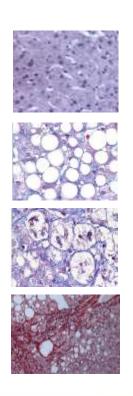
$$\cosh x + b_0 \sinh x \right) \widetilde{\varsigma}^2(\varepsilon) = \widetilde{\varsigma}^2(\varepsilon) = \frac{\sum_{i=1}^{j=1}}{n-2n}, (1)$$

NASH and bariatric surgery

Francois Pattou
University Hospital, Lille, France

Definition: NASH vs NAFLD (MASLD)



```
Normal liver

NAFL
(steatosis)

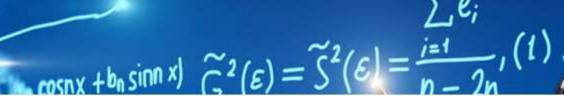
NASH
(inflammation necrosis)

Cirrhosis
(fibrosis)
```

Liver histology

NASH / NAFLD Activity score (0-8) Fibrosis Kleiner score (0-4)

- Non invasive biomarkers (NFS, APRI, FIB-4)
- Non invasive imaging (MRI, Elastography)



Efficacy: NASH after surgery

Meta-analysis of cohort studies

Chavez Tapia Cochrane Review 2010

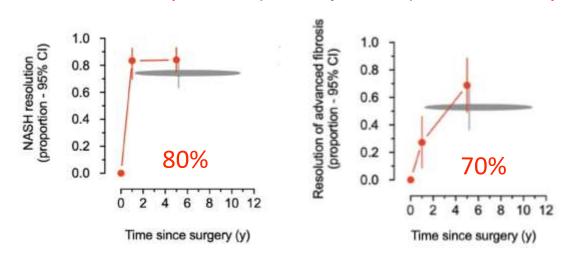
Prospective cohort study

Baseline	1 year	5 years		
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Lassailly et al. Gastroenterology 2020

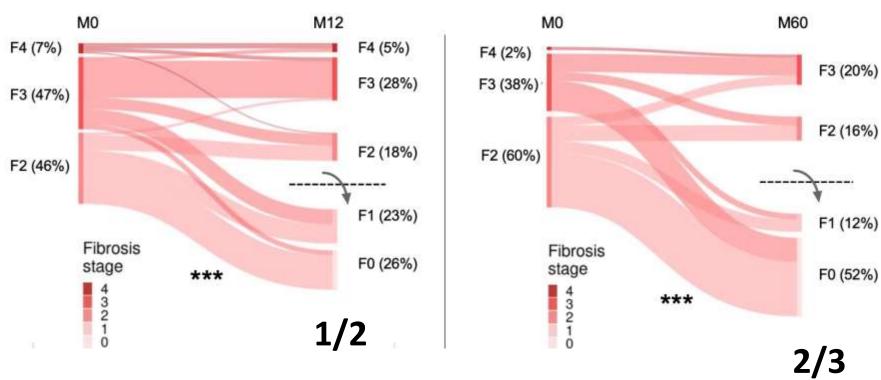
NASH Advanced Fibrosis

Prospective (retrospective) cohort study

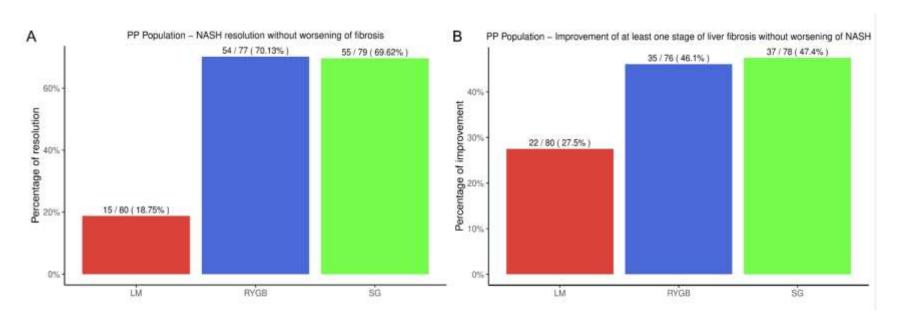


Pattou et al. Hepatology 2022

Efficacy: fibrosis remission after surgery



Efficacy: RCT - Surgery vs Medical treatment

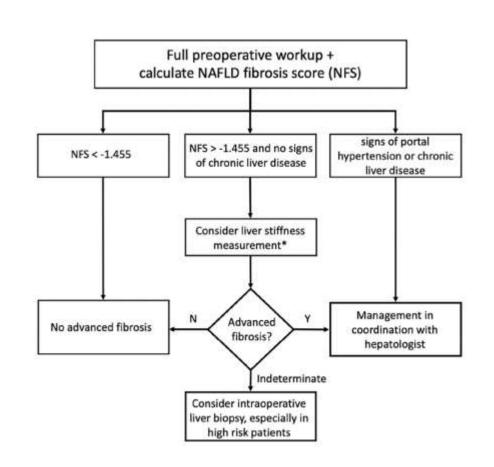


Verrastro et al. Lancet 2023

NASH diagnosis

ASMBS position statement

Mazzini et al. SOARD 2021



$$\frac{\sum_{i=1}^{n} e_i}{\sum_{i=1}^{n} e_i} \lesssim e_i = \frac{\sum_{i=1}^{n} e_i}{\sum_{i=1}^{n} e_i} \lesssim e_i$$

