 19 Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias		14-15	
Interpretation			12-15
Generalisability	21	Discuss the generalisability (external validity) of the study results	14-15
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	n/a

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

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Productivity loss due to menstruation-related symptoms: a nationwide cross-sectional survey among 32,748 women

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Primary Subject Heading :	Obstetrics and gynaecology
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Keywords:	Menstruation, Community gynaecology < GYNAECOLOGY, Menstrual cycle, Presenteeism, Absenteeism



1 2						
2 3 4	1	Productivity loss due to menstruation-related symptoms: a nationwide cro				
5 6 7	sectional survey among 32,748 women					
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28 29 30	31						
31 32 33 34	32	Word count: 4,058					
35 36 37	33						
38 39	34	Key Words: Absenteeism, Presenteeism, Menstrual cycle, Menstruation, Community					
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1 2		
3 4 5	36	Abstract
6 7	37	Objective
8 9	38	To evaluate age-dependent productivity loss caused by menstruation-related
10 11 12	39	symptoms, measured in absenteeism (time away from work or school) and
13 14	40	presenteeism (productivity loss while present at work/school).
15 16 17 18	41	
19 20	42	Methods
21 22	43	Design/Setting: Internet-based, cross-sectional survey conducted in the Netherlands
23 24 25 26	44	from July to October 2017.
20 27 28 29	45	Participants: 32,748 women aged 15 to 45 years, recruited through social-media.
30 31	46	Outcome measures: Self reported lost productivity in days, divided into absenteeism
32 33	47	and presenteeism; impact of menstrual symptoms; reasons women give when calling
34 35 36	48	in sick; and women's preferences regarding the implications of menstruation related
37 38	49	symptoms for schools and workplaces.
39 40 41	50	<u>Results</u>
42 43	51	A total of 13.8% (n=4,514) of all women reported absenteeism during their menstrual
44 45 46	52	periods with 3.4% (n=1,108) reporting absenteeism every or almost every menstrual
47 48	53	cycle. The mean absenteeism related to a woman's period was 1.2 days per year. A
49 50	54	total of 80.7% (n=26,438) of the respondents reported presenteeism and decreased
51 52 53	55	productivity a mean of 23.2 days per year. An average productivity loss of 33%
54 55	56	resulted in a mean of 8.9 days of total lost productivity per year. Women under 21
56 57	57	were more likely to report absenteeism due to menstruation-related symptoms (OR
58 59 60	58	3.3, 95% confidence interval 3.1 to 3.6). When women called in sick due to their
00	59	periods, only 20.1% (n=908) told their employer or school that their absence was due

to menstrual complaints. Notably, 67.7% (n=22,154) of the participants wished they had greater flexibility in their tasks and working hours at work or school during their periods. Conclusions Menstruation-related symptoms cause a great deal of lost productivity, and presenteeism is a bigger contributor to this than absenteeism. There is an urgent need for more focus on the impact of these symptoms, especially in women aged under 21, for discussions of treatment options with women of all ages, and, ideally, more flexibility for women who work or go to school. Strengths and limitations of this study This is the largest cohort study to analyse the impact of menstruation-related symptoms on work and school productivity. The survey was performed among the general female population and is consequently not per se related to one specific gynaecological condition. Due to the way of recruitment of participants, there may have been some degree of selection bias. Outcomes are based on self reported data and consequently lack objectivity regarding productivity loss The generalisability of the study may be limited to employment and school systems comparable to the Dutch.

Background

Menstruation-related symptoms (MRSs) are diverse and widespread among women. Symptoms include dysmenorrhea, heavy menstrual bleeding and premenstrual mood disturbances with reported prevalence of 45% to 90%, 14% to 25%, and 20% to 29% respectively.¹⁻³ Studies show that women with MRSs have lower scores on several domains of quality of life such as general health and physical, mental, social, and occupational functioning during their periods.^{1 4-7} Furthermore, these symptoms may create considerable financial burdens on patients and their families as well as on society.^{5 6 8-12} Such financial burdens are related to the costs of visits to the doctor, over-the-counter drugs, and medical or surgical treatment. However, costs related to productivity loss could be the largest cost driver. Productivity costs are defined as costs associated with paid and unpaid production loss and the replacement of productive people due to illness or disability.¹³ Productivity costs can be divided into costs related to absenteeism and costs related to presenteeism. Absenteeism represents the total amount of time off work or away from school, and presenteeism represents the loss of productivity while present at a job or school.

Although the literature is scarce and the results are variable, studies on specific patient groups generally show that MRSs can cause absenteeism.¹⁴⁻¹⁶ Research on the association between MRSs and presenteeism is even more limited. It has been suggested that research into possible impairments in quality of life caused by menstrual symptoms should not focus on single symptoms but rather on a complex of symptoms that vary widely but that are all related to the menstrual cycle. This complex includes both standard symptoms, like heavy menstrual bleeding and abdominal cramps, and also less common symptoms, like nausea and cold sweats.

Taking all symptoms into account, it seems likely that the real impact of MRSs is underestimated in the general population. Despite being almost two decades into the 21st century, discussions about MRSs may still be rather taboo. This survey-based exploratory study aimed to quantify the burden of MRSs in the general female population, with burden defined as the number of lost days at work or school due to absenteeism and presenteeism. Furthermore, it was aimed to study the impact of specific symptoms on absenteeism and presenteeism.

116 Methods

This cross-sectional study consisted of an online survey that was distributed from July 12 to October 11, 2017. Approval for this study was obtained from the local medical ethics committee "Commissie Mensgebonden Onderzoek (CMO)" under number file number 2017-3387 on July 12th 2017. All data were anonymously collected and stored under the privacy rules of the Radboud University Medical Center. Patients gave informed consent when they initiated the questionnaire.

