

SLOVENIA

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VHPB BALKAN MEETING

*CURRENT SITUATION:
EPIDEMIOLOGY, BURDEN OF
DISEASE, SCREENING &
PREVENTION, CASCADE OF
CARE*



DECLARATION OF CONFLICT OF INTEREST

- Lecturer: Abbvie, Bayer, Gilead, Merck, Sandoz
- Manuscript preparation: Abbvie, Gilead, Merck
- Travel/accommodational meeting expences: Abbvie, Gilead, Merck

No conflict of interest regarding this presentation

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EPIDEMIOLOGY - HBV

PREVALENCE	HBsAg (year)
General population	est. 0.8% ? (2019)
Blood donors (first time)	0.004 % (2018-2020); N=250 640
Pregnant women	NA (a national study in progress)
Risk Groups	
PWID	2.5% (2003)
MSM	0.25% (2019-2022); N=1 600
Prisoners	NA
Other	-
Healthcare workers	NA

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EPIDEMIOLOGY - HBV

INCIDENCE	Acute: HBsAg (most common age group)	Chronic: HBsAg (most common age group)
General population	2019: 0.4/100 000 (65-74 y) 2020: 0.1/100 000 (55-64 y) NO INFANTS* (since 1996)	2019: 2.5/100 000 (55-64 y) 2020: 1.2/100 000 (45-54 y) NO INFANTS* (since 1996)

*born to parents with Slovenian origin

WHO 2021	
Elimination of chronic HBV infection as a public health problem	
Incidence 95% reduction	Mortality 65% reduction
HBV EMTCT ≤0.1% HBsAg prevalence in ≤5 year olds ^{a,b}	Annual mortality ^a (HBV) ≤4/100 000
Additional target: ≤2% MICT rate (where use of targeted HepB-BD) ^c	

Number of annually reported cases (cases /100 000) of acute and chronic HBV infection in Slovenia, period 2016-2020					
LETO	2016	2017	2018	2019	2020
	št. primerov (št.primero / 100.000)				
akutni hepatitis B	18 (0,9)	15 (0,7)	10 (0,5)	9 (0,4)	2 (0,1)
kronični hepatitis B	22 (1,1)	37 (1,8)	45 (2,2)	52 (2,5)	26 (1,2)



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EPIDEMIOLOGY - HCV

HCV-seroprevalence by selected groups, period 1992-2015

Population	HCV RNA + (%)	N	Year
Haemophiliacs	26.7%	374	1992
Blood donors	0.025%	1,4 mill	1993-2000
Haemodialysis	1.1%	1343	1995
PWID on OAT	16.5%	1,050	2007
Prison inmates	26%	378	2006
Pregnant women	0.08%	9,574	2013

Maticic M. JEADV 2008; Kastelic A et al. 2nd Slovenian Conference on HCV Infection in IVDU, March 2007; Skamperle M et al. Acta Dermatoven APA 2014; 23: 25-6.

EMCDDA. <http://www.emcdda.europa.eu/publications/country-overviews/sj>; Kopilovic B et al. Euro Surveill 2015; 20(22): 21144. Seme K, et al. Nephron. 1997;77(3):273-8.

