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## 1 Obesity and Overweight Associated with Lower Rates of Colorectal

## 2 Cancer Screening in Switzerland

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#### 27 Abstract

Background: Screening for colorectal cancer (CRC) is associated with reduced CRC mortality,
but low screening rates have been reported in several settings.

30 **Objective:** To assess predictors of low CRC screening in Switzerland.

31 Design & Participants: Retrospective cohort of a random sample of 940 patients aged 50–80

32 years followed for 2 years from 4 Swiss University primary care settings. Patients with illegal

33 residency status and history of CRC or colorectal polyps were excluded.

34 Main measures: We abstracted socio-demographic data of patients and physicians, patient

35 health status and indicators derived from RAND's Quality Assessment Tools from medical

36 charts. We defined CRC screening as colonoscopy in last 10 years, flexible sigmoidoscopy in last

37 5 years, or fecal occult blood testing in last 2 years. We used bivariate and multivariate logistic

38 regression analyses.

39 Key results: Of 940 patients (mean age 63.9 years, 42.7% women), 316 (33.6%) had CRC

40 screening. In multivariate analysis, birthplace in a country outside of Western Europe and North

41 America (OR 0.65, 95% CI 0.45-0.97), male gender of the physician in charge (OR 0.67, 95% CI

42 0.50-0.91), BMI 25.0-29.9 kg/m<sup>2</sup> (OR 0.66, CI 0.46-0.96) and  $\geq$  30.0 kg/m<sup>2</sup> (OR 0.61, CI 0.40-

43 0.90) were associated with lower CRC screening rates.

44 Conclusion: Obesity, overweight, birthplace outside of Western Europe and North America, and
45 male gender of the physician in charge were associated with lower CRC screening rates in Swiss
46 University primary care settings. Physician perception of obesity and its impact on their
47 recommendation for CRC screening might be a target for further research.

48 Keywords: Colorectal Cancer; Cancer Screening; Obesity

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#### 49 Background

50 According to the World Health Organization (WHO), colorectal cancer (CRC) is the fourth 51 leading cause of cancer death worldwide. CRC screening using biennial fecal occult blood 52 testing has been shown to reduce CRC related mortality by about 14% [1]. Screening colonoscopy every 10 years, flexible sigmoidoscopy every 5 years, or annual fecal occult blood 53 54 testing (FOBT) have been recommended by United States (U.S.) Guidelines [2] and a European 55 Panel [3] for patients older than 50. Several studies also support cost effectiveness of CRC 56 screening [4,5]. However, reported screening rates stay around 55% in the U.S. [6,7] and around 57 45% in European countries [8]. 58 In search for a better understanding of these low rates, recent studies examined the factors 59 associated with CRC screening. Socio-economic factors such as white ethnicity, male gender, 60 lower age, higher education level and higher household income, and insurance coverage, were 61 found to be associated with higher CRC screening rates in several studies in the U.S. and Canada 62 [9–12]. Whereas presence of a chronic condition increased odds for CRC screening in one study 63 [11], no association with the number of chronic conditions was found in another [12]. A recent 64 review on the relationship between obesity and cancer screening found a complex pattern of 65 positive and negative associations between CRC screening and weight status throughout the 66 literature [13]. Several studies found that annual influenza vaccine was associated with higher 67 CRC screening rates [9,11]. The main reasons not to perform FOBT screening were patient

68 unawareness and physicians not recommending it [10].

69 Outside of Northern America, publications on correlates of CRC screening are rare. A recent

70 Spanish study found higher education level, periodic screening for breast and cervical cancer in

vomen, and knowledge about CRC and CRC screening to increase odds for initial participation

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72	in a population-based screening program [14]. One study from Hong Kong found that physician
73	factors such as experience, academic appointment and agreement with CRC screening were
74	associated with higher screening rates [15]. However, we found no studies assessing the factors
75	associated with CRC screening in European countries without population-based screening
76	program, such as Switzerland.
77	The aim of this study was to assess patient and physician factors associated with lower rate of

78 CRC screening, in University primary care settings in Switzerland, a country without a

79 population-based screening program.