124 Patient and Public Involvement

A group of women, among which were several members of the Dutch Patient Endometriosis Foundation, women with a linguistic education and women with a medical origin, was involved in the conduct of this study at several stages; i.e. in the development and dissemination of the questionnaire and in the analysis and interpretation of the results. One of the authors of this manuscript, BD, is the chair of the Dutch Patient Endometriosis Foundation. Additional contributions are noted in the Acknowledgements section.

1 2		
3 4	132	
5 6 7	133	Questionnaire development
8 9 10	134	The questionnaire had several parts, and Appendix 1 provides details about the
11 12	135	questions. Part 1 consisted of questions about each woman's basic characteristics.
13 14	136	Part 2 had questions about menstrual symptoms, and part 3 had questions related to
15 16 17	137	absenteeism and presenteeism. Adaptive questioning was used with a maximum of 6
18 19	138	questions per page. Participants were asked in a lay manner how long their
20 21	139	menstrual cycle was and what the exact meaning of a menstrual cycle was. The
22 23 24	140	duration of the cycle was divided in 5 categories (25 days or less, 26-30 days, 31-35
25 26	141	days, 36-40 days and 41 days or more). Furthermore participants could indicate if
27 28	142	they had an irregular cycle, meaning more than 10 days difference per cycle, if they
29 30 31	143	were amenorrheic due to the use of an Intra-Uterine Device (IUD) or the continuous
32 33	144	use of oral contraceptives, or the option "I do not know". Additional questions about
34 35	145	absenteeism and presenteeism were included that were based on the Productivity
36 37 38	146	Cost Questionnaire from the Institute for Medical Technology Assessment (iMTA-
39 40	147	PCQ). ¹⁹ We modified the iMTA-PCQ-recommended recall period of four weeks to
41 42	148	three months so that it was in line with the relevant time period for this study and so
43 44	149	we could include multiple menstrual periods. We assumed the amount of
45 46 47	150	presenteeism to be larger than the amount of absenteeism. Therefore, the recall
48 49	151	period for absenteeism was extended to six months to maintain accuracy. Visual
50 51	152	analogue scales (VAS) were used to quantify the amount of pain, or the intensity of
52 53 54	153	the symptom, and the impairment due to pain or the other symptom. Presenteeism
55 56	154	was measured by asking women to what extent they were able to be as productive
57 58	155	as possible compared to a day without MRSs. This was scored on a scale from 0 to
59 60	156	10, with 0 being totally unproductive and 10 fully productive. In separate questions,

participants were asked to quantify their absenteeism that was related to MRSs and
absenteeism for any other reason than MRSs. For the latter, we did not specifically
ask the underlying reason.

161 Target population and recruitment

The study population comprised of women between 15 and 45 years old. The upper age limit was chosen to avoid interference from menopausal symptoms; the lower to have a time margin after the average menarche age, since it is known that the first periods are irregular and often accompanied with discomfort and uncertainty. A large number of women were approached with the aim of obtaining a cohort that was representative of the general female population in terms of level of education, medical history, and/or gynaecological diagnosis. Women were invited to complete a survey using an online questionnaire tool²⁰ through a campaign on social media platforms Facebook and Twitter. Patient organisations, colleagues, and visitors of the Facebook page of one of the authors (TN) were asked to share the link to the questionnaire in order to reach the widest possible audience.

On July 12, 2017, a link to the questionnaire was posted on Facebook and Twitter through the account of one of the authors (TN). In the post both women with and without MRSs were encouraged to fill in the questionnaire. Within 24 hours of the first posting on social media, over 6,000 respondents had filled in the questionnaire, and by July 18, there were 15,000 respondents, which was announced by a re-post of the link to the guestionnaire. A third post was made on Facebook and Twitter on September 16, 2017, to reach women who may have been on holiday when the first posts were created.

1 2		
3 4 5	181	
6 7 8 9 10 11 12	182	Data analysis
	183	The outcome measures were presented in a descriptive way; we used valid
	184	percentages in case of missing values where necessary. We distinguished between
13 14 15	185	women who were mainly working or mainly studying. Therefore, we present these
16 17	186	data for two groups i.e. for women who worked more than 5 hours per week
18 19 20	187	("working group") and for women who studied more than 5 hours per week ("studying
21 22	188	group").
23 24 25	189	We used binary logistic regression to calculate odds ratios. Absenteeism and
26 27	190	presenteeism were used as dependant variables. As independent variables we used
28 29	191	the following parameters: women younger than 21 versus women aged 21 and older,
30 31 32	192	smoking yes or no, reports of absenteeism not related to MRSs, educational level,
32 33 34	193	the use of oral contraception and the use of an levonorgestrel-releasing intra uterine
35 36	194	device. All independent variables were used in an univariate as well as a multivariate
37 38 39	195	analysis. We also studied the association between pain scores and both
40 41	196	absenteeism and presenteeism, given that the literature shows that pain scores of 0
42 43	197	to 4, 5 to 6, and 7 or higher have a different impact on activity, mood, and sleep. ^{21 22}
44 45 46	198	Analyses were performed using IBM SPSS Statistics version 22.00.
47 48 49 50 51 52 53 54 55	199	
	200	Assumptions and transformation of the original data
	201	To present data on level of education in an international format we had to transform
56 57	202	the original data, which was based on the Dutch school system.23 The categorical
58 59 60	203	data of participants' length of menstrual cycle were transformed into averages.

With regard to the evaluation of absenteeism and presenteeism, "the guideline for economic evaluations in healthcare in the Netherlands" was used.²⁴ A work-day accounts for 8 hours. For most sectors in the Netherlands, a full-time work-week is 36 hours. The maximum amount of working hours per year was set at 1558 when they were working full time. We asked women to report their absenteeism due to MRSs per cycle and used a recall period of six months.