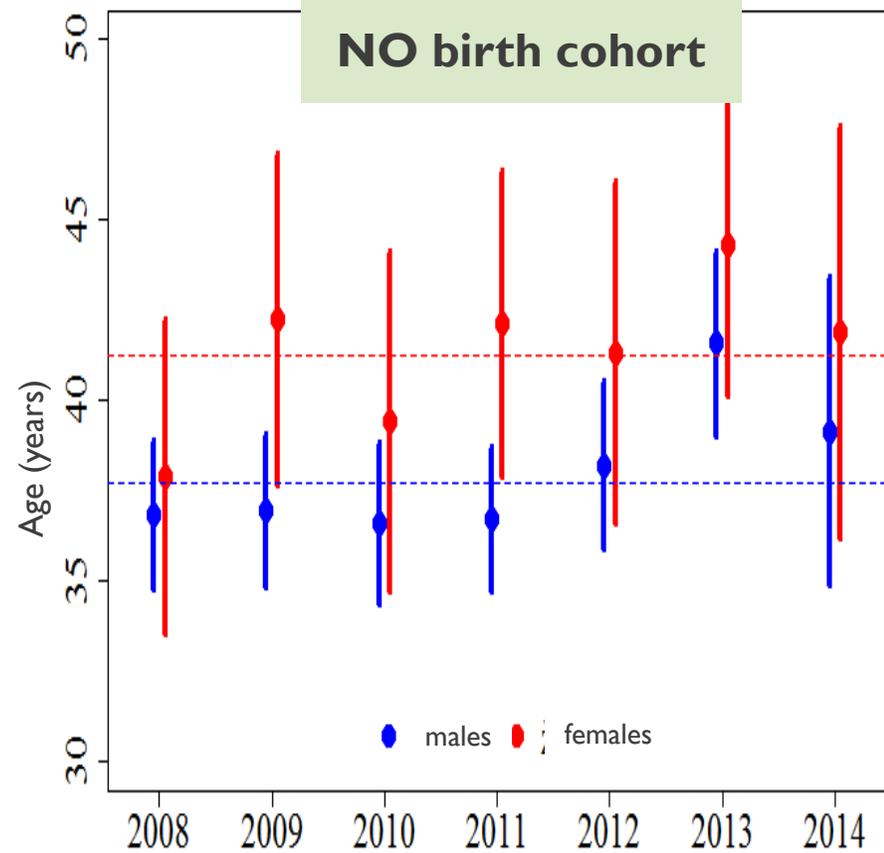
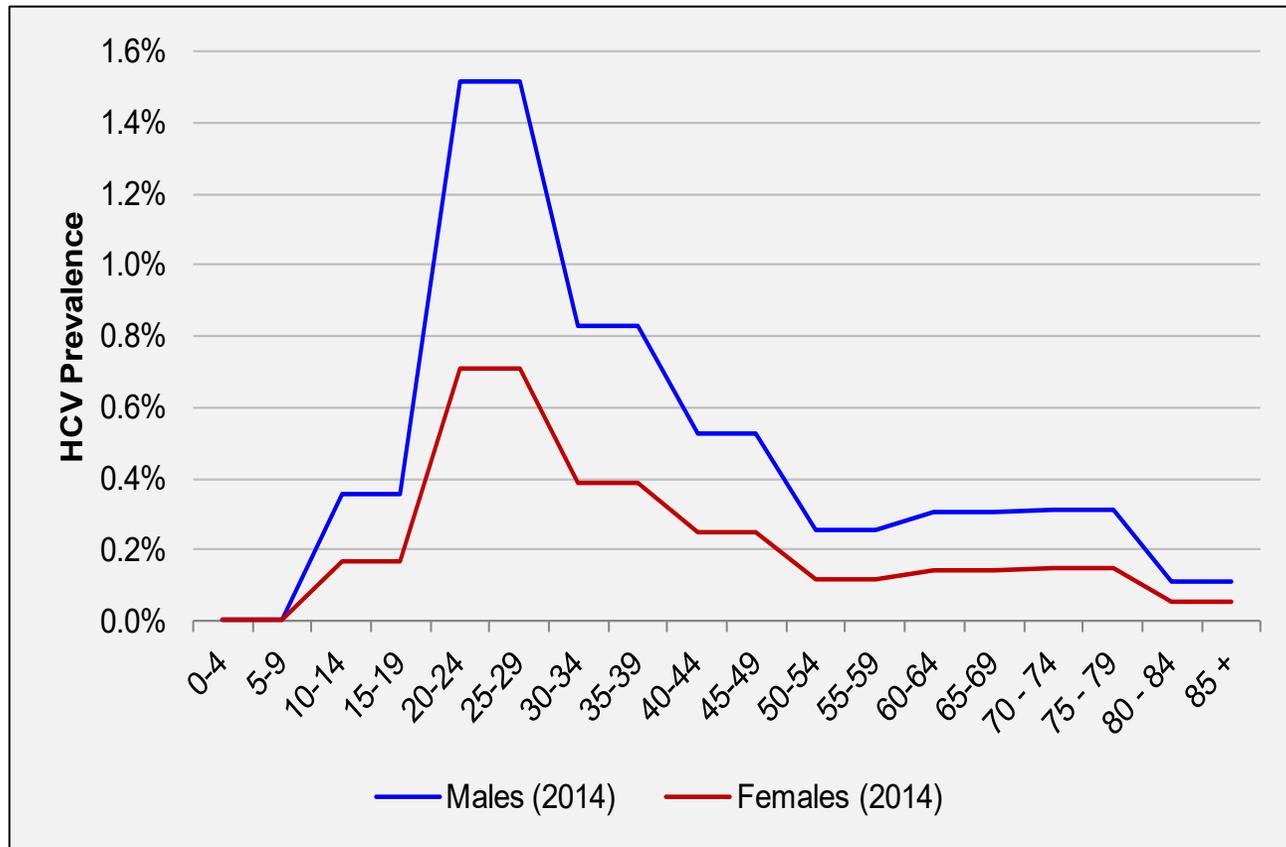
Gregorčič S et al. ECCMID 2018. Madrid, April 23, 2018. Poster #Mon-13. Institute for transfusion Medicine Slovenia; Annual Reports 1993-2000.



