

MOET ANTIBIOTISCHE PROFYLAXE WORDEN AANGEPAST BIJ HOOG RISICO GROEPEN?

Marjan Wouthuyzen-Bakker

Internist-infectioloog

University Medical Center Groningen, the Netherlands



umcg



NINJA.

Northern Infection Network Joint Arthroplasty

HOOG RISICO GROEPEN VOOR PJI

Algemeen:

Risico PJI 1-2%

1.5 – 10 hoger risico
op PJI

Second INTERNATIONAL
CONSENSUS MEETING (ICM)
on MUSCULOSKELETAL INFECTION



TABLE 1. Summary of risk factors associated with development of SSI/PJI

Modifiable Host Factors	Factors with Limited Evidence of Associations with SSI/PJI
<ul style="list-style-type: none">BMI – StrongSmoking – StrongHigh alcohol intake (alcohol abuse) – StrongLow income – StrongMalnutrition (low serum albumin) – StrongHistory of DM – StrongHistory of CVD – ModerateHistory of CHF – StrongHistory of cardiac arrhythmia – StrongHistory of PVD – StrongChronic pulmonary disease – StrongChronic obstructive pulmonary disease – StrongHistory of renal disease – StrongHistory of liver disease/cirrhosis – StrongHistory of RA – StrongHistory of cancer/malignancy – StrongHistory of osteonecrosis – StrongHistory of depression – StrongHistory of psychosis – StrongHistory of HIV/AIDS – StrongNeurologic disease (hemiplegia, paraplegia) – ModerateHistory of corticosteroid administration – StrongHistory of intra-articular corticosteroid injection – ModeratePrevious joint surgery – StrongRevision arthroplasty – StrongPrevious joint infection – ModerateFrailty – ModeratePreoperative anemia – StrongASA grade > 2 – StrongCharlson comorbidity index (high) – StrongPreoperative hyperglycemia and high HbA1c – ModerateAllogenic blood transfusion – StrongProphylaxis with warfarin or low molecular weight heparin – Moderate	<ul style="list-style-type: none">Age (as a continuous exposure) – LimitedHispanic ethnicity – LimitedNative American and Eskimo ethnicity – LimitedAsian race – LimitedHistory of drug abuse – LimitedRural location vs. non-rural location – LimitedUnderweight – LimitedHistory of hypertension – LimitedHistory of osteoarthritis – LimitedHistory of post-traumatic arthritis – LimitedLow- or high-risk dental procedures – LimitedHistory of UTI – LimitedHistory of dementia – LimitedHypercholesterolemia – LimitedPeptic ulcer disease – LimitedValvular disease – LimitedMetastatic tumor – LimitedHistory of coagulopathy – LimitedHistory of venous thromboembolism – LimitedPulmonary circulatory disorders – LimitedHypothyroidism – LimitedHepatitis (B or C) – LimitedElectrolyte imbalance – LimitedAutogenous blood transfusion – Limited
Non-modifiable Host Factors	<ul style="list-style-type: none">Age (≥ 75 years) – ModerateMale sex – StrongBlack race – StrongTKA vs. THA – Strong

HOOG RISICO GROEPEN VOOR PJI

Algemeen:

Risico PJI 1-2%

1.5 – 10 keer
op

WAAR TE BEGINNEN?
EN HOE?

Second INTERNATIONAL
CONSENSUS MEETING (ICM)
on MUSCULOSKELETAL INFECTION



TABLE 1. Summary of risk factors associated with development of SSI/PJI

Modifiable Host Factors	Factors with Limited Evidence of Associations with SSI/PJI
<ul style="list-style-type: none">BMI – StrongSmoking – StrongHigh alcohol intake (alcohol abuse) – StrongLow income – StrongMalnutrition (low serum albumin) – StrongHistory of DM – StrongHistory of CVD – ModerateHistory of CHF – StrongHistory of cardiovascular disease – Strong	<ul style="list-style-type: none">Age (as a continuous exposure) – LimitedHispanic ethnicity – LimitedNative American and Eskimo ethnicity – LimitedAsian race – LimitedHistory of drug abuse – LimitedRural location vs. non-rural location – LimitedUnderweight – Limited
<ul style="list-style-type: none">Previous history of HIV/AIDS – StrongNeurologic disease (hemiplegia, paraplegia) – ModerateHistory of corticosteroid administration – StrongHistory of intra-articular corticosteroid injection – ModeratePrevious joint surgery – StrongRevision arthroplasty – StrongPrevious joint infection – ModerateFrailty – ModeratePreoperative anemia – StrongASA grade > 2 – StrongCharlson comorbidity index (high) – StrongPreoperative hyperglycemia and high HbA1c – ModerateAllogenic blood transfusion – StrongProphylaxis with warfarin or low molecular weight heparin – Moderate	<ul style="list-style-type: none">History of coagulopathy – LimitedHistory of venous thromboembolism – LimitedPulmonary circulatory disorders – LimitedHypothyroidism – LimitedHepatitis (B or C) – LimitedElectrolyte imbalance – LimitedAutogenous blood transfusion – Limited
Non-modifiable Host Factors	<ul style="list-style-type: none">Age (≥ 75 years) – ModerateMale sex – StrongBlack race – StrongTKA vs. THA – Strong

Extended Oral Antibiotic Prophylaxis in High-Risk Patients Substantially Reduces Primary Total Hip and Knee Arthroplasty 90-Day Infection Rate

Avinash Inabathula, MD, Julian E. Dilley, BS, Mary Ziembra-Davis, BA, Lucian C. Warth, MD,
Khalid A. Azzam, MD, Philip H. Ireland, MD, and R. Michael Meneghini, MD

Investigation performed at the Department of Orthopaedic Surgery, Indiana University School of Medicine, Indianapolis, Indiana

J Bone Joint Surg Am. 2018;100:2103-9

VERLENGDE ORALE ANTIBIOTICA PROFYLAXE IN HOOG RISICOGROEPEN?