#### 80 Methods

#### 81 Study design and patients

82 The study design was previously published [16]. Briefly, this retrospective cohort study aimed to 83 assess the quality of preventive care and control of cardiovascular risk factors, and included a 84 random sample of all patients aged 50 to 80 years old, followed up by primary care physicians 85 (PCPs) in 4 Swiss University primary care settings in Basel, Geneva, Lausanne and Zurich 86 during 2005 and 2006. We limited the sample to this age group in order to have a high enough prevalence of examined preventive indicators and eligibility for cancer screening. From the 87 88 initial random sample of 1889 patients, we excluded patients whose medical charts could not be 89 found (54), patients with emergency visits or nurse appointment only (125), those followed up in 90 specialized clinics only (117) or for less than one year (591), because of a possible lack of 91 adequate time to provide preventive care (Appendix Figure 1). For this study, we also excluded 92 12 patients known for CRC or colorectal polyps at the beginning of the review period, as the 93 focus was to assess cancer screening in average risk patients and not to follow up. In addition, 94 we excluded 50 patients with irregular residency status, as insurance coverage is not guaranteed 95 for these patients. The final sample consisted of 940 abstracted medical charts. A similar sample 96 size was used in previous studies on quality of care based on chart abstraction [17,18]. The 97 Institutional Review Board approved the study protocol at each site.

#### 98 Chart abstraction

99 Based on previous studies on factors associated with CRC screening [9,10,15], we examined the 100 following indicators: patient demographic characteristics, including age, gender, civil status, 101 birthplace and occupation; characteristics of the physician in charge (defined as the physician 102 with whom the patient had the most visits over the review period), including gender, position, and number of visits. BMI was calculated as reported body-weight in kilograms divided by
squared height in meters, or directly abstracted from the medical chart. As a surrogate to patient
health status, we calculated the Charlson combined age-comorbidity index [19,20] from reported
comorbid conditions and the patient's age. As this index only captures a limited list of conditions
that have a potential impact on survival, we abstracted the number of prescribed medications
over the two years review period.

109 As previously described [16], patient data were abstracted from medical charts by trained 110 medical students using a questionnaire form similar to previous studies [17,18,21]. The 111 questionnaire assessed the 33 selected indicators for chronic and preventive care derived from 112 RAND's Quality Assessment Tool System [21], as well as other covariates (demographics, co-113 morbid conditions) based on the chart abstraction form from the Translating Research Into 114 Action For Diabetes Study [22]. Performed CRC screening was defined as documented 115 colonoscopy in the last 10 years, flexible sigmoidoscopy or double contrast barium enema in the last 5 years, or fecal occult blood testing in the last 2 years, based on the U.S. Preventive 116 117 Services Task Force 2002 recommendation statement [23]. We considered any procedure or test 118 regardless of their indication, as data on indications was not always available. We used the 2002 119 recommendation statement instead of the 2008 statement [2] to measure compliance with 120 guidelines at the time of the review period (2005–2006). Patients who refused CRC screening 121 (n=22) were grouped with those not screened, even though screening was offered by the 122 physician, as this information might not be systematically documented in medical charts and 123 therefore not adequately captured by chart. To measure physician-initiated care, we repeated the 124 analysis categorizing these patients with the performed CRC screening group in a secondary 125 analysis.

126 As chart abstraction might underestimate quality of care by 5% to 10% compared to clinical vignettes and standardized patients [24], inter-rater reliability was assessed by repeating the chart 127 128 abstraction on a random sample of patients (n=45) to detect a significant kappa value [16,25]. 129 For CRC screening, the kappa value was 0.85. Influenza immunization indicators were validated 130 with an external administrative register at Lausanne, as previously reported [16]. For 230 patients, no BMI value could be abstracted or calculated from medical charts; we imputed these 131 missing values by the Gaussian normal regression imputation method from known values of 132 133 patient's age, gender, weight and height. We report median, lower and upper quartile of BMI 134 before and after imputation, and proportion of imputed BMI values for each BMI category. We 135 categorized age of the patients in three life decades, and categorized birthplace as Switzerland or 136 foreign country, further divided into Western European and North American countries, defined as 137 country in the child and adult mortality stratum A according to the Annex 1 of the WHO 2003 138 World Health Report [26], and outside of Western Europe and North America, mortality strata B 139 - E. We dichotomized continuous variables without normal distribution, i.e. Charlson combined 140 age-comorbidity index, the number of prescribed medications, and the number of visits to the 141 physician in charge, according to the median, and report interquartile ranges (IQR).