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EPIDEMIOLOGY - HCV

Anti-HCV positive individuals by age and gender, period 2008 - 2014



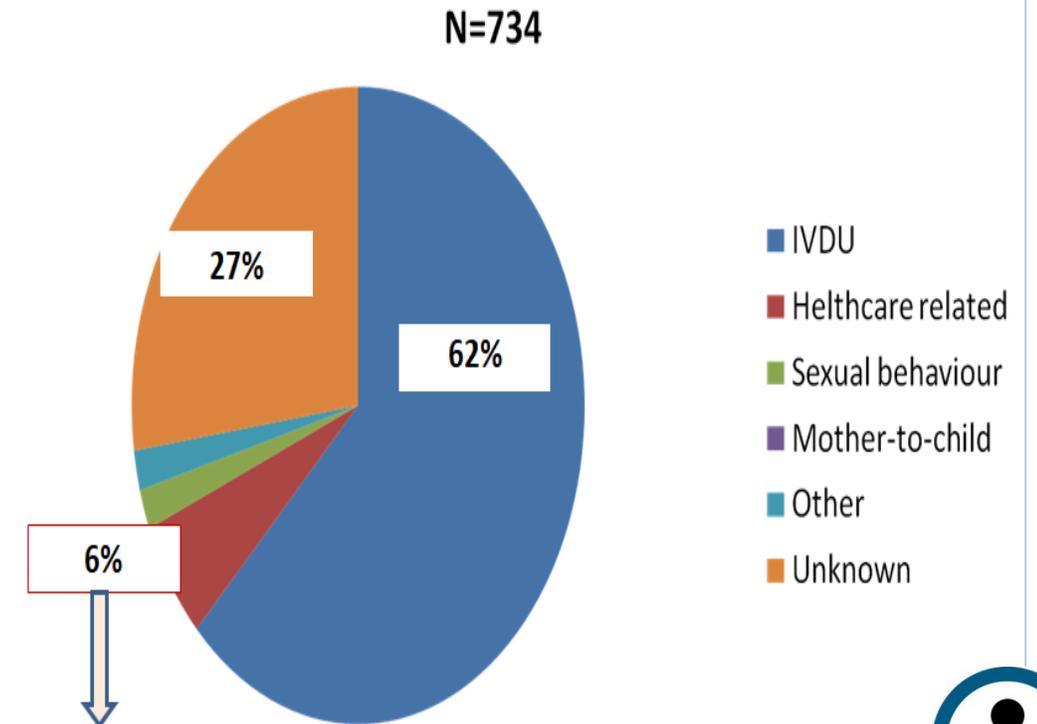
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EPIDEMIOLOGY - HCV