2181 primaire TJA

Retrospectieve studie (2011-2016)

2015: 7 dagen orale AB profylaxe geïmplementeerd

Outcome: PJI < 90 days

High risk groups, ≥ 1 :

- BMI $\geq 35 \text{ kg/m}^2$
- Diabetes mellitus
- Active smoker
- Chronic kidney disease
- Autoimmune disease
- Nasal colonization with MRSA or MSSA

VERLENGDE ORALE ANTIBIOTICA PROFYLAXE IN HOOG RISICOGROEPEN?

2181 primaire TJA

Retrospectieve studie (2011-2016)

2015: 7 dagen orale AB profylaxe geïmplementeerd

Outcome: PJI < 90 days

High risk groups, ≥ 1:

- BMI $\geq 35 \text{ kg/m}^2$
- Diabetes mellitus
- Active smoker
- Chronic kidney disease
- Autoimmune disease
- Nasal colonization with MRSA or MSSA

	Low risk	High risk
TKA	0.3% (1/369)	2.1% (10/468)
THA	1.5% (5/336)	4.3% (12/282)
Total	0.9% (6/705)	2.9% (22/750)

VERLENGDE ORALE ANTIBIOTICA PROFYLAXE IN HOOG RISICOGROEPEN?

2181 primaire TJA

Retrospectieve studie (2011-2016)

2015: 7 dagen orale AB profylaxe geïmplementeerd

Outcome: PJI < 90 days

2015

High risk groups, ≥ 1:

- BMI $\geq 35 \text{ kg/m}^2$
- Diabetes mellitus
- Active smoker
- Chronic kidney disease
- Autoimmune disease
- Nasal colonization with MRSA or MSSA

	Low risk	High risk	High risk, extended AB	P-value
TKA	0.3% (1/369)	2.1% (10/468)	0.4% (2/450)	0.009
THA	1.5% (5/336)	4.3% (12/282)	1.1% (3/276)	0.020
Total	0.9% (6/705)	2.9% (22/750)	0.7% (5/726)	0.006

VERLENGDE ORALE ANTIBIOTICA PROFYLAXE IN HOOG RISICOGROEPEN?

2181 primaire TJA

Retrospectieve studie (2011-2016)

2015: 7 dagen orale AB profylaxe geïmplementeerd

Study limitations:

- **67.7% geclassificeerd als hoog risico (NNT 47)**
- **Bias: incidentie PJI laag risico groep vanaf 2015?**
- **Is infectie voorkomen of uitgesteld?**

- Nasal colonization with MRSA or MSSA

Total	0.9% (6/705)	2.9% (22/750)	0.7% (5/726)	0.006
-------	-----------------	------------------	-----------------	-------

Perioperative Antibiotic Prophylaxis in Total Joint Arthroplasty

A Single Dose Is as Effective as Multiple Doses

Timothy L. Tan, MD, Noam Shohat, MD, Alexander J. Rondon, MD, MBA, Carol Foltz, PhD, Karan Goswami, MD,
Sean P. Ryan, MD, Thorsten M. Seyler, MD, PhD, and Javad Parvizi, MD, FRCS

Investigation performed at The Rothman Institute at Thomas Jefferson University, Philadelphia, Pennsylvania

J Bone Joint Surg Am. 2019;101:429-37

1 GIFT VERSUS MULTIPLE GIFTEN

20682 primaire TJA

2006-2017

SINGLE DOSE
(n=4523)

MULTIPLE DOSE
(n=16159)