Non invasive liver tests

NFS, NAFLD fibrosis score

FIB-4, Fibrosis-4 index

APRI, AST to platelet ratio index

HFS, hepamet fibrosis index

FNI, fibrotic NASH index

Non-invasive test	FIB-4	NFS	APRI	HFS	FNI
First author, year of publication (# ref)	Sterling, 2006 (27)	Angulo, 2007 (28)	Wai, 2003 (29)	Ampuero, 2020 (30)	Tavaglione, 2023 (31)
Reference lower cut- off	1.3	-1.445	1	0.12	0.1
Variables					
Age	X	X		x	
Gender				X	
BMI		X			
AST	X	X	X	X	X
ALT	X	X			
Albumin		X		X	
Platelets	x	x	X	X	
HbA1c					X
Type 2 diabetes		X		X	
HOMA-IR				X	
HDL cholesterol					X

Diagnosing significant fibrosis in BS

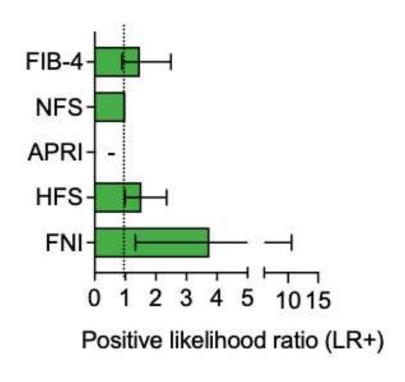
Non-invasive tests	AUC M0	AUC M12	AUC M60
FIB-4	0.71 (0.67-0.74)	0.75 (0.69-0.80)	0.69 (0.59-0.80)
NFS	0.68 (0.64-0.72)	0.68 (0.61-0.75)	0.75 (0.67-0.83)
APRI	0.75 (0.72-0.79)	0.75 (0.69-0.80)	0.73 (0.61-0.84)
HFS	0.74 (0.70-0.78)	0.72 (0.64-0.80)	0.82 (0.74-0.91)
FNI	0.79 (0.76-0.82)	0.74 (0.68-0.80)	0.84 (0.77-0.92)

Numbers in brackets are 95% CI.

Diagnosing remission of significant fibrosis

Non-invasive tests	AUC M12	AUC M60		
FIB-4	0.69 (0.59-0.78)	0.70 (0.54-0.85)		
NFS	0.59 (0.47-0.70)	0.70 (0.55-0.85)		
APRI	0.67 (0.58-0.77)	0.72 (0.57-0.87)		
HFS	0.68 (0.57-0.69)	0.83 (0.71-0.96)		
FNI	0.67 (0.56-0.77)	0.83 (0.71-0.95)		

Diagnosing 5-year remission of significant fibrosis



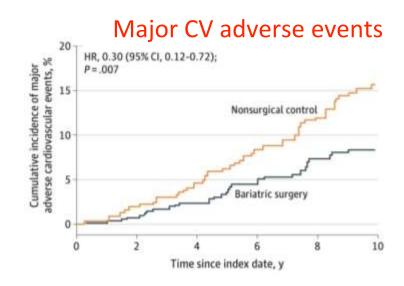
Safety: Surgery in NASH

Meta analysis

Type of surgery	30 day mortality				
	Overall Child-Pugh class			MELD	
	(%)	A (%)	B (%)	C (%)	score
Appendectomy	9	n.a.	n.a.	n.a.	n.a.
Cardiac	16-17	0-3	42-50	100	n.a.
Cholecystectomy	1-3	0.5	3	n.a.	<8 = 0% ≥8 = 6%
Colorectal cancer surgery	12.5	6	13	27	n.a.
Esophagectomy	17	n.a.	n.a.	n.a.	n.a.
Major abdominal surgery	26-30	10	30-31	76-82	n.a.

Bhangui et al. J Hepatol 2012

Retrospective controled cohort study



Aminian et al. JAMA 2021

$$|\cos(x) + b_n \sin(x)| \approx 2(\varepsilon) = \frac{\zeta^2}{2}(\varepsilon) = \frac{i-1}{n-2n}, (1)$$

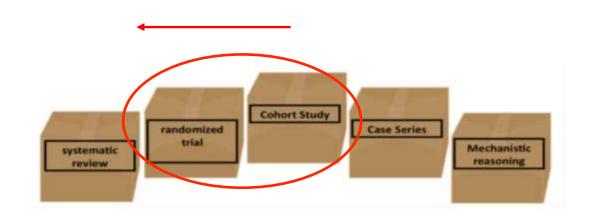
Surgery for NASH?

- -> Randomized controlled trials (ongoing)
 - 1. BRAVES (BS Versus Non-alcoholic Steato-hepatitis) NCT03524365, Italy
 - 2. BariaNash (BS for NASH With Advanced Liver Fibrosis) NCT03472157, France
 - 3. VSG and Lifestyle Modification for the Treatment of NASH NCT03587831, USA
 - 4. BeLEANeR (BS vs. Lifestyle Modification for NASH) NCT04298736, Brazil
 - 5. NASH-APOLLO (Endoscopic Gastric Tubulization in Patients With NASH) NCT03426111, Spain

$$|\cos(x)| + |\cos(x)| \approx 2 (\varepsilon) = \frac{\zeta^2(\varepsilon)}{\zeta^2(\varepsilon)} = \frac{\zeta^2(\varepsilon)}{|x-2n|} = \frac{\zeta^2(\varepsilon)}{|x-2n|}$$

Surgery for NASH?

"The Oxford 2011 Levels of Evidence". http://www.cebm.net



- Does surgery help to treat NASH? -> Level 1-2
- What are the common (rare) harms of surgery in NASH? -> Level 2-3