Characteristics of anti-HCV positive individuals in two observational periods

	Period		p-value
	1993-2007	2008-2015	
Number	2719	1398	
	4117		
Sex (n, %)			0.27
Males	1811 (66.6%)	955 (68.3%)	
Females	908 (33.4%)	443 (31.7%)	
Age at diagnosis			<0.001*
mean, SD	36.65 (15.29)	39.33 (14.40)	
Risk factors (n, %)			<0.001*
Intravenous drug use	1029 (37.8%)	818 (58.5%)	
Healthcare	300 (11.9%)	80 (5.7%)	
Sexual contacts	38 (1.4%)	29 (2.1%)	
Other	24 (0.9%)	37 (2.6%)	
Unknown	1328 (48.8%)	434 (31.1%)	

Proportion of HCV-seropositive persons by risk groups, period 2008-2014



• Healthcare related: 80% blood-transfusion recipients before 1992



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EPIDEMIOLOGY - HCV

PREVALENCE	HCV RNA (year)
General population	0.01% (2022)
Blood donors (first time)	0.000% (2018-2020); N=250 640
Pregnant women	0.08% (2013); N=9 574
Risk Groups	
PWID	16.5% (2007)
MSM	0.13% (2019-2022); N=4 783
Prisoners	26% (2006)
Other	-
Healthcare workers	0.7% (2000-2013)

Maticic M. JEADV 2008; Kastelic A et al. 2nd Slovenian Conference on HCV Infection in IVDU, March 2007; Baklan Z, et al. Wien Klin Wochenschr 2004; 116(2): 5-7.

EMCDDA. <http://www.emcdda.europa.eu/publications/country-overviews/si>; Kopilovic B et al. Euro Surveill 2015; 20(22): 21144. Gregorčič S et al. ECCMID 2018. Madrid, April 23, 2018. Poster #Mon-13.

Rahne U et al. Institute for transfusion Medicine Slovenia; Report 2018-2020. Legebitra. Annual Report on STI testing in MSM, 2019.-2022.



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EPIDEMIOLOGY - HCV

INCIDENCE	Acute: HCV RNA (most common age group)	Chronic: HCV RNA (most common age group)
General population	2019: 0.1/100 000 (25-34 y) 2020: 0.1/100 000 (35-44 y)	2019: 2.5/100 000 (35-44 y) 2020: 1.1/100 000 (35-44 y)

WHO 2021	
Elimination of chronic HCV infection as a public health problem	
Incidence 80% reduction	Mortality 65% reduction
Annual incidence (HCV) ≤5/100 000 ≤2/100 (PWID)	Annual mortality ^B (HCV) ≤2/100 000

Number of annually reported cases (cases / 100 000) of acute and chronic HCV infection in Slovenia, period 2016-2020					
LETO	2016	2017	2018	2019	2020
	št. primerov (št.primerov / 100.000)				
akutni hepatitis C	7 (0,3)	6 (0,3)	5 (0,2)	1 (0,1)	2 (0,1)
kronični hepatitis C	108 (5,2)	111 (5,4)	114 (5,5)	53 (2,5)	24 (1,1)

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EPIDEMIOLOGY - HEV

PREVALENCE	HBsA (year)	HCV RNA (year)
General population	?	0.01% (2022)
Blood donors (first time)	0.004 % (2018-2020) N=250 640	0.000% (2018-2020) N=250 640
Pregnant women	NA	0.08% (2013) N=9 574
Risk Groups		
PWID	2.5% (2003); 2-8.1% (2010-2014)	16% (2007), 13% (2022)
MSM	0.25% (2019-2022); N=1 600	0.13% (2019-2022); N=4 783
Prisoners	NA	26 (2006)
Other	-	-
Healthcare workers	NA	0.7% (2000-2013)
	In risk-population (specify)	Prevalence (year)
HEV RNA	Blood donors	0.057% (2019, a study); N=8 817
Anti HEV (IgG)	Blood donors	7.6% (2019, a study); N=1264