PJI < 1 jaar



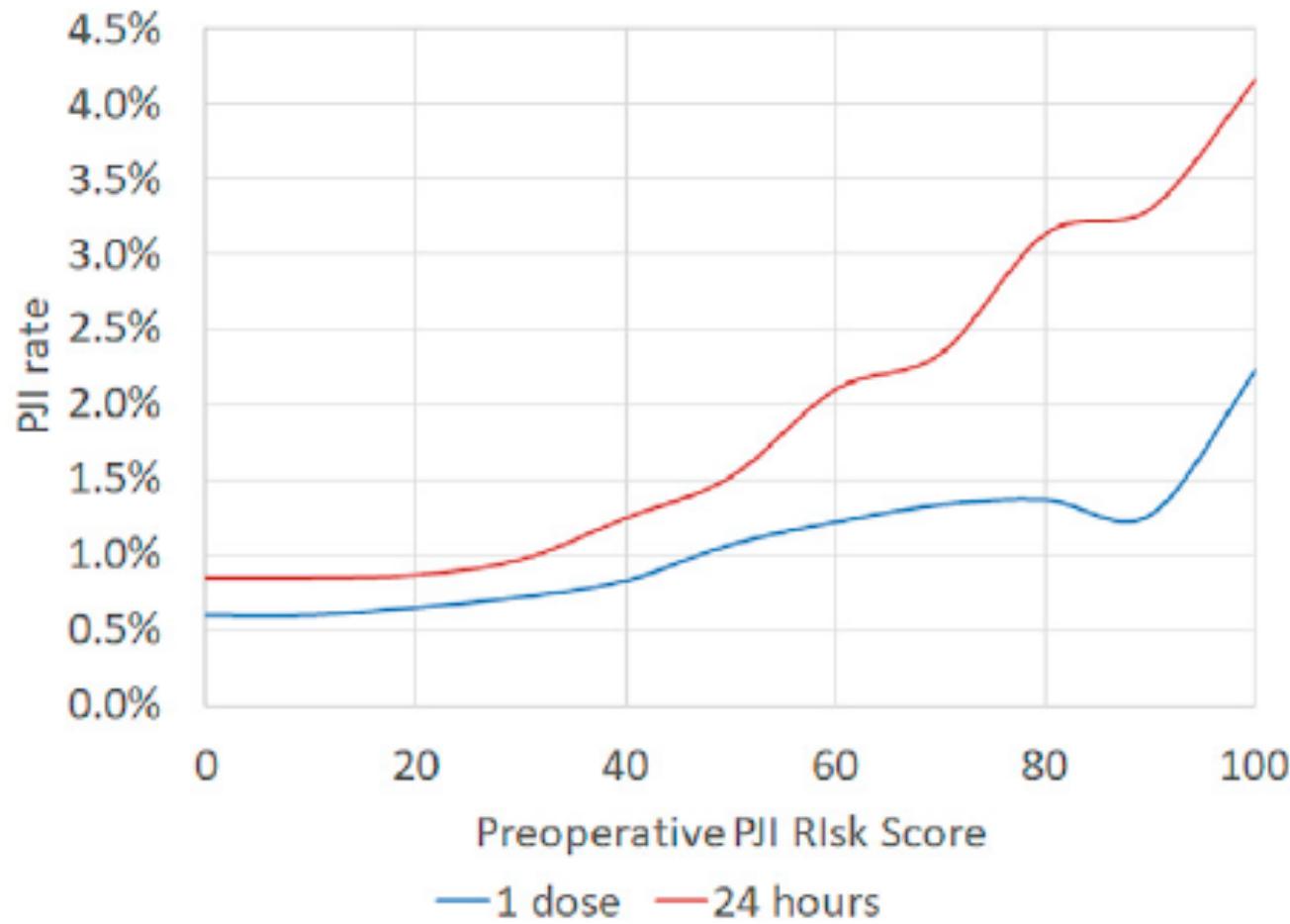
0.6%



0.9%

$$P = 0.66$$

HOOG RISICO PATIËNTEN



UITZONDERING TUMOR CHIRURGIE?

Systematic review 4836 patiënten

Deep infection rate 13% (<24h) → 8% (>24h)



BREDERE OF ANDERE PROFYLAXE NODIG?

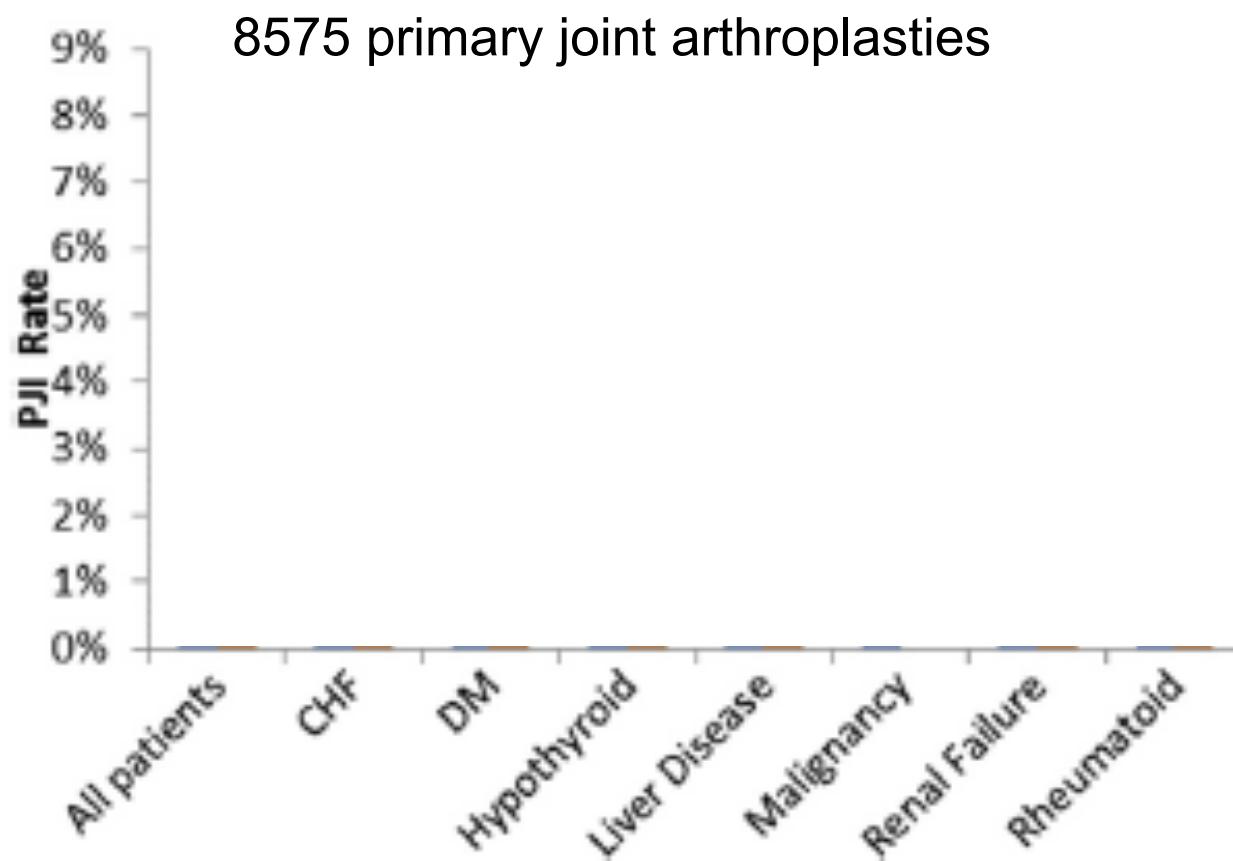


Fig. 2. Rate of PJI by patient's comorbidity and perioperative antibiotic. DM, diabetes mellitus; CHF, congestive heart failure.