To calculate the percentages for absenteeism, one day of absenteeism accounted for 8 hours of lost productivity. When a woman reported to study or work more than 40 hours per week, we transformed these hours to 40 for reasons of clarity in the calculations and comparability with the data of the Dutch Central Bureau of Statistics, CBS. We made a few other transformations for categorical data. For absenteeism related to MRSs, the category "more than three days per cycle" was considered to be 4 days per cycle. For absenteeism that was not related to MRSs, the category "more than ten days in the past six months" was considered to be 11 days in the past six months.

To present yearly data, we multiplied some of these data based on the original recall period. The number of days for absenteeism related to MRSs was based on days per cycle, which were therefore multiplied by 12.7 based on the reported average menstrual cycle of 28.8 days, see table 1. These values were then multiplied by one if the woman reported that she called in sick "every period", 0.75 if she reported "almost every period", 0.5 if she reported "half of all periods", and 0.25 if she reported calling in sick "only once in a while". Values for absenteeism that was not related to MRS were based on a recall period of 6 months and were therefore multiplied by two in order to obtain the number of days per year. The values for presenteeism were based on a recall period of three months and were therefore multiplied by four.

2 3 4	229				
5 6 7	230	Results			
8 9 10	231	A total of 44,173 women initiated the ques	tionnaire. We ex	cluded partic	ipants who
11 12	232	did not report a date of birth or whose age d	id not fulfil the ind	clusion criteria	a (figure 1).
13 14 15	233	There were no duplicates of IP addresses	. Women who di	d not answei	questions
16 17	234	related to absenteeism and presenteeism	were excluded. F	Furthermore,	cases with
18 19	235	impossible results (e.g. 10,000,000 days o	of presenteeism	in three mon	ths or 140
20 21 22	236	changes of sanitary pads a day) were exc	luded. This resul	ted in a total	l of 32,748
22 23 24	237	women in the final analysis.			
25 26	238	Table 1 summarizes the basic characteris	stics of the parti	cinants We	found that
27 28					
29 30	239	45.4% (n=14,839) had visited a doctor for	menstrual compl	aints in the p	ast, with a
31 32	240	total of 3017 (9.2%) women reporting a diagnosis of a menstrual disorder, such as			
33 34	241	endometriosis or fibroids.			
35 36 37	242	The mean age of women in the working g	roup was higher	than the me	an ager of
38 39	243	women in the studying group. The mean number of working hours per week in the			
40 41 42	244	working group was 27.0 (SD 11.4), and the mean number of study hours in the			
43 44	245	studying group was 27.4 (SD 12.1). A tota	l of 7,335 wome	n (22.4%) rep	oorted both
45 46	246	working and studying more than five hours	s per week. In th	iis group, 3,0	01 women
47 48 40	247	were working more than 16 hours a week, and 5,284 women in the study group were			
49 50 51	248	studying more than 16 hours a week.			
52 53 54 55			Number (percentage)	Mean ± SD	Median
56 57		Age, years		28.6±8.6	28
58 59		15-19	6,141 (18.8%)		
60		20-24	6,118 (18.7%)		
					11

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25-29	5,825 (17.8%)		
30-34	5,483 (16.7%)		
35-40	4,687 (14.3%)		
41-45	4,494 (13.7%)		
Level of education			
Low	4,020 (12.3%)		
Medium	12,335 (37.9%)		
High	16,229 (49.8%)		
Hours/week			
Paid work		21.7±14.7	24
Study		7.4±13.6	0
Voluntary work		0.8±3.1	0
Menstrual cycle			
Regular cycle	25,717 (78.5%)		
Duration	4.	28.8±3.0	28
Amenorrhoea due to LG-IUD/OC	3,675 (11.2%)		
Irregular, variation >10 days per cycle	2,495 (7.6%)		
Do not know	861 (2.6%)		
Days with blood loss per cycle	0,	5.4±1.6	5
Visited a doctor for MRSs			
Νο	17,873 (54.6%)		
Yes, general practitioner	10,141 (31.0%)		
Yes, gynaecologist	4,698 (14.4%)		
Diagnosis for MRSs*			
Νο	29,731 (90.8%)		
Yes	3,017 (9.2%)		
Endometriosis	1,120 (3.4%)		
PCOS	588 (1.8%)		
Adenomyosis	103 (0.3%)		

Fibroids	275 (0.8%)		
Other	1,901 (5.8%)		
Contraception*			
Hormonal contraception	11,993 (36.6%)		
OC	8,650 (26.4%)		
LG-IUD	2,752 (8.4%)		
Other hormonal: injection, transdermal etc.	882 (2.7%)		
No hormonal contraception	20,755 (63.4%)		
Cu-IUD	771 (2.4%)		
Female sterilisation	423 (1.3%)		
No female contraception	19,639 (60.0%)		
Nulliparous	21,585 (66.0%)		
Paid work >5 hr a week	26,104 (79.7%)		
Age		29.7±8.3	
Hours of paid work/week	2	27.0±11.4	
Hours spent on study/week		7.5±13.4	
>40 hours of paid work/week	1,047 (3.2%)		
Study >5 hr a week	8,764 (26.8%)		
Age		22.0±6.2	
Hours spent on study/week		27.4±12.1	
Hours of paid work/week		15.5±11.3	
>40 hours spent on study/week	322 (1.0%)		

Table 1 | Basic characteristics of study participants (n=32,748) Mean duration of cycle based on women with a regular cycle. SD = standard deviation, MRSs = menstruation-related symptoms, PCOS = polycystic ovary syndrome, OC = oral contraceptive, LG-IUD = levonorgestrel-releasing intrauterine device, Cu-IUD = copper intrauterine device. *More than one answer possible.