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EPIDEMIOLOGY - HEV

PREVALENCE	HBsA (year)	HCV RNA (year)
General population	?	0.01% (2022)
Blood donors (first time)	0.004 % (2018-2020) N=250 640	0.000% (2018-2020) N=250 640
Pregnant women	NA	0.08% (2013) N=9 574
Risk Groups		
PWID	2.5% (2003); 2-8.1% (2010-2014)	16% (2007), 13% (2022)
MSM	0.25% (2019-2022); N=1 600	0.13% (2019-2022); N=4 783
Prisoners	NA	26 (2006)
Other	-	-
Healthcare workers	NA	0.7% (2000-2013)
	In risk-population (specify)	Prevalence (year)
Anti HAV	NA	NA
Anti HEV	Blood donors	7.6% (2019, a study); N=1264



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EPIDEMIOLOGY - HAV



Spread of hepatitis A virus strains of genotype IB in several EU countries and the United Kingdom

News

29 Sep 2022

An **OUTBREAK**

March-September 2022:

48 cases

25 hospitalised (a few severe cases)

MSM → general population

Source: Budapest (“Hungarian HAV”)



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VACCINATION PROGRAM

Hepatitis A	Y/N	population + Schedule	Since/period
Universal	N	-	-
Risk group	Y	Professional / epidemiological exposure MSM	1993 2015
Hepatitis B	Y/N	population + Schedule	Since/period
Universal	Y	Children	Birth cohort ≥1992
Catch-up	N	-	-
Risk Group	Y	HCW PWID Prisoners MSM Medical indication Family members/sex partners Refugees from Ukraine	1986 1995 2016 2015 1993 1993 2022



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NATIONAL PROGRAM FOR CHILDREN VACCINATION

- 3rd dose vaccination coverage:

YEAR	3rd dose (%)
2016/2017	88,7
2017/2018	87,2
2018/2019	87,5
2019/2020	79,9
2020/2021	79,9

- Birth cohort ≥ 1992

WHEN?

- Untill 2020: at age 5-6 y
- From Jan 2020: at age 3 mth
(6-valent vaccine)



SLOVENIA SCREENING

Recommended for following groups:	Hep B	Hep C
General population	N	N
Birth cohorts	N	N
Blood and organ donors	Y	Y
Pregnant women	Y	N
PWID	Y	Y
STI clinic patients	Y	Y
Haemodialysis patients	Y	Y
Health care workers	Y	Y
Men having sex with men	Y	Y
Prison population	Y	Y
Migrants	N	N
Others: Refugees from Ukraine	Y	Y

SLOVENIA SCREENING

Recommended for following groups:	Hep B	Hep C
General population	N	N
Birth cohorts	N	N
Blood and organ donors	Y	Y
Pregnant women	Y	N
PWID	Y	Y
STI clinic patients	Y	Y
Haemodialysis patients	Y	Y
Health care workers	Y	Y
Men having sex with men	Y	Y
Prison population	Y	Y
Migrants	N	N
Others: Refugees from Ukraine	Y	Y

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ACTIVE SEARCH FOR HBV/HCV SCREENING IN REFUGEES FROM UKRAINE

A QUESTIONNAIRE on possible HBV, HCV, HIV infections for refugees from Ukraine For personal use only (Engl/Ukr)

A QUESTIONNAIRE for displaced persons from Ukraine

regarding infections with HIV, hepatitis B virus, hepatitis C virus

(ONLY FOR PERSONAL USE)

By answering to the following questions you may importantly affect your health and receive appropriate medical care in Slovenia if needed.