VERWEKKERS IN RELATIE TOT COMORBIDITEIT

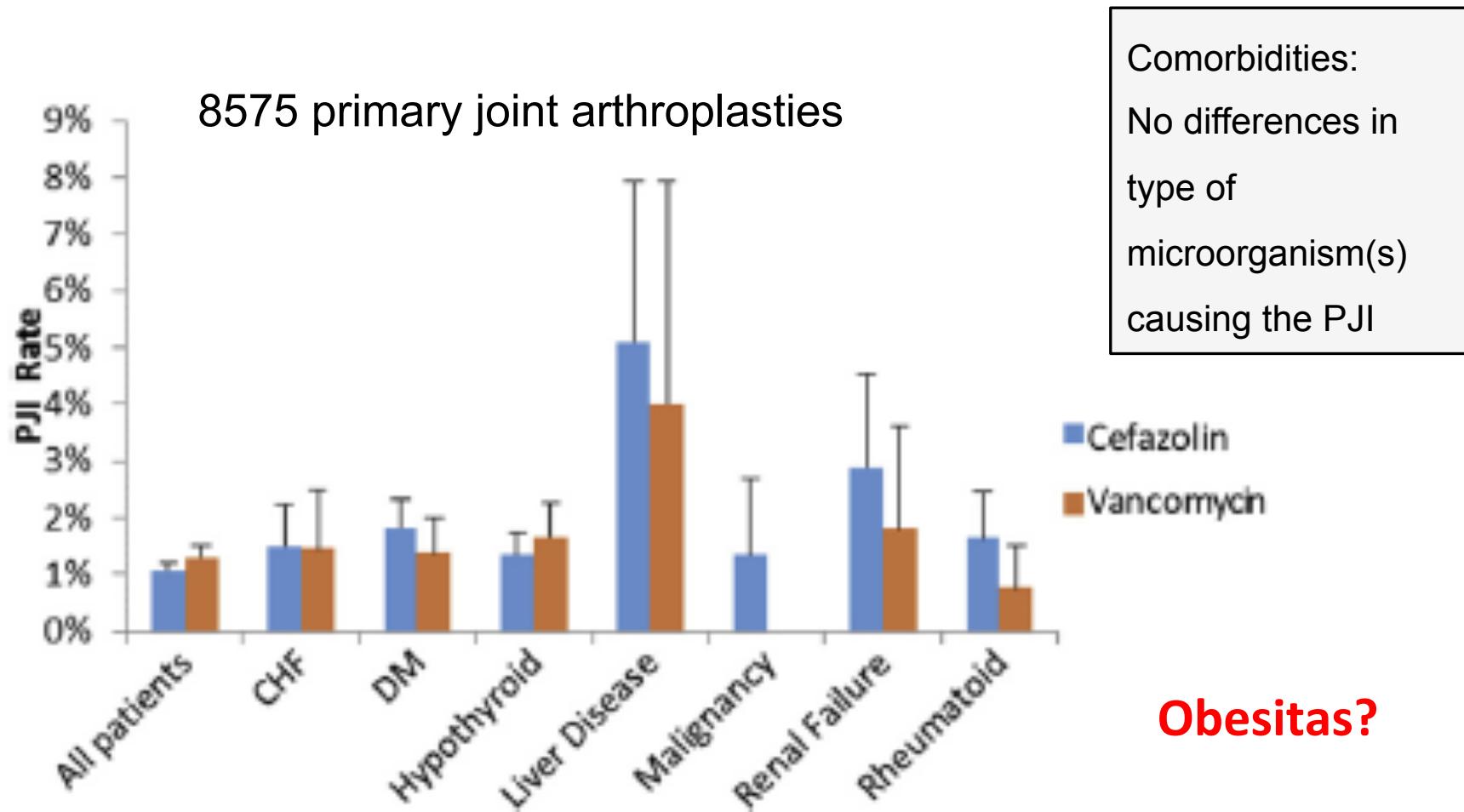
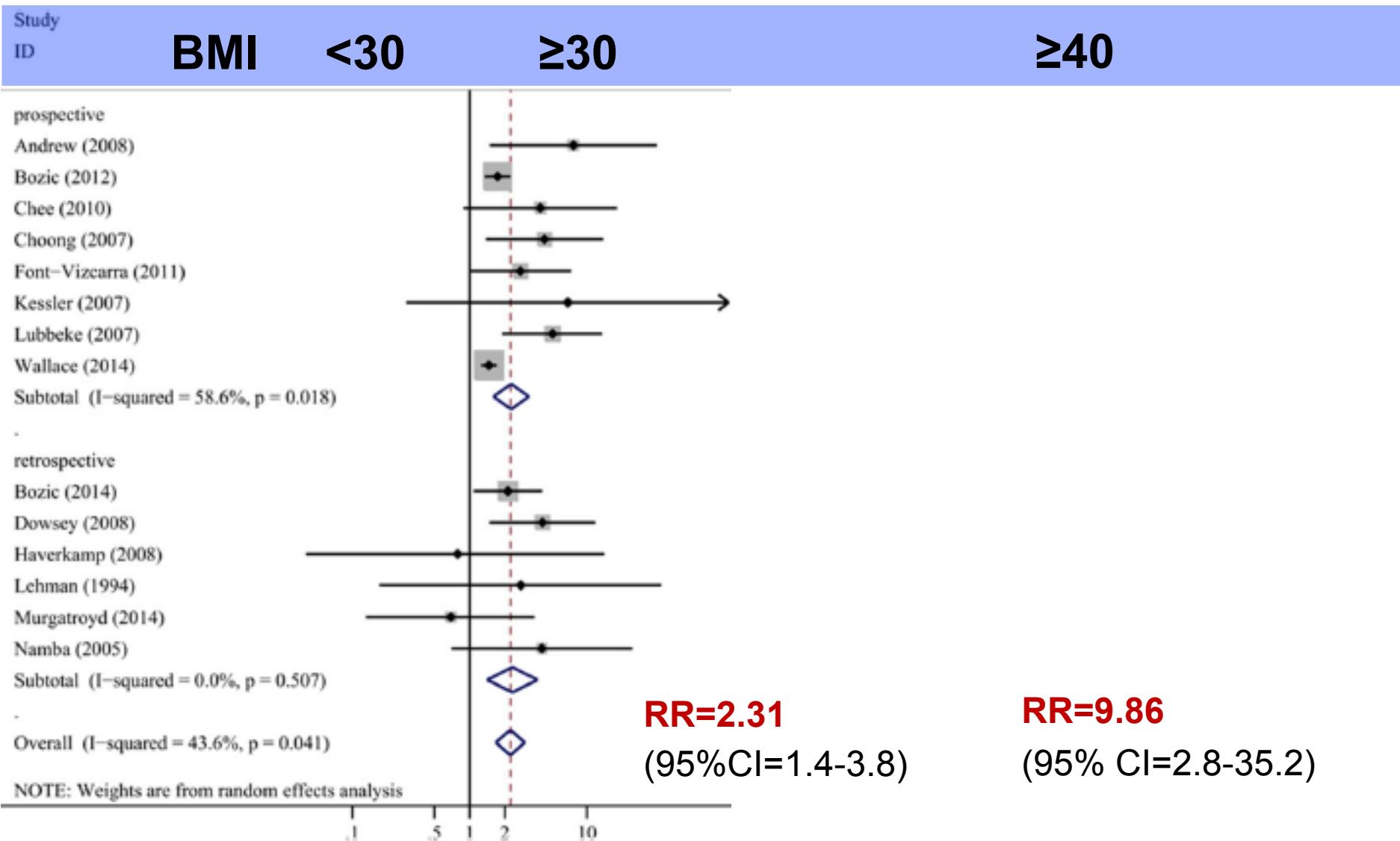


Fig. 2. Rate of PJI by patient's comorbidity and perioperative antibiotic. DM, diabetes mellitus; CHF, congestive heart failure.