60 255

256 Absenteeism

Table 2 shows the results on absenteeism due to MRSs. Although 13.8% of the women (n=4,514) reported absenteeism due to MRSs, only 1,108 women (3.4%) reported absenteeism every cycle or almost every cycle. The percentage of absenteeism in every cycle or almost every cycle was 2.4% in the working group and 4.5% in the studying group. The mean absenteeism due to MRSs was 0.9 days per year for the working group and 1.6 day per year for the study group.

We also calculated the mean total absenteeism that was not related to MRSs. For the entire group, this was 3.3 days per year; for the working group, it was 3.5 days, and for the studying group, it was 4.3 days. The mean percentage of absenteeism that was not related to MRSs was 3.5% in the working group and 3.7% in the studying group. Consequently, absenteeism due to MRSs in our cohort accounted for, on average, 22% of the total absenteeism in the working group and 24% in the studying elez oni group.

	Number (percentage)	Mean ± SD
All (n=32,748)		
Absenteeism	4,514 (13.8)	
≤0.5 day	538 (1.6%)	
1 day	2,259 (6.9%)	
2 days	1,171 (3.6%)	
3 days	349 (1.1%)	
>3 days	184 (0.6%)	
Total days of absenteeism per year		1.3±5.9
Work (n=26,104)		
Absenteeism	2,926 (11.2%)	
≤0.5 day	374 (1.4%)	
1 day	1,476 (5.7%)	
2 days	757 (2.9%	
3 days	211 (0.8%)	
>3 days	98 (0.4%)	
Total days of absenteeism per year	2	0.9±3.9
Study (n=8,764)	0	
Absenteeism	1,715 (19.6%)	
≤0.5 day	234 (2.7%)	4
1 day	921 (10.5%)	
2 days	423 (4.8%)	
3 days	92 (1.0%)	
>3 days	41 (0.5%)	
Total days of absenteeism per year		1.6±5.0

Table 2 | Reported absenteeism caused by menstruation-related symptoms. Women
were asked to report the average amount of days on which they were absent due to
menstruation-related symptoms per cycle. The total days of absenteeism per year

3 4 5 6	275 276 277	was calculated. The added numbers of wome the total amount of participants, since 2,120 more than 5 hours/week. SD = standard devi	women reported	, , ,	
7 8	278				
9 10 11	279	Presenteeism			
12 13 14	280	Table 3 shows the numbers reported for	presenteeism. O	ver 80% of a	III women
15 16	281	reported presenteeism during their periods	. The difference	s between the	e working
17 18	282	group and the study group were not large in	terms of prevalen	ice and lost pro	oductivity.
19 20 21	283	The mean number of lost productive days p	per year due to p	presenteeism	was more
22 23	284	than seven-fold greater than the mean n	umber of lost p	roductive day	rs due to
24 25	285	absenteeism.			
26 27					
28 29			Number (percentage)	Mean ± SD	Median
30 31		All (==22,749)			
32		All (n=32,748)			
33 34 35		Presenteeism	26,438 (80.7%)		
36 37		Number of days in the past 3 months	4	5.8±5.3	5.0
38 39		Percentage of productivity loss per day		33.0±24.8	30.0
40 41		Days/year of lost productivity		8.9±11.0	5.6
42 43 44		Work (n=26,104)			
44 45		Presenteeism	21,252		
46			(81.4%)		
47				5 7 . 5 0	= 0
48 49		Number of days in the past 3 months		5.7±5.2	5.0
50 51		Percentage of productivity loss per day		31.7±24.7	30.0
52 53		Days/year of lost productivity		8.4±10.6	4.8
54 55		Study (n=8,764)			
56 57		Presenteeism	7,385 (84.3%)		
58 59		Number of days in the past 3 months		6.3±5.3	5.0
60		Deveenters of productivity loss per dev		20.0124.2	40.0

Percentage of productivity loss per day

1 2

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16

36.8±24.2

40.0

Smoking^b

Absenteeism not

related to MRSs in

the past six months^c

Level of education^d

54 55 56

57

58

59 60

	Days/year of lost proc	ductivity		10.5±11.8	7.2
286			<u> </u>		
287	Table 3 Reported pres	enteeism caused by r	menstruation-r	elated symptom	s. Wome
288	were asked to report the	e amount of days on w	which they we	e less productiv	e and to
289	what extent. The total da	ays of lost productivity	y per year was	s calculated. The	added
290	numbers of women in th	e work and study gro	up exceed the	e total amount of	
291	participants, since 2,120) women reported to b	both study and	I work more thar	1 5
292	hours/week. SD = stand	lard deviation			
293					
204	Eastern approxisted with	obcontaciom and pro	aantaaiam		
294	Factors associated with	absenteeism and pre	SenleelSM		
295	Figure 2 shows the ass	ociation between repo	orted pain or c	liscomfort score	s and bot
296	absenteeism and prese	nteeism. As seen in	detail in table	4, high VAS so	ores wer
297	significantly associated	with higher levels of	of absenteeisr	n and presente	eism. Th
298	strongest relationship w	as found for abdomin	al pain scores	that were 7 or h	nigher on
299	scale from 0 to ten. Od	ds ratios were 5.6 for	r absenteeism	(95% confidence	ce interva
200	E(0, to, G(2)) and $Q(0, for)$	proportagion (DEO/	opfidance int	much 0,1 to 0,5	
300	5.0 to 6.2) and 8.8 for	presenteeism (95% C	confidence inte	erval: 8.1 to 9.5). Figure
301	shows the association	between age and be	oth presentee	ism and absent	eeism. A
202	abour is both figure	2 and table 4 wa	found that		n ohouvo
302	shown in both figure	5 and table 4, We	e iound that	younger wome	n snowe
303	significantly higher rat	es of absenteeism	and present	eeism. A levor	orgestrel
304	releasing intrauterine de	evice is associated wi	th especially le	ess presenteeisr	n.
		Absenteeism		Presenteeism	
		OR (95% CI)	OR after	OR (95% CI)	OR af
			correction (95% CI)		correct (95% (
A	ge<21 years ^a	3.7 (3.4-3.9)	3.3 (3.1-3.6)	1.4 (1.3-1.5)	1.3 (1.2-
_					

