# Amputation in Patients with PAD with and without Diabetes: Insights from the EUCLID Trial

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### BACKGROUND

- Amputation is a major complication of peripheral artery disease (PAD)
- Multifactorial nature is increasingly recognized, particularly in those with concomitant diabetes mellitus (DM)
- Elucidating the drivers of amputation in PAD with and without DM may be important in developing strategies for prevention



#### WIfl Classification Wound Ischemia 0: No ulcer or gangrene Ankle Pressure/Toe Pressure 1: Small ulcer & no gangrene (mmHg) 2: Deep ulcer & toe gangrene 0: >100 / ≥60 3: Extensive ulcer/gangrene 1:70-100/40-59 2: 50-70 / 30-39 3: <50 / <30 foot Infection **0:No infection** 1:Mild: skin/subcut. 2:Moderate: deep to subcut. 3:Severe: systemic symptoms

# **M**ETHODS

- EUCLID randomized 13,885 patients with PAD. Investigators prospectively reported all amputations
- In this post-hoc analysis, amputations (major – ankle and above, minor – distal to ankle) were retrospectively adjudicated using safety data when available to characterize the drivers including infection, ischemia, or multifactorial
- Etiologies were evaluated by DM status at baseline





# **COHORT CHARACTERISTICS**

#### 415 patients (3% of total) underwent 533 amputations over a median of 30 months



\*10 amputations were unknown regarding major or minor, 3 in diabetics and 7 in non-diabetics

Characteristic	At least one	No
	amputation	amputation
Ν	415	13,470
Age, mean (SD), years	66 (8.7)	67 (8.4)
Female	21%	28%
Current/former smoker	68%	79%
Diabetes Mellitus	67%	38%
Insulin requirement	59%	35%
Oral agent alone	37%	57%
Diet control alone	4%	8%
HbA1c (%), mean (SD)	7.8 (2.1)	6.8 (8.4)
PAD Characteristics		
ABI, mean (SD)	0.64 (0.27)	0.72 (0.21)
Prior revascularization	62%	56%
Prior major amputation	11%	2%
Prior minor amputation	25%	4%
Rutherford Classification		
0. Asymptomatic	13%	19%
1-2. Mild/Mod claudication	39%	54%
3. Severe claudication	27%	23%
4. Rest pain	9%	3%
5. Minor tissue loss	9%	1%
6. Major tissue loss	3%	<1%

# TIMING OF AMPUTATIONS







# RESULTS

- 172 out of 533 non-traumatic amputations with sufficient documentation to determine drivers
- Ischemia was the primary driver overall (51%) followed by infection (27%) and multifactorial (22%)
- Primary driver varied by DM status





RESULTS

Diabetes No Diabetes 2% 6% 5% Major Amputations 9% 26% 43% 26% 83% 7% Minor Amputations 22% 46% 29% 78% 18%

Infection alone

Ischemia alone

Infection primary, ischemia contributing
Ischemia primary, infection contributing

 The etiology varied for major and minor with the former driven by ischemia (65%) and the latter driven by infection (59%)

 Infection was the predominant driver in patients with diabetes for both major (52%) and minor (64%) amputations

# LIMITATIONS & CONCLUSION

# LIMITATIONS

- Subgroup analysis of RCT
- Critical limb ischemia (CLI) underrepresented in overall clinical trial population
- Incomplete data for adjudication of amputation drivers

# CONCLUSION

- Amputations in PAD appear to have different primary drivers depending on concomitant Diabetes
- Infection may have a larger role in patients with Diabetes and ischemia in patients without Diabetes