OBESITAS ALS RISICO FACTOR VOOR PJI



VERWEKKERS PJI OBESE VS NIET OBESE PATIËNTEN

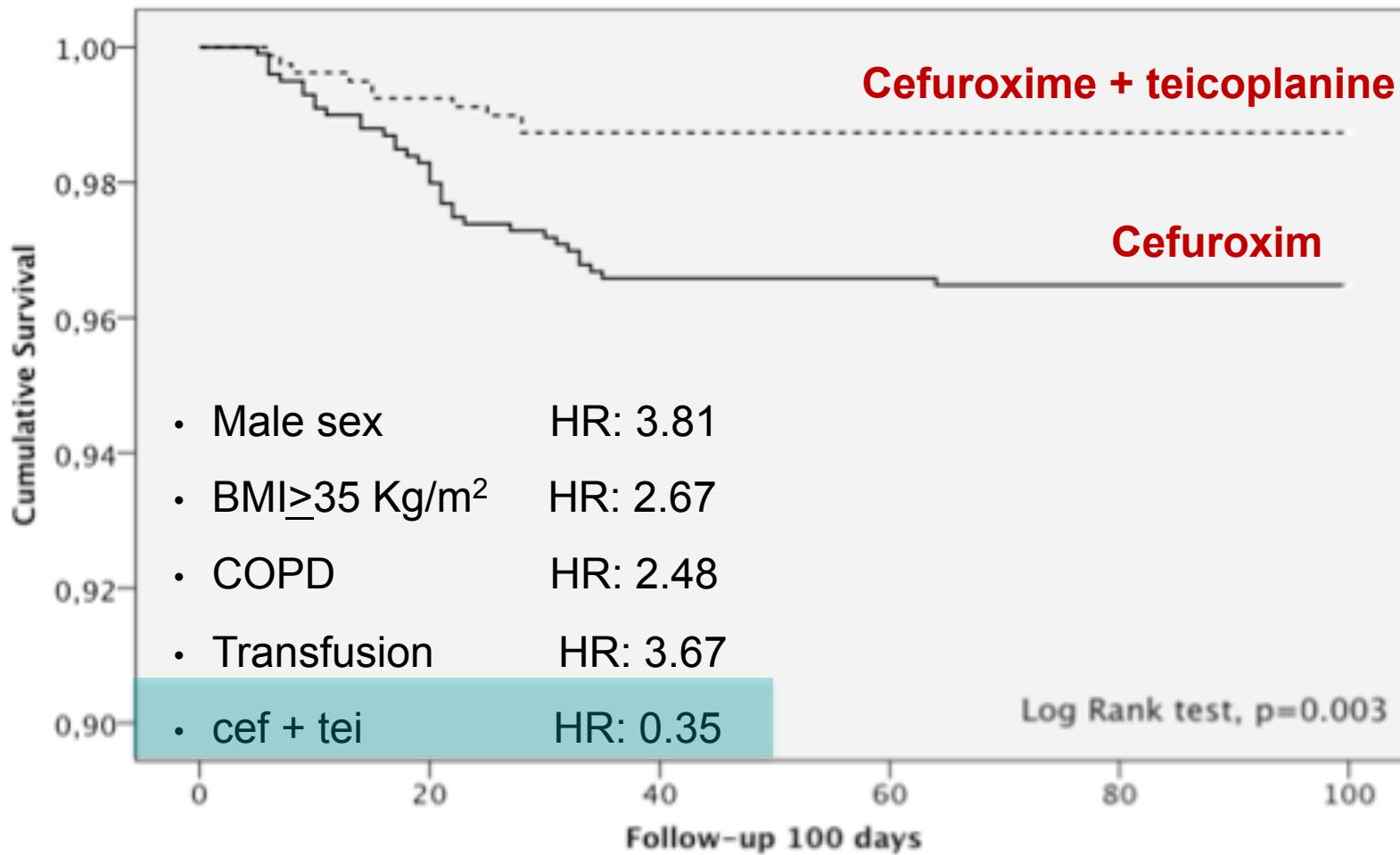
EARLY PJI	HIP	BMI <30 (n=79)	BMI 30-35 (n=43)	BMI ≥35 (n=38)	P value
Polymicrobial infection		34.2%	67.4%	68.4%	.001
Gram-positives		96.2%	95.3%	97.4%	.892
	<i>Staphylococcus aureus</i>	41.8%	41.9%	36.8%	.863
	<i>Staphylococcus epidermidis</i>	31.6%	41.9%	44.7%	.308
	<i>Corynebacterium</i> species	8.9%	34.9%	13.2%	.008
	<i>Enterococcus</i> species	12.7%	29.4%	36.8%	.007
	<i>Streptococcus</i> species	25.3%	27.9%	10.5%	.122
Gram-negatives		12.7%	16.3%	34.2%	.005
	<i>Escherichia coli</i>	2.5%	2.3%	5.3%	.685
	<i>Enterobacter cloacae</i>	2.5%	2.3%	5.3%	.685
	<i>Proteus</i> species	2.5%	0.0%	18.4%	<.001
	<i>Pseudomonas</i> species	3.8%	4.7%	2.6%	.892
	<i>Morganella morganii</i>	0.0%	0.0%	7.9%	.002
Anaerobes		3.8%	2.3%	7.9%	.445

VERWEKKERS PJI OBESE VS NIET OBESE PATIËNTEN

EARLY PJI	HIP	BMI <30 (n=79)	BMI 30-35 (n=43)	BMI ≥35 (n=38)	P value
Polymicrobial infection		34.2%	67.4%	68.4%	.001
Gram-positives		96.2%	95.3%	97.4%	.892
	<i>Staphylococcus aureus</i>	41.8%		36.8%	.863
	<i>Staphylococcus epidermidis</i>			44.7%	.308
	<i>Corynebacterium</i>			13.2%	.008
				36.8%	.007
			27.9%	10.5%	.122
Gram-negatives		12.7%	16.3%	34.2%	.005
	<i>Escherichia coli</i>	2.5%	2.3%	5.3%	.685
	<i>Enterobacter cloacae</i>	2.5%	2.3%	5.3%	.685
	<i>Proteus species</i>	2.5%	0.0%	18.4%	<.001
	<i>Pseudomonas species</i>	3.8%	4.7%	2.6%	.892
	<i>Morganella morganii</i>	0.0%	0.0%	7.9%	.002
Anaerobes		3.8%	2.3%	7.9%	.445

No differences observed
in knees

DUO PROFYLAXE?