1.3 (1.2-1.5)

2.2 (2.1-2.4)

1.3 (1.2-1.4)

1.7 (1.6-1.9)

1.5 (1.3-1.6)

1.4 (1.3-1.5)

1.4 (1.3-1.6)

1.3 (1.2-1.4)

	Low	4.5 (4.1-4.9)	2.7 (2.4-3.0)	1.3 (1.2-1.4)	1.1 (1.0-1.2
	Medium	2.2 (2.1-2.4)	1.7 (1.5-1.8)	1.3 (1.2-1.4)	1.2 (1.1-1.2
	High	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
Oral contraception ^e					
	No	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	Yes	1.2 (1.1-1.3)	1.0 (0.9- 1.1)**	0.9 (0.9-1.0)	0.9 (0.8-0.9
LG-IUD ^e					
	No	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	Yes	0.7 (0.6-0.8)	0.9 (0.8-1.0)	0.5 (0.5-0.6)	0.5 (0.5-0.6
Abdominal pain score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	2.6 (2.3-2.9)	2.2 (1.9-2.4)	5.2 (4.8-5.7)	5.3 (4.9-5.7
	>7	7.0 (6.4-7.8)	5.6 (5.0-6.2)	8.7 (8.0-9.4)	8.8 (8.1-9.5
Headache pain score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	1.5 (1.3-1.6)	1.5 (1.4-1.6)	3.0 (2.7-3.3)	3.1 (2.8-3.4
	>7	2.0 (1.8-2.1)	2.3 (2.1-2.5)	3.5 (3.2-3.9)	3.7 (3.4-4.1
Backache pain score ^e				1	
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	1.6 (1.5-1.7)	1.4 (1.3-1.5)	3.5 (3.2-3.9)	3.5 (3.2-3.8
	>7	2.7 (2.5-2.9)	2.2 (2.1-2.4)	4.7 (4.2-5.2)	4.5 (4.0-5.0
Tiredness intensity score ^e					
	0-4	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)	1.0 (n/a)
	5-6	1.8 (1.7-2.0)	1.8 (1.6-2.0)	3.3 (3.1-3.6)	3.3 (3.1-3.6
	>7	3.0 (2.8-3.2)	2.8 (2.6-3.1)	5.1 (4.7-5.6)	5.2 (4.7-5.7
Psychological complaints intensity					

	1.0 (n/a) 1.5 (1.4-1.7)	1.0 (n/a)	1.0 (n/a)
	1.5 (1.4-1.7)	0.7/0.5.00	
>7 2.2 (2.0-2.4)		2.7 (2.5-2.9)	2.6 (2,5-2.9)
	2.1 (2.0-2.3)	4.4 (4.0-4.7)	4.3 (4.0-4.7)
305			
Table 4 Odd ratios (ORs) and 95% confidenc	e intervals (9	5% CIs) for fact	tors
related to absenteeism and presenteeism. OR	s >1 correlate	e with a higher p	orevalence
of absenteeism or presenteeism. ORs <1 corre	elate with a lo	wer prevalence	e of
absenteeism or presenteeism.			
LG-IUD = Levonorgestrel-releasing intra uterin	e device		
^a Correction for smoking and absenteeism that	was not relat	ed to menstrua	tion-
12 related symptoms (MRSs); ^b Correction for age	, absenteeisr	n that was not r	elated to
13 MRSs, and level of education; Correction for a	age, smoking	, and level of ed	Jucation;
¹⁴ ^d Correction for age, smoking, and absenteeism	n that was no	t related to MRS	Ss;
¹⁵ ^e Correction for age, smoking, absenteeism tha	it was not rela	ated to MRSs, a	and level of
education. *p=0.26, **p=0,73 For all other OR	s, p<0.05		
17			
Menstruation and suggested implications for so	chools and w	orkplaces	
19 From the respondent who had been calling in	sick due to N	MRSs, 20.1% (r	n=908) told
their employer or school menstrual symptoms	were the rea	son, 46.4% (n=	2092) only
21 mentioned the presenting symptom. No reaso	on was given	by 27.7% (n=1)	250), while
5.8% (n=260) made up another reason. Co	omparing wor	nen aged belc	w 21 with
women aged 21 and above, we found that y	ounger wom	en were less c	open about
their MRSs being the reason for calling in sick	k (12.0%) ver	rsus women old	ler than 21
(27.0%). Women were asked to report su	iggestions oi	n how work p	laces and
		nction better d	uring their
326 conditions could be changed in order for	them to tur		<u> </u>
conditions could be changed in order for menstrual periods. There were 32,708 respon			C

(67.7%, n=22,154) preferred more flexibility during their periods, such as the possibility of doing less physical work (32.1%, n=10,499), the ability to work from home (39.5%, n=12,917), more time for personal care (28.3%, n=9,241), or the ability to take a day off and make up for it later (11.5%, n=3,756). In addition, 32.9% wished they could take a complete day off without any consequences. 27.2% (n=8,890) did not wish for any changes in policy. Many women (79.7%, n=26,072) were open to -> ompany .tention to MR: discussing MRSs with their company doctor, and 56.7% (n=18,579) thought that doing so would draw more attention to MRS-related matters.

Discussion

This survey-based study showed that menstruation-related absenteeism and, to a greater extent, presenteeism are widespread in the general female population. In our cohort, MRSs accounted for up to 24% of total absenteeism for women who were working and studying. The annual productivity loss due to presenteeism was seven-fold times more than the annual productivity loss due to absenteeism and women younger than 21 years experience the largest burden. Symptom severity scores showed significant and strong associations with both absenteeism and presenteeism. When women called in sick due to MRSs, only one in five stated openly that menstrual symptoms were the main reason. A majority of women prefers more flexibility during their periods when it comes to work or school.