#### 142 Statistical analyses

We used descriptive statistics and bivariate analyses to characterize the sample, and performed multivariate logistical regression to analyze the predictors of adequate CRC screening. Results are reported as percentages with 95% binomial exact confidence intervals (CI). To account for clustering within each of the four University setting, we treated each site as a fixed effect. Multivariate logistic regression analysis was repeated in sensitivity analyses using measured BMI without imputation, and number of prescribed medications in addition to Charlson index.

- 149 We used Stata software (version 12.1, Stata Corp., College Station, Texas) for all statistical
- 150 analyses.

#### 151 **Results**

#### 152 **Patient description**

- 153 Table 1 shows characteristics of the 940 included patients. Mean age was 63.9 years (standard
- deviation 8.8), and 402 patients (42.7%) were women. Median BMI was 27.9 (lower quartile:
- 155 24.9, upper quartile: 31.2) before imputation and 28.2 kg/m<sup>2</sup> (IQR 24.8 32.1) after imputation.
- 156 Most of the care was delivered by residents (89.5% vs. 10.5% by senior residents or attending
- 157 physicians) and 55.1% of patients were followed by female physicians. Annual influenza
- 158 vaccination was delivered to 22.3% of patients, whereas 77.7% did not receive vaccination.

#### 159 Colorectal cancer screening rates

160 Table 2 shows CRC screening rates and describes screening methods. Only 316 (33.6%) of

161 patients eligible for CRC screening had adequate CRC screening. Refusal of CRC screening was

162 documented in 22 patients (2.3%). Fifty six percent had a colonoscopy, 41.3% had FOBT alone

- 163 or combined with endoscopy, and only 3.6% had flexible sigmoidoscopy alone. Double contrast
- 164 barium enema was not used for any patient.
- 165 Results of bivariate and multivariate logistical regression analyses are reported in Table 3. In
- bivariate analyses, being overweight (BMI 25.0 to 29.9 kg/m<sup>2</sup>) and obese (BMI  $\geq$  30.0 kg/m<sup>2</sup>)
- 167 were associated with lower CRC screening rates (OR 0.64, CI 0.46 0.90, and OR 0.61, CI 0.43
- 168 0.88, respectively; p for trend = 0.007). Male gender of the physician in charge (OR 0.70, CI
- $169 \quad 0.53 0.93$ ) was also negatively associated with CRC screening. Place of birth in a country
- 170 outside of Western Europe and North America was only borderline significant (OR 0.73, 95% CI
- 171 0.53 1.00).
- 172 In multivariate logistic regression analysis, being overweight (OR 0.66, CI 0.46 0.96) or obese
- 173 (OR 0.61, CI 0.40 0.90), and male gender of the physician in charge (OR 0.67, 95% CI 0.50 –
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0.91; p for trend = 0.011) remained significantly associated with lower screening rates, and place
of birth outside of Western Europe and North America became significant (OR 0.65, 95% CI
0.45 - 0.97).

177 The proportion of imputed BMI values was considerably higher in normal and overweight than

178 in obese patients (23% and 34% vs. 13% respectively). A sensitivity analysis using measured

179 BMI without imputation confirmed the association of CRC screening with obesity (OR 0.63,

180 95% CI 0.51 - 0.99) and birthplace in a country outside of Western Europe and North America,

181 but not with overweight and physician gender. Including the total number of medications in

182 addition to Charlson combined age-comorbidity index did not significantly change the results. In

183 a secondary analysis considering CRC screening proposed by the physician rather than

184 performed screening, being overweight and obese (OR 0.65, 95% CI 0.45 – 0.93 and OR 0.62,

185 95% CI 0.42 - 0.91), and male gender of the physician in charge (OR 0.67, 95% CI 0.50 - 0.91)

186 were associated with CRC screening, but not with place of birth in a country outside of Western

187 Europe and North America.

188

### 189 **Discussion**

190 We found that only about one in three patients followed up in Swiss University primary care 191 settings were screened for CRC. This relatively low CRC screening rate is comparable to those 192 reported in some underserved populations in the U.S. [27]. We found three characteristics to be 193 associated with lower CRC screening rates: being overweight or obese, male gender of the 194 physician in charge, and being born in a country outside of Western Europe and North America. 195 None of the other measured socio-demographic factors (patient gender, occupation, civil status), 196 nor patient health status estimated by the Charlson combined age-comorbidity index reached 197 statistical significance.