DUO PROFYLAXE?

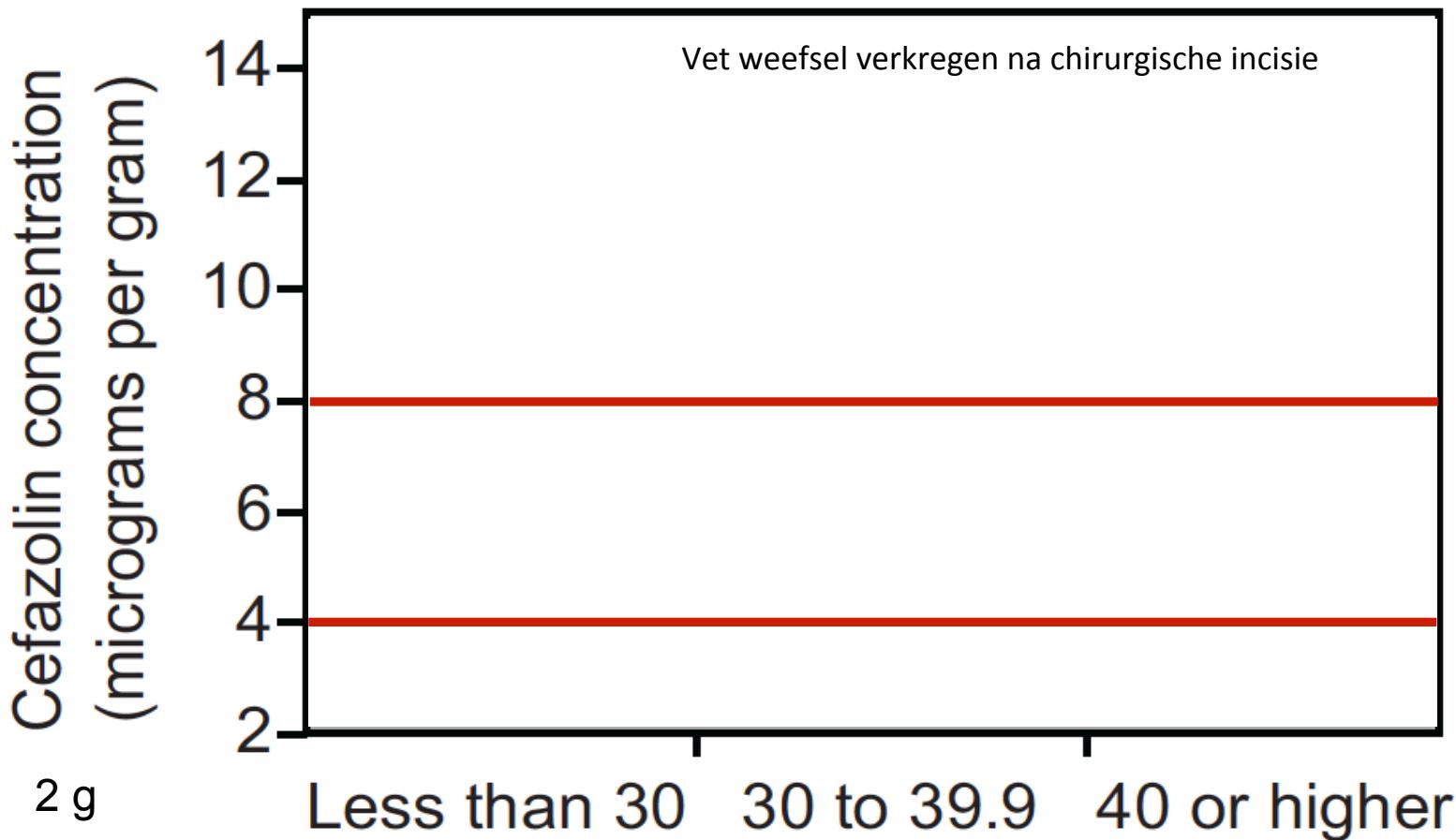
Patient subgroup (no. of patients) and PJI subgroup	C group (n = 995)	CT group (n = 791)	P value ^a
BMI of <30 kg/m ² (n = 960) ^b			
PJI	13 (2.6)	8 (1.8)	0.383
PJI due to GP ^c	9 (1.8)	6 (1.3)	0.558
BMI of ≥30 kg/m ² (n = 822) ^b			
PJI	22 (4.5)	2 (0.6)	0.001
PJI due to GP	20 (4.1)	1 (0.3)	0.001

* afname was met name in MSSA, doserings probleem?