There have been few studies on absenteeism and presenteeism related to MRSs in the general female population. To our knowledge, Tanaka's study ²⁵ is the only other published study on absenteeism and presenteeism due to MRSs in the general female population. In a cohort of 19,254 Japanese women, a total of 3,311 (17.2%) reported work productivity lost in the prior 3 months, mostly in the form of decreased efficiency (62.0%, n=2,052). Of these 2,052 subjects, the mean number of workdays lost due to decreased efficiency was 5.7 days in 3 months. After recalculation, this accounts for 2.4 days per year for the entire population. This is fewer days than the 8.9 days per year in our cohort. On the other hand, the numbers for absenteeism were more similar, with a mean of 1.0 day of absenteeism per year in the entire Japanese cohort compared to 1.2 days in our cohort. Differences in regulations of social services, a difference in attitude towards sick leave and a different method of data collection might explain these differences. It has been suggested in research on musculoskeletal symptoms that rates of absenteeism might be lower in Japan

362 compared to European countries and the United States. Consequently, presenteeism
 363 might therefore be a more representative variable.^{26 27}

More data are available regarding the impact of dysmenorrhoea on guality of life and absenteeism. De Sanctis et al. reviewed studies on dysmenorrhea in multiple countries, some of which included menstruation-related absenteeism data. They found that the prevalence of school absences in adolescents that was due to dysmenorrhea varied between 7.7% and 57.8%. Since the review included 41,140 women in 27 countries, and there was a high degree of heterogeneity in the outcome measures, no firm conclusions could be drawn. Hailemeskel et al. evaluated 440 female university students in Ethiopia.²⁸ Among students with dysmenorrhea, 66.8% reported a loss of concentration in class, and 56.3% reported class absences during the last month. In a questionnaire-based study of 706 Hispanic female adolescents, 38% reported missing school due to dysmenorrhea during the 3 months prior to the survey, and 59% reported a decrease in concentration in class due to dysmenorrhea.

Absenteeism and presenteeism due to endometriosis in other studies was greater than in our study, which was not surprising.^{9 14 30} However, some interesting parallels can be drawn to findings from a recent study by Soliman et al.¹⁴ They found that the average number of hours of presenteeism, 5.3 hours per week, was far greater than the number of hours of absenteeism, which was 1.1 hours per week. Furthermore, younger women had significantly higher levels of lost productivity than their older counterparts, and more severe symptoms were associated with more absenteeism and presenteeism. This was in line with our findings, since we also found higher rates of both absenteeism and presenteeism in younger women. A taboo on talking openly about MRS, under-treatment and less flexibility at school might be possible

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explanations for these differences. Comparing our outcomes with studies on other non-gynaecological conditions is hard due to differences in methods and presentation of findings and the cyclic character of MRSs. However, the incidence of presenteeism seems to be as high as it is in patients with inflammatory bowel disease.³¹ Moreover, the amount of impairment is comparable with severe gastroesophageal reflux (31.9%), moderate irritable bowel syndrome (36.6%) and allergic rhinitis (33.4%-39.8%).³²

Our finding that only 20.1% of women were open about their menstrual symptoms as a reason for calling in sick may confirm the general idea that women tend not to speak openly about MRSs. Wong et al. found that in a cohort of schoolgirls in Malaysia, 76.1% considered dysmenorrhoea a normal part of the menstrual cycle.¹⁵ In the context of the findings noted above, our study also suggests there is a taboo for women in terms of discussing menstrual problems with their employers. The latter may therefore conclude that the impact of MRSs on their employees is negligible. Considering the fact that we also found that 68% of women wish that they had greater flexibility during their periods, either at school or at work, more openness about MRSs in the employment setting seems desirable. The reasons underlying this taboo are likely to have a historical basis; indeed, since ancient times, menstruation has been surrounded with mythical stories and has not been well understood. However, in recent years, the lay literature in developed countries has focused more attention on MRSs. ³³⁻³⁵ The prevalence and the impact of MRSs on the general population and the number of women who are asking for a different approach all reflect the need to change the view of menstrual symptoms and the way they are addressed in society.

This study consisted of a large cohort, and it reached a large number of women within the age range that was aimed for. The questionnaire was developed in collaboration with patient representatives to make it understandable by and relevant to most women. The cohort appeared to be a representative sample of the general female population based on the number of working hours. ³³ When we compare our data with the national registries the total amount of absenteeism is found to be comparable, regardless of whether it was related to MRSs.^{36 37} It is difficult to compare our numbers on women with a diagnosis explaining their MRSs with numbers found in other studies. We found that only 9% of the participants had such a diagnosis, which seems about as expected or even somewhat low.³ ³⁸⁻⁴⁰ In contrast, 45% of the women in the study reported consulting a physician for their MRSs. This number was relatively high compared to other studies in which, for example, the percentage of women with dysmenorrhoea who sought medical advice was approximately 15%.¹⁵ ¹⁶ An important factor might be the Dutch health system in which general practitioners are available free of charge. Women with a low level of education were relatively underrepresented.⁴¹ As our results show, especially absenteeism related to MRSs is associated with a low level of education, and this might have biased our results. We expect women with lower educational levels to do more physical jobs or jobs with less flexibility. Therefore, our findings on work productivity loss might be underestimated. On the other hand, our finding could be overestimated due to the possibility that women with more MRSs might be more likely to respond to a questionnaire, as it may seem more relevant to them. Moreover, we were not able to provide data on presenteeism not related to MRSs nor were we able to correct for comorbid health conditions. Thus, these results must be interpreted with caution. Due to the way that the questionnaire was distributed through social media, there may have been some selection bias. However, a recent