198 As previously hypothesized [16], the absence of a population-based CRC screening program, or 199 the lack of systematic performance monitoring in Switzerland, might explain these low CRC 200 screening rates. This hypothesis is also supported by findings of a Canadian study, where higher 201 CRC screening rates were found in regions with recent implementation of a CRC screening program [11]. Also, influenza vaccination has been shown to be associated with higher CRC 202 203 screening rates [9,11], but was not significant in our study, possibly due to the small sample size. 204 The negative association between BMI and CRC screening has been reported in previous studies 205 [28], but a recent review article by Fagan *et al.* showed a mixed pattern of association between 206 weight status and CRC screening without a clear trend towards a positive or negative association 207 throughout the literature [13]. The authors hypothesized that factors associated with CRC 208 screening might differ according to race and gender of the studied population. Overweight 209 persons might be less health conscious and therefore less demanding of health care. However, 210 previous studies have shown that lack of recommendation by the physician is the most frequent 211 reason to skip CRC screening, instead of patients not asking or refusing it [10]. Also, physicians

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might have been less likely to recommend CRC screening to overweight and obese patients,
either because they think that these patients would not want screening or because of higher
priority on other medical conditions. However, in the latter case, one would expect the same to
be true for patients with higher Charlson combined age-comorbidity index. This potential
stigmatization of overweight and obese patients by their physicians has been documented
[29,30].

Female gender of the physician in charge has been associated with higher rates of screening mammographies and cervical smears [31]. However, physician gender was not associated with CRC screening in previous studies [15]. Female physicians might be more sensitized to cancer prevention through their participation as patients in regional, population-based breast cancer screening programs in Switzerland, and well-implemented screening for cervical cancer, whereas no such program exists for cancer in men.

224 Our finding that CRC screening is associated with birthplace outside of Western Europe and 225 North America is consistent with previous studies that found ethnicity to be associated with CRC 226 screening [9–12]. These studies also found other socio-demographic factors such as gender and 227 educational level, to be predictors of CRC screening that did not reach statistical significance in 228 our results. This might be due to the relatively small sample size. A recent review by Naylor et 229 *al.* suggests that patient education and physician training in communication with patients of low health literacy are paths for improving adherence to CRC screening in ethnic minorities [32]. 230 231 Our study has several limitations. A first limitation was the potential underreporting in medical 232 charts of data such as patient refusal of screening. Process-based quality scores using abstraction 233 of medical charts have been found to underestimate the quality of care compared to clinical 234 vignettes and standardized patients by 5% and 10%, respectively [24]. Second, we considered

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235 any initial sigmoidoscopy in the last 5 years or colonoscopy in the last 10 years as CRC 236 screening, because of lack of complete data on the indication, and excluded follow-up 237 colonoscopies for CRC or adenomatous polyps only. We therefore cannot exclude that some of 238 these endoscopic procedures were conducted for symptomatic indications, such as rectal 239 bleeding. Third, our results rely on imputed BMI and sensitivity analysis confirmed the 240 association between CRC screening and obesity only, but not for overweight and physician gender. This might be due to loss of power in the statistical model (n=688) and potential 241 242 underreporting of BMI in normal weight and overweight, but not for obese patients. Fourth, our 243 sample in University primary care settings might not be representative of the general Swiss 244 population. In fact, our sample had a high proportion of patients with migratory background 245 (51%) and men (57%), and a higher prevalence of obesity than in the general Swiss population 246 (32% in our sample vs. 13% in the general Swiss population aged 55-74 years) [33]. Our model was adjusted for these characteristics. Fifth, most of the care was delivered by residents in 247 248 postgraduate training. Similar to our study, a previous report on CRC screening showed no 249 difference in screening rates between physicians with or without completed postgraduate training 250 [15], but we did not find studies directly comparing screening rates between community-based 251 PCPs and University-based residents within a health care system with universal coverage. Sixth, 252 it was not possible to measure patient and physician knowledge and beliefs regarding CRC and 253 CRC screening, as this study was based on a retrospective review of medical charts. Seventh, due 254 to the relatively small sample size, the absence of statistical significance has to be interpreted 255 with caution, as the study might not have enough power to detect associations in subgroups. 256 These results may be important for clinical practice, as obesity has been suggested as a risk 257 factor for colorectal cancer [34,35]. Physician perception of obesity and its impact on their