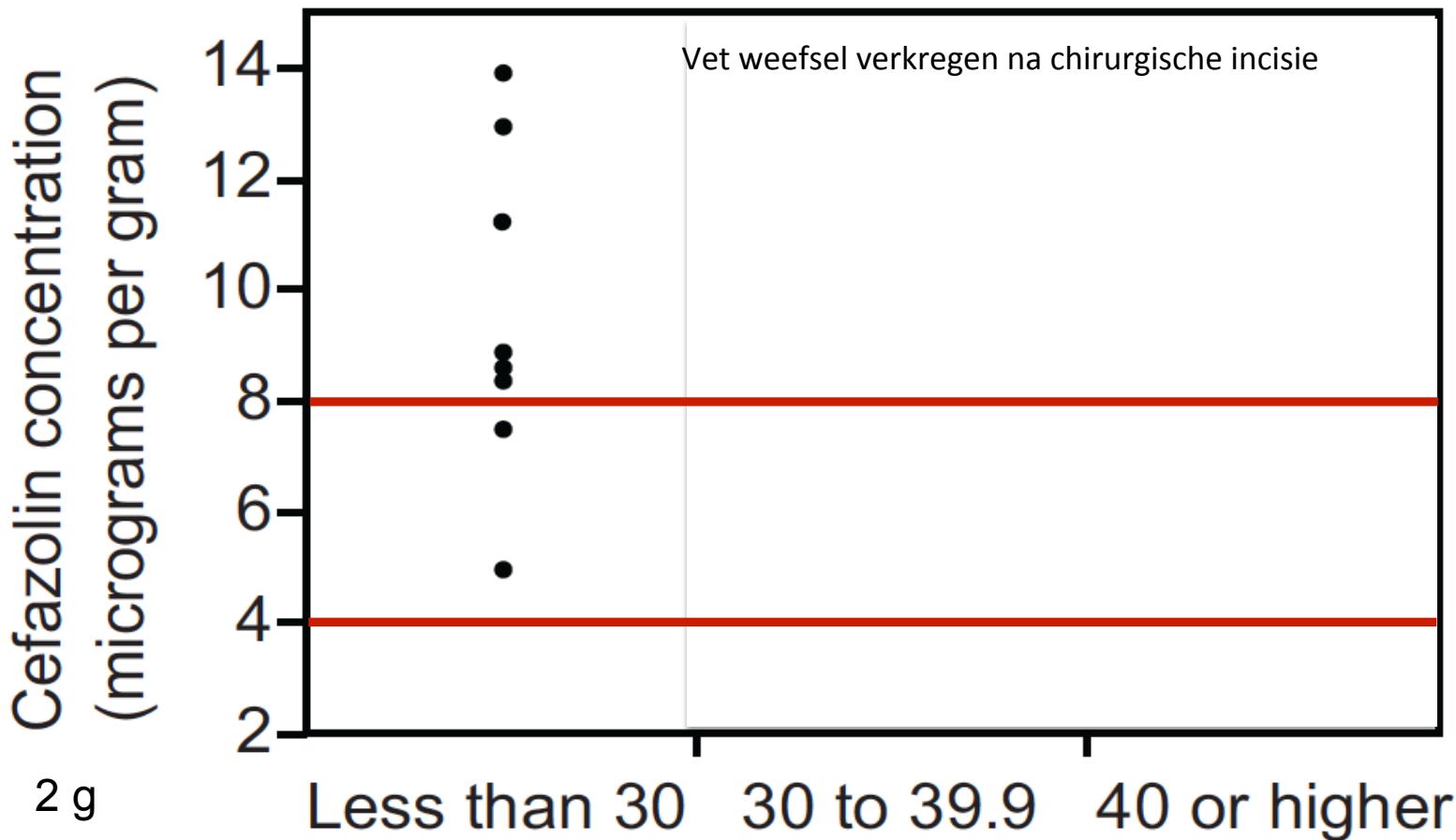
BLOED PERFUSIE IN VET WEEFSEL

Clinical feature	Control subjects (n° 10)	Obese patients before RYGB (n° 16)	Obese patients after RYGB (n° 16)
Age (years)	46 ± 9	45 ± 11	46 ± 11
Sex M/F	1/9	2/14	2/14
Body-mass index	21 ± 2	46 ± 6	33 ± 6
Adipose blood perfusion (PU)	79 (34)	4.8 (2.7) ↓	10 (6) ↓

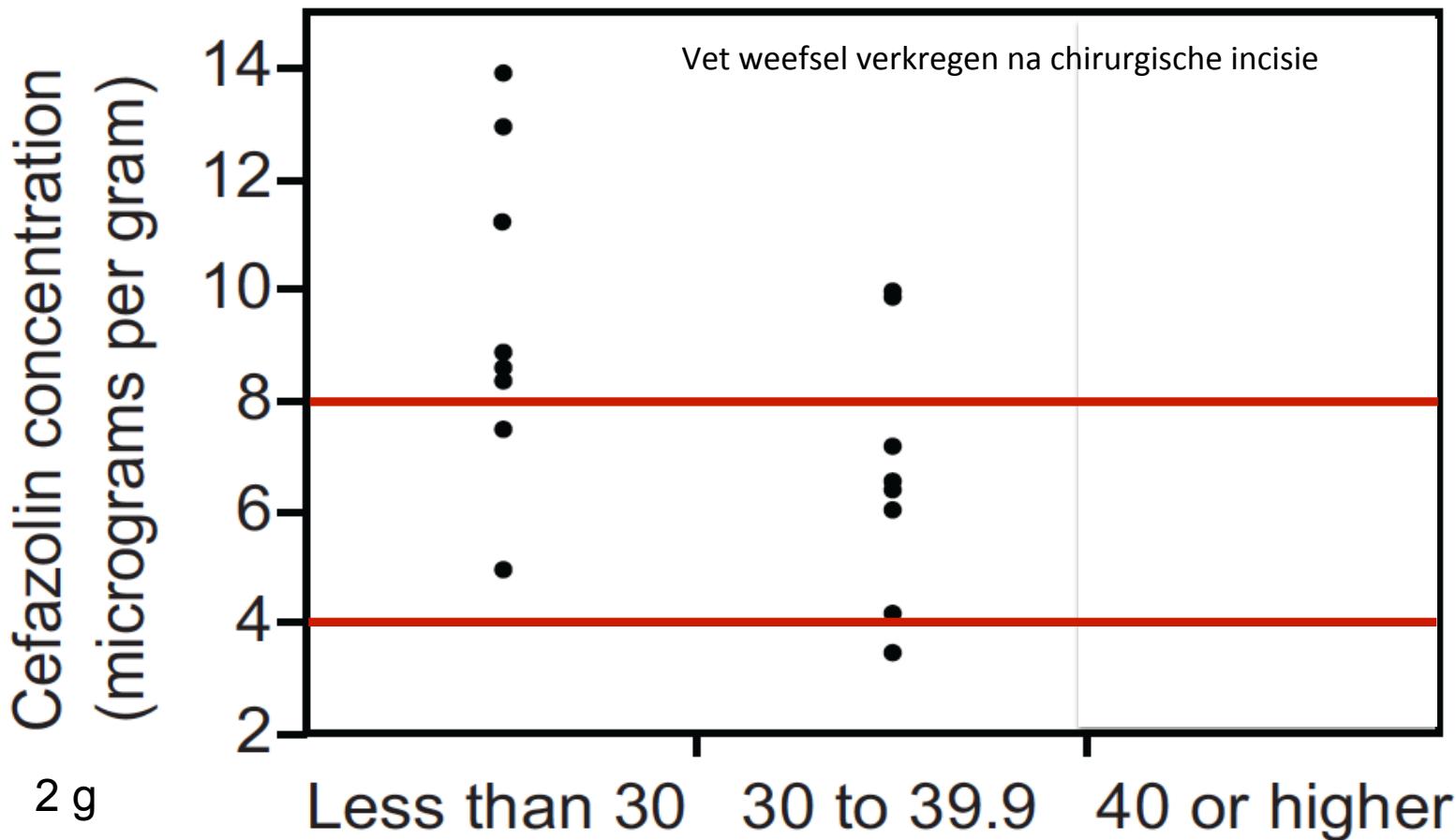
LAGERE CEFAZOLINE CONCENTRATIES IN VETWEEFSEL