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review stated that Facebook is a useful recruitment tool for healthcare research.42 Although we did not use a validated guestionnaire, our most important outcomes were based on questions derived from the PCQ, which itself is based on validated questions and which is recommended by guidelines for cost research.²⁴ Self-reported absenteeism generally shows a good correlation with official records, although accuracy decreases with increasing recall period.⁴³ This might have initiated a recall bias in our cohort. It is unknown to what extend recall bias affects reports on presenteeism.⁴⁴ In general, although results vary among studies on premenstrual complaints, a prospective collection of data on symptoms is advisable.^{45 46} Furthermore, an extrapolation of a three and six months timeframe to a yearly basis may intrinsically imply some degree of uncertainty, for example when the influenza season is not included in the original analysis. Finally, these results may not be generalized internationally due to variability in the regulation of social services in different countries, and this is also a limitation of our study. In The Netherlands, wages are paid during sick leave that has duration of less than 1 year, but women in other countries may not have this benefit. Since we know that many factors influence menstrual symptoms, including biological, cultural, and environmental factors, these differences might well influence both absenteeism and presenteeism.⁶¹⁴⁴⁷

In conclusion, we have found that the impact of MRSs on work and school productivity is considerable and that presenteeism contributes significantly more to the matter than absenteeism. Future research should identify how women affected by MRSs could be helped best and how their productivity can be improved in order to reduce the societal impact regarding absenteeism and presenteeism.

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618 Competing interest statement

619 All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi disclosure.pdf and declare: no support from any organisation for 620 621 the submitted work; no financial relationships with any organisations that might have an interest in the submitted work in the previous three years; no other relationships or 622 activities that could appear to have influenced the submitted work. 623

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625 Contributorship statement

TN, BD and JA conceived the study. MS wrote the first and successive drafts of the manuscript. MS, TN and EA modelled and analysed the data. TN, EA, JM, BD and JA contributed to study conception and design. MS and TN collected the data. All authors revised the manuscript for important intellectual content. MS and TN had full access to the data and take responsibility for the integrity of the data and the accuracy of the data analysis. TN is the guarantor.

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633 Data sharing statement

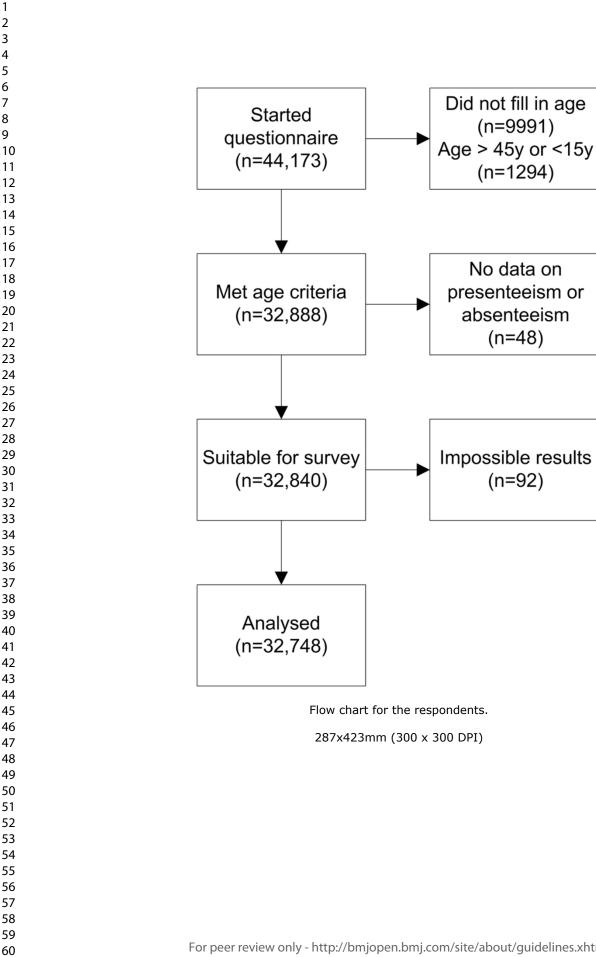
634 No additional data from this study are available from a repository. Data are available
 635 on request from the corresponding author.

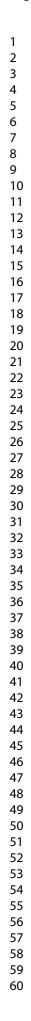
636 Acknowledgements

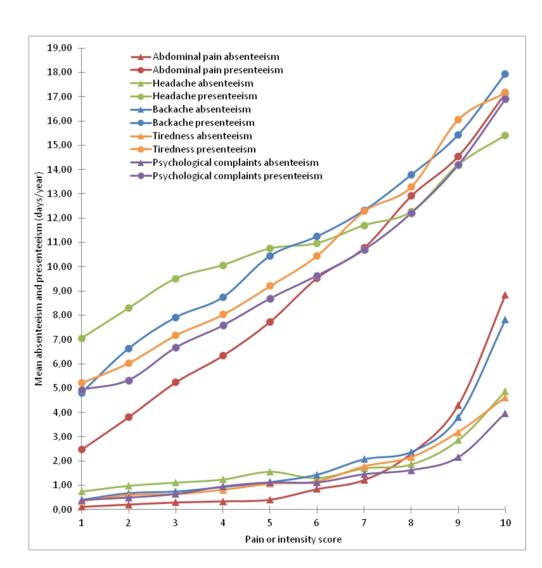
The authors would like to thank Reinoud Oomen, Peter de Vroed, Steven Giesbers,
Elsbeth Teeling, Paula Kragten and Annemarie Haverkamp for their valuable
contributions in the development and dispersion of the questionnaire.