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- 258 recommendation for CRC screening might be a target for further research. A population-based
- 259 CRC screening program and a systematic performance monitoring might help improve the rate
- 260 of CRC screening.

261

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- 266 Swiss Society of General Internal Medicine 79<sup>th</sup> Annual Conference in Lausanne, Switzerland,
- 267 under the title of "Obesity and overweight are negative predictors of colorectal cancer screening
- 268 in a health care system combining fee for service and universal coverage".

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## Tables

			<b>N</b> X 0.40
Table 1: Patient charac	cteristics		N=940
Age (years, n, %)			
50 – 59 years		345	36.7%
60 – 69 years		359	38.2%
70 – 80 years		236	25.1%
Gender (n, %)			
female		402	42.7%
Civil Status (N=931, n, %)			
married		490	52.6%
divorced or separated		214	23.0%
single		133	14.3%
widowed		94	10.1%
Birth place (N=930, n, %)			
Switzerland		453	48.7%
Other country, within We	estern Europe and North America*	194	20.9%
Other country, outside of	Western Europe and North America*	283	30.4%
Occupation (N=921, n, %)	1		
employed		251	27 3%
retired		364	39.5%
freelance at home or in a	education	112	12.2%
social aid		109	11.8%
unemployed		85	9.2%
Body mass index* (n %)		05	9.270
$< 18.5 \text{ kg/m}^2$		14	1 5%
$18.5 - 24.9 \text{ kg/m}^2$		230	24 5%
$25.0 - 29.9 \text{ kg/m}^2$	391	42.6%	
$>30.0 \text{ kg/m}^2$		305	32 4%
<u>Charlson combined age-con</u>	norhidity index	505	52.470
median interquartile rand	3	2 - 5	
Number of prescribed med	5	2-5	
median interquartile rand	1	26	
Physician in charge*		т Т	2-0
$n_{\rm N} = 0.21 n_{\rm N}^{-0.21}$	resident	833	80 50/
position (N=931, II, 70)	senior resident	833 78	89.570 8 10/-
	attending / feaulty	78	0.470 0.10/
ander(N=020, n, 9/)	formals	20	2.170 55 10/
gender (N-939, II, 70)	interguartile range)	517	33.170
number of visits (median, interquartile range)			4-9
Annual Influenza vaccinati	0n (n, %)	210	22.20/
uone		210	22.5%
not done		730	11.1%

\* Defined as countries in the child and adult mortality stratum A based on World Health Organization (WHO) mortality estimates [26].

<sup>†</sup> Two hundred and thirty missing values were imputed from patient age, gender, height and weight. The median body mass index (BMI) was 28.2 kg/m<sup>2</sup> (interquartile range 24.8 – 32.1) before imputation and 27.9 kg/m<sup>2</sup> (interquartile range 24.9 - 31.3) after imputation.

<sup>‡</sup> Physician in charge is defined as the physician the patient had the most visits to.

Table 2: Colo-Rectal Cancer (CRC) screening				
CRC screening (n, %)				
not done *	624	66.4%		
done	316	33.6%		
Screening method (N=315, n, %)				
fecal occult blood test	130	41.3%		
colonoscopy†	173	54.9%		
flexible sigmoidoscopy:	12	3.8%		
double contrast barium enema	0	0%		

\* Including 22 (2%) patients who had refused CRC screening.

In 47 patients with FOBT or sigmoidoscopy and subsequent colonoscopy, the latter was considered follow-up, not screening.