1. Are infected with HIV: NO YES Not sure
2. Do you have hepatitis B: NO YES Not sure
3. Do you have hepatitis C: NO YES Not sure

If you answered to any of the questions with "YES", you can get medical care at one of the following centers for managing these diseases:

Ljubljana: Clinic for Infectious Diseases and Febrile Illnesses, Japljeva 2
Tel: 01 522 44 54 Email: narocanje.kibvs@kclj.si
Maribor: University Clinical Centre Maribor
Tel: 02 321 27 40 Email: infekcijska.ambulanta@ukc-mb.si
Celje: Splošna bolnica Celje
Tel: 03 423 30 08 Email: enarocanje@sb-celje.si
Murska Sobota: Splošna bolnica Murska Sobota
Tel: 02 512 34 55 Email: info@sb-ms.si
Novo mesto: Splošna bolnica Novo mesto
Tel: 07 391 65 82 Email: adm.infekcijski@sb-nm.si

Please continue if you answered to the questions above with "NO":

4. Is there any possibility that you are infected with HIV, hepatitis B virus or hepatitis C virus?
NO YES I don't know
5. Have you ever had a sexually transmitted infection (syphilis, gonorrhoea, chlamydia, etc.)?
NO YES I don't know
6. Have you or any of your family members had tuberculosis (now or in the past)?
NO YES I don't know

7. Have you ever injected drugs? NO YES
8. Have you had unprotected sex with multiple partners? NO YES
9. If you are male: have you had unprotected sex with men?
NO YES
10. Are you a sex worker? NO YES
11. Have you been sexually abused? NO YES
12. Have you ever been incarcerated? NO YES
13. Do you have a sexual partner, close family member, joint household member who has HIV infection or/and hepatitis B or/and hepatitis C?
NO YES I don't know
14. Have you ever received a blood transfusion? NO YES I don't know
15. Have you ever been treated with haemodialysis? NO YES I don't know
16. Are you taking medications that weaken your immune system?
NO YES I don't know
17. Do you have abnormal liver test results? NO YES I don't know
18. Are you pregnant? NO YES I don't know

If you answered to any of the questions with "YES" or "I don't know", you are advised to test for infections with HIV, hepatitis B virus and hepatitis C virus.

You can get tested at general practitioners or you can be tested anonymously and free-of-charge at: Clinic for Infectious Diseases and Febrile Illnesses, Poljanski nasip 58, Ljubljana (on Mondays between 12.00-14.00) or at Blood Transfusion Centers (<http://www.ztm.si/testiranje/>).

If you are an active drug user, you can get tested at one of 22 Centers for Prevention and Treatment of Drug Addiction (CPZOPD) where you can also get a medical care for drug addiction (<https://www.infodroga.si/kam-po-nasvet/seznam-centrov-cpzopd/>).

In case the result of your test is POSITIVE, you will be provided with the necessary medical care at one of five centers for managing HIV, viral hepatitis B and C infections.

Accessible

On-line:

- Ministry of health RS
- National Institute of Public Health
- University Medical Centers
- General Hospitals
- Centers for Refugees

In paper:

- When applying for a Temporary Protection
- Centers for Refugees



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TREATMENT

National guidelines available	Y/N (year)
Hepatitis B	Y (2000 + updates)
Hepatitis C	Y (1997+ updates)
Management of HCV infection in PWID	Y (2007)
Prevention of HBV reactivation in persons on immunosuppressive therapy	Y (2008)
Management of HBV in patients with chronic kidney disease – an update	Y (2021)

National Viral Hepatitis Expert Board. CPG for HCV treatment. Ljubljana, 1997-2021.

National Viral Hepatitis Expert Board. CPG for HBV treatment. Ljubljana, 2000-2019.

Maticic M, Kastelic A. Zdrav Vestn 2009; 78: 529-39.

Matičič M, Poljak M. Zdrav Vestn 2010; 79: 599-608.

Mlinšek G, Oblak M, Arnol M, Matičič M. Clin Nephrol 2021 Suppl;96(1):119-126.