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641	Legends to figures
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642	Figure 1
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644	Flow chart for the respondents.
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646	Figure 2
647	The relationship between pain and intensity scores, related to absenteeism and
647	The relationship between pain and intensity scores, related to absenteeism and
619	presenteeism, in lost days per year.
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CE 1	The relationship between age and everage abcenteeiem and presenteeiem
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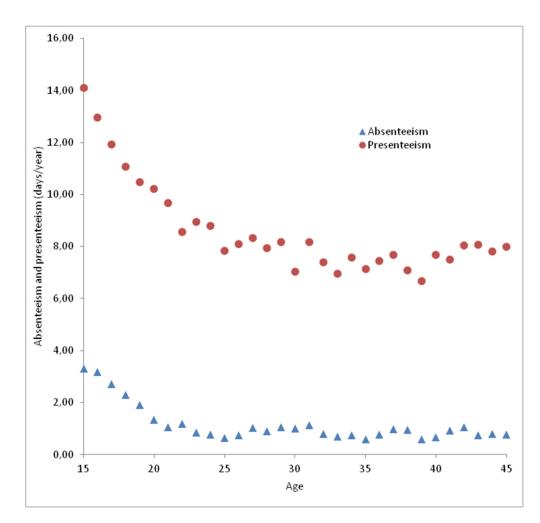






The relationship between pain and intensity scores, related to absenteeism and presenteeism, in lost days per year.

202x209mm (300 x 300 DPI)



The relationship between age and average absenteeism and presenteeism.

206x199mm (300 x 300 DPI)

Work, voluntary work and study: hours per week, sector, experienced stress

Menstrual cycle: total length, number of days with blood loss, number of times

Symptoms: abdominal pain, heavy menstrual blood loss, headache, fatigue,

backache, nausea and vomiting, tender breasts, problems with stool, psychological

Per symptom: number of days the symptom was present; number of days that the

symptom influenced daily functioning; the magnitude of the symptom's influence on

daily functioning; pain or intensity score (not for heavy menstrual bleeding, nausea

and vomiting); continuation of daily activities despite symptoms; and the extent to

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3	Appendix
4	Appendix
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9	Commence of the acception point
10	Summary of the questionnaire
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10	Part 1: Questions about basic characteristics
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20	 Age, marital status, nationality, level of education
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22	 Work, voluntary work and study: hours per week, sector,
23	. , , , , , , ,
24	- Smoking
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27	 Menstrual cycle: total length, number of days with blood
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29	needing to change pad or tampon 📈
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31 32	 Medical and obstetric history
33	intedical and obstetric history
34	Contraction
35	- Contraception use
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38	Part 2: General questions about menstruation-related symptoms
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41	 Symptoms: abdominal pain, heavy menstrual blood loss,
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43	backache, nausea and vomiting, tender breasts, problems
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48	Der symptom: number of days the symptom was present
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55	and vomiting); continuation of daily activities despite sym
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57	which woman forced themselves to continue their deily of
58 50	which women forced themselves to continue their daily a
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00	 Usage of over-the-counter drugs, narcotics, painkillers, al

Usage of over-the-counter drugs, narcotics, painkillers, alternative medicine

which women forced themselves to continue their daily activities

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Part 3: Questions related to absenteeism and presenteeism (full text)

- How often did you call in sick due to menstrual complaints in the past six months? _
 - Every menstrual cycle
 - Almost every menstrual cycle
 - Half of all menstrual cycles
 - Sometimes
 - Never Ο
- What is the mean number of days you called in sick due to menstruation-related _

symptoms per cycle in the past six months?

- More than three days
- Three days Ο
- Two days Ο
- One day Ο
- ė Liez Half a day or a few hours
- What is the number of days you were present at work while being hindered by menstrual complaints?
- How was your performance during these days?
 - o A Visual Analog Scale by means of sliding a bar was used to score the amount of productivity loss, where the left end said "I could not do anything" and the right end said "I could do just as much as I normally do".

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3	- What was the number of days you called in sick because of non-menstruation-related
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6	symptoms in the past six months?
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11	 What reason do you give when you call in sick due to menstruation-related
12	
13	symptoms?
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16	 Menstrual complaints
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18	 Only the presenting symptom, for example abdominal pain or headache
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20	
21	 You do not give a reason
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23	 You think up another reason
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28	- What measures would be desirable for women on the workplace or at school during
28	
30	their menstrual period? It is possible to give more than one answer.
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32	 A day off without any consequences
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35	 A day off to catch up later
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37	 Flexibility to perform less physical work
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39	 More time for personal care
40	 More time for personal care
41	 The possibility to work from home
42	 The possibility to work from home
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STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of cross-sectional studies

Section/Topic	ltem #	Recommendation	Reported on page #
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	2
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	4
Objectives	3	State specific objectives, including any prespecified hypotheses	5
Methods			
Study design	4	Present key elements of study design early in the paper	5
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	5-7
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants	6
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	8, appendix
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	7-8
Bias	9	Describe any efforts to address potential sources of bias	7-9
Study size	10	Explain how the study size was arrived at	7
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	7-9
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	7-8
		(b) Describe any methods used to examine subgroups and interactions	7-8
		(c) Explain how missing data were addressed	7,9
		(d) If applicable, describe analytical methods taking account of sampling strategy	n/a
		(e) Describe any sensitivity analyses	n/a
Results			

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Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility,	9
		confirmed eligible, included in the study, completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	9/Figure 1
		(c) Consider use of a flow diagram	Figure 1
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	9/Table 1
		(b) Indicate number of participants with missing data for each variable of interest	Not reported, can be reported if desirable.
Outcome data	15*	Report numbers of outcome events or summary measures	9-11/Table 2,3
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence	Table 4
		interval). Make clear which confounders were adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	Table 1,4
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	n/a
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	n/a
Discussion		6	
Key results	18	Summarise key results with reference to study objectives	11-12
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	14-15
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	12-15
Generalisability	21	Discuss the generalisability (external validity) of the study results	14-15
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	n/a

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.