‡ Nine patients screened with FOBT and sigmoidoscopy grouped with FOBT.

	Bivariate OR (95% CI)		Multivariable adjusted OR§ (95% CI)	
Age		( )		,
50 - 59 years	1 (ref.)		1 (ref.)	
60 - 69 years	1.45	(1.06 - 1.99)	1.29	(0.87 - 1.88)
70 - 80 years	1.18	(0.82 - 1.68)	0.90	(0.55 - 1.57)
Gender		(0.02 0.000)		(0000 000))
female	1 (ref.)		1 (ref.)	
male	1.04	(0.79 - 1.37)	1.01	(0.72 - 1.41)
Civil status		× ,		
married	1 (ref.)		1 (ref.)	
divorced or separated	1.02	(0.73 - 1.44)	0.94	(0.66 - 1.37)
widowed	1.34	(0.85 - 2.11)	1.19	(0.72 - 1.95)
single	1.05	(0.70 - 1.58)	0.94	(0.60 - 1.47)
Birth place				
Switzerland	1 (ref.)		1 (ref.)	
Other country, within Western Europe and	1			
North America*	1.02	(0.72 - 1.46)	1.02	(0.70 - 1.49)
Other country, outside of Western Europe				
and North America*	0.73	(0.53 - 1.00)	0.65	(0.45 - 0.97)
Occupation				
employed	1 (ref.)		1 (ref.)	
retired	1.28	(0.91 - 1.79)	1.32	(0.83 - 2.09)
freelance, at home or in education	0.78	(0.48 - 1.28)	1.00	(0.55 - 1.81)
social aid	1.10	(0.68 - 1.77)	1.10	(0.65 - 1.86)
unemployed	1.11	(0.66 - 1.86)	1.13	(0.62 - 2.06)
Body mass index <sup>D</sup>				
$<18.5 \text{ kg/m}^2$	0.56	(0.17 - 1.83)	0.32	(0.09 - 1.17)
$18.5 - 24.9 \text{ kg/m}^2$	1 (ref.)		1 (ref.)	
$25 - 29.9 \text{ kg/m}^2$	0.64	(0.46 - 0.90)	0.66	(0.46 - 0.96)
$\geq 30 \text{ kg/m}^2$	0.61	(0.43 - 0.88)	0.61	(0.40 - 0.90)
Charlson combined age-comorbidity index	[			
$\leq$ 3 points	1 (ref.)		1 (ref.)	
> 3 points	0.85	(0.65 - 1.11)	0.80	(0.58 - 1.11)
Physician in charge‡				
position resident	1 (ref.)		1 (ref.)	
senior resident	1.05	(0.65 - 1.71)	1.25	(0.74 - 2.11)
attending /faculty	1.32	(0.53 - 3.28)	1.33	(0.50 - 3.57)
gender female	1 (ref.)		1 (ref.)	
male	0.70	(0.53 - 0.93)	0.67	(0.50 - 0.91)
$\leq 6$ visits during 2 years follow-up	1 (ref.)			
> 6 visits during 2 years follow-up	1.18	(0.90 – 1.55)	1.15	(0.86 - 1.55)
Annual Influenza vaccination	1 ( 0)		1 ( 0)	
done	1 (ref.)	(0, 4, 1, 0, 1)	1 (ref.)	(0, 50, 1, 20)
not done	0.74	(0.4 - 1.01)	0.84	(0.58 - 1.20)

#### Table 3: Factors associated with colorectal cancer screening

\* Defined as countries in the child and adult mortality stratum A based on World Health Organization (WHO) mortality estimates [26].

<sup>†</sup> Two hundred and thirty missing values were imputed from patient age, gender, height and weight.

\* Physician in charge is defined as the physician the patient had the most visits to.

§ Adjusted for age, gender, civil status, occupation, birth place, BMI, Charlson combined age-comorbidity index, annual influenza vaccination, number of visits to the physician in charge, and function, gender and age of the physician in charge, using a multilevel generalized linear model and treating each site as a random effect. N=897 due to missing values.

## Appendix

Appendix Figure 1: Flowchart from the electronic administrative data to the final sample