SLOVENIA TREATMENT

Drugs available for HCV treatment*	Y/N
PegInterferon	N
Ribavirin	N
Boceprevir	N
Telaprevir	N
Simeprevir	N
Sofosbuvir	N
Daclatasvir	N
Sofosbuvir/Ledipasvir	N
Sofosbuvir/Velpatasvir	Y
Glecaprevir/Pibrentasvir	Y
Sofosbuvir/Velpatasvir/Voxilaprevir	Y
Other (specify)	-

Drugs available for HBV treatment*	Y/N
Lamivudine	Y
Adefovir	N
Telbivudine	N
Entacavir	Y
TDF	Y
TAF	Y

Number of patients treated (cumulative)

Hep B	801 (2000-2020)
Hep C	2 823 (1997-2020)

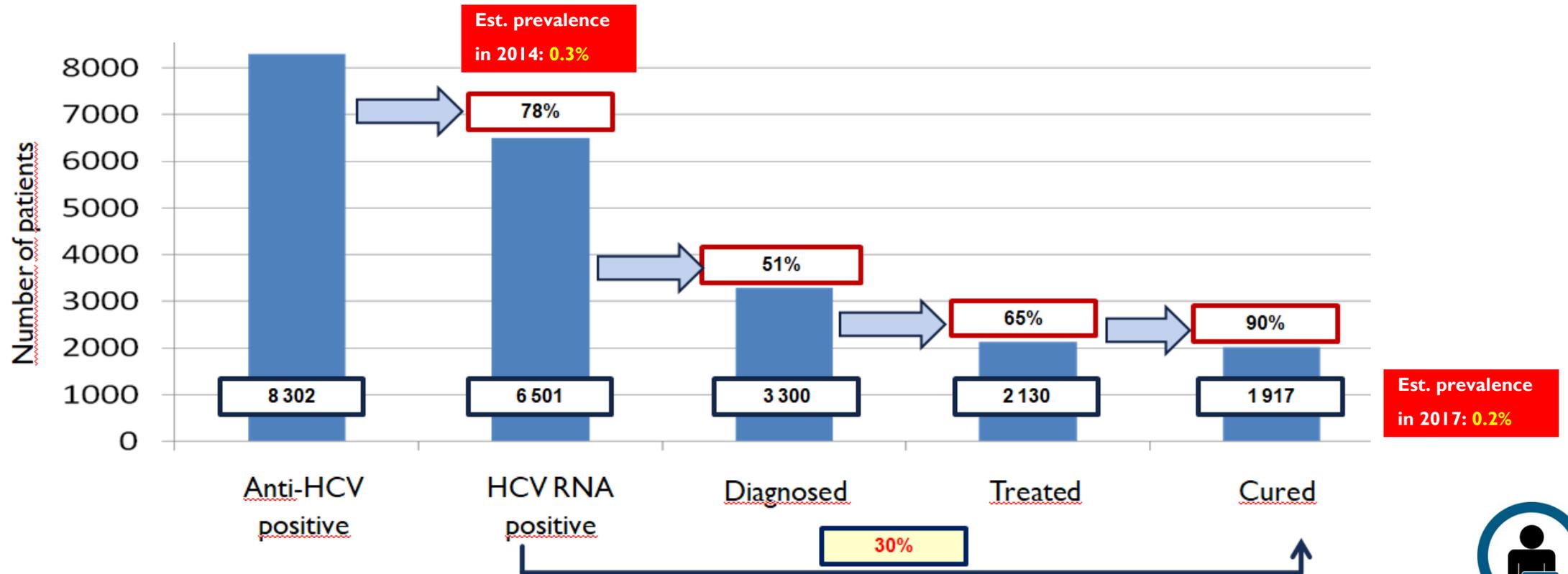
*included on the national essential medicines list