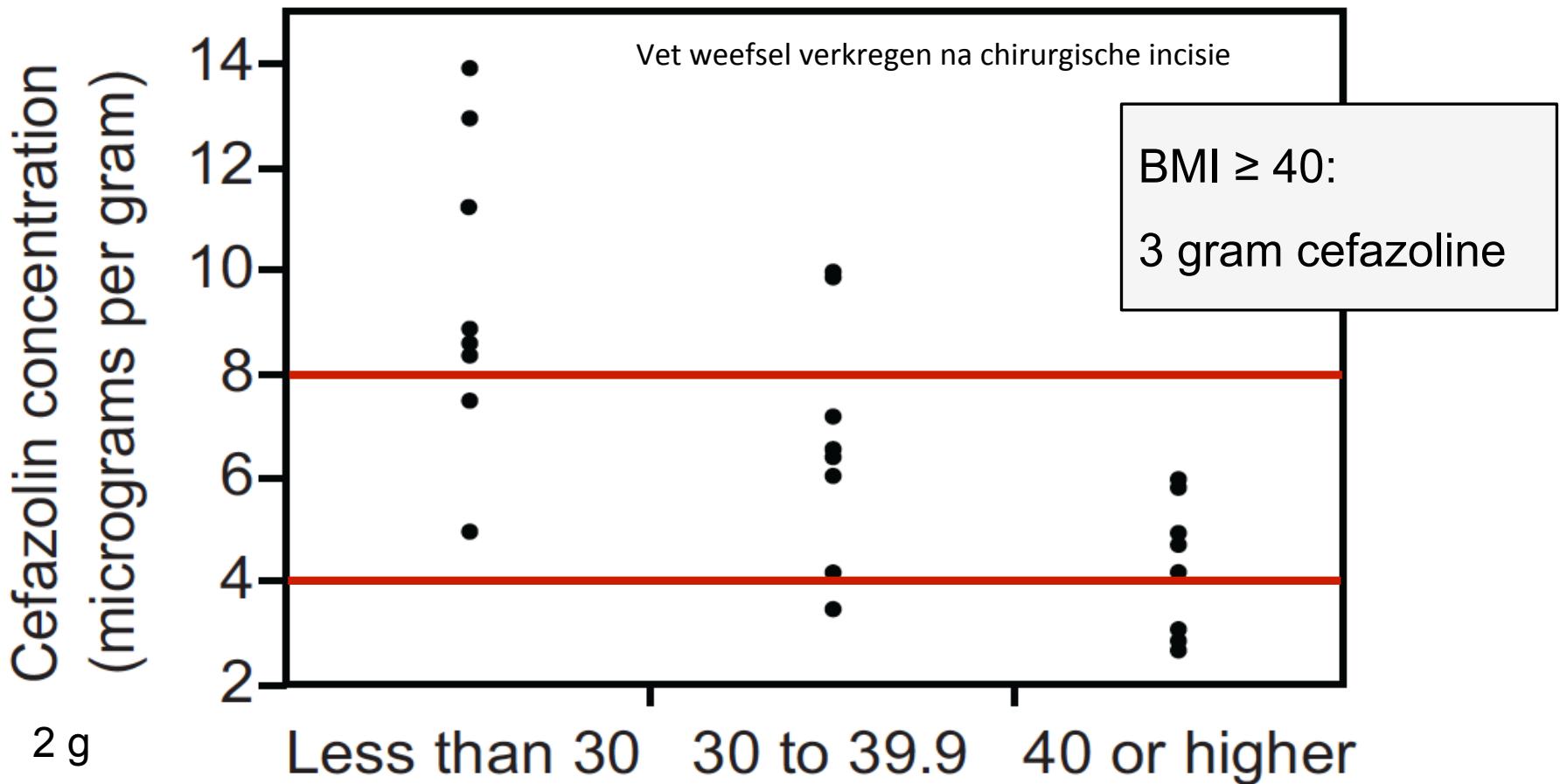
LAGERE CEFAZOLINE CONCENTRATIES IN VETWEEFSEL



LAGERE CEFAZOLINE CONCENTRATIES IN VETWEEFSEL



LAGERE CEFAZOLINE CONCENTRATIES IN VETWEEFSEL



CONCLUSIES

- Behoudens in tumor chirurgie, vooralsnog geen overtuigende aanwijzingen dat verlenging van de antibiotica profylaxe het risico op PJI verlaagt.
- Onvoldoende data dat antibiotica profylaxe aangepast of verbreed moet worden in hoog risico groepen.
- De dosering van antibiotica profylaxe moet worden geoptimaliseerd in obese patiënten.
- Andere preventie maatregelen belangrijker dan antibiotica profylaxe?