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HCV BURDEN - 2014

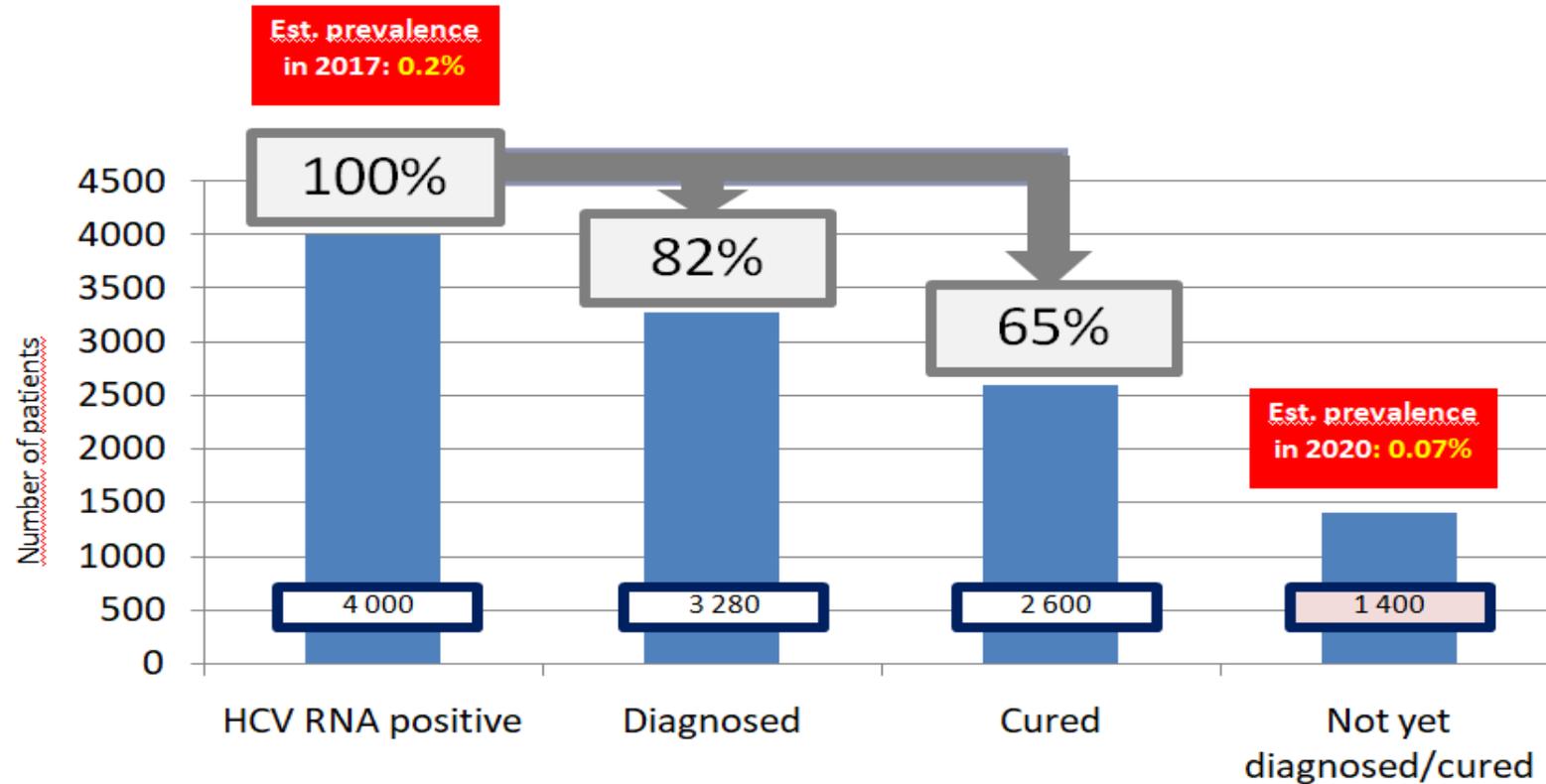
A Cascade-of-Care (1997-2014): est. HCV RNA prevalence **0.3%**



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HCV BURDEN - 2020

A Cascade-of-Care (2017-2020): est. HCV RNA prevalence **0.07%**



THANK YOU

