

Prevalence of Endogenous Hypercortisolism in Individuals With Resistant Hypertension Stratified by Kidney Function: Results From the MOMENTUM Study

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Speaker Disclosures

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Alynlam, Regeneron

Grants:

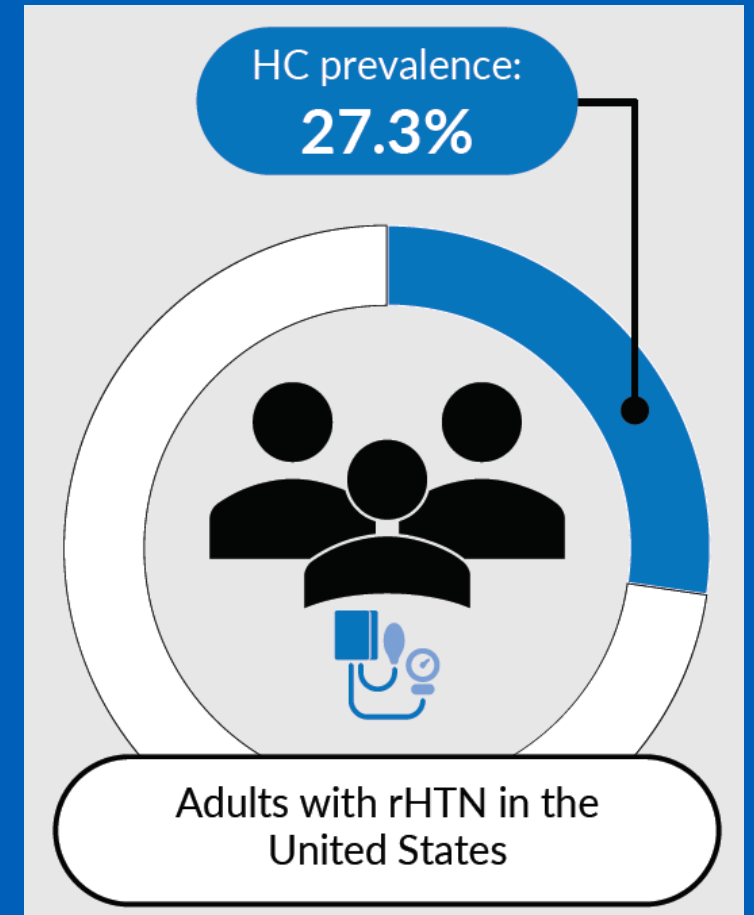
Astra Zeneca, Eli Lilly, Regeneron

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MOMENTUM Primary Result

- The first US-based trial to assess hypercortisolism prevalence in individuals with resistant hypertension (rHTN)
- Screening test used: 1-mg overnight DST with post-DST morning cortisol >1.8 mg/dL



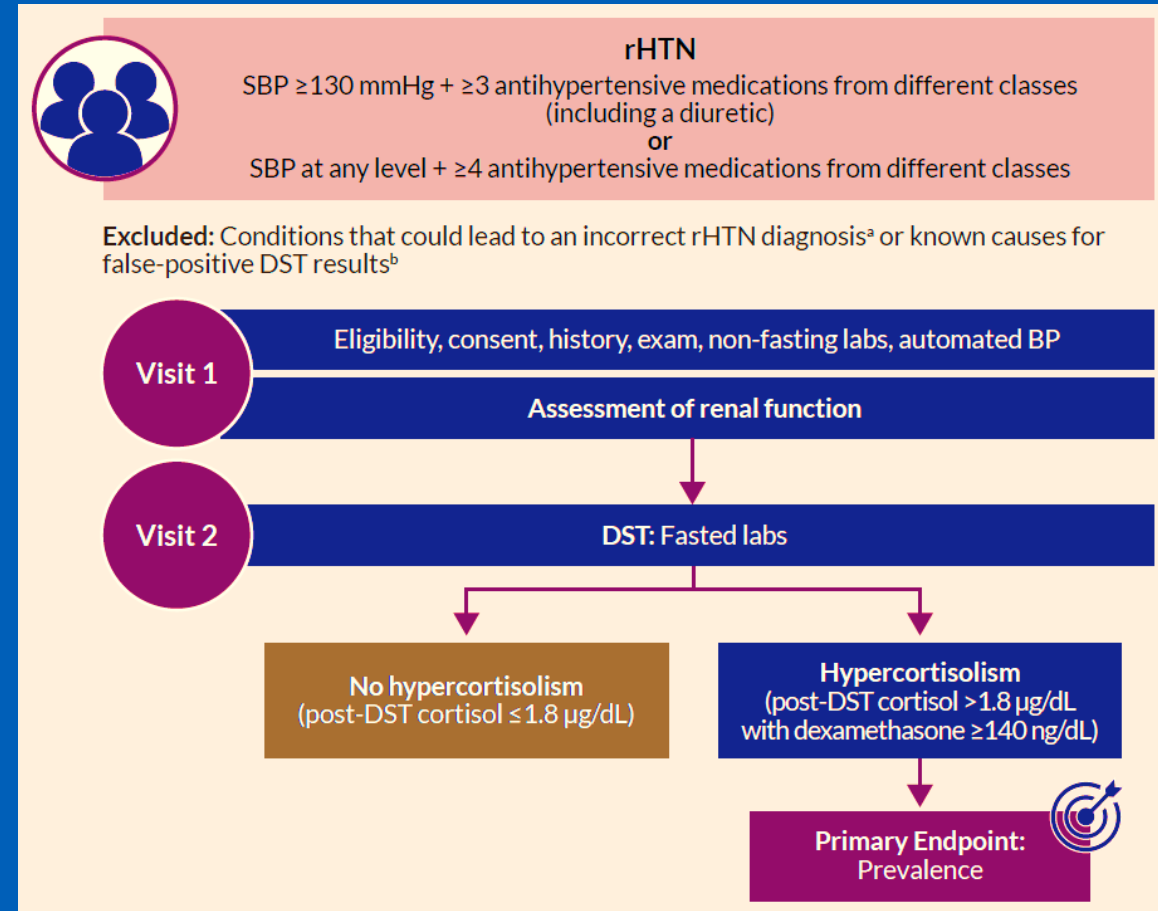
CKD & HC in People with rHTN

- Uremia is an insulin-resistant state, and hypercortisolism also causes insulin resistance^{1,2}
 - However, the relationship between CKD and hypercortisolism in rHTN is not well characterized
 - Many adverse effects associated with the uremic state (eg, insulin resistance, cardiometabolic complications, bone disease) are also characteristic of hypercortisolism, suggesting a potential pathophysiologic connection
- If eGFR is reduced, there is a prolongation of cortisol half-life³ and potential for cross-reactivity of cortisol metabolites in the DST immunoassay

► **We analyzed the frequency of post-DST cortisol >1.8 µg/dL and characteristics of participants in MOMENTUM by eGFR**

Methods: The MOMENTUM Trial (N=1,086)

- A US-based, multicenter, prospective, observational study
- Screened adults with rHTN per AHA criteria using the 1-mg overnight DST
- Causes of false-positive DSTs were excluded
- Cortisol measured by electrochemiluminescence immunoassay
 - The assay used to measure cortisol was not tested for the potential effect of low eGFR



Baseline Demographics & Characteristics Across eGFR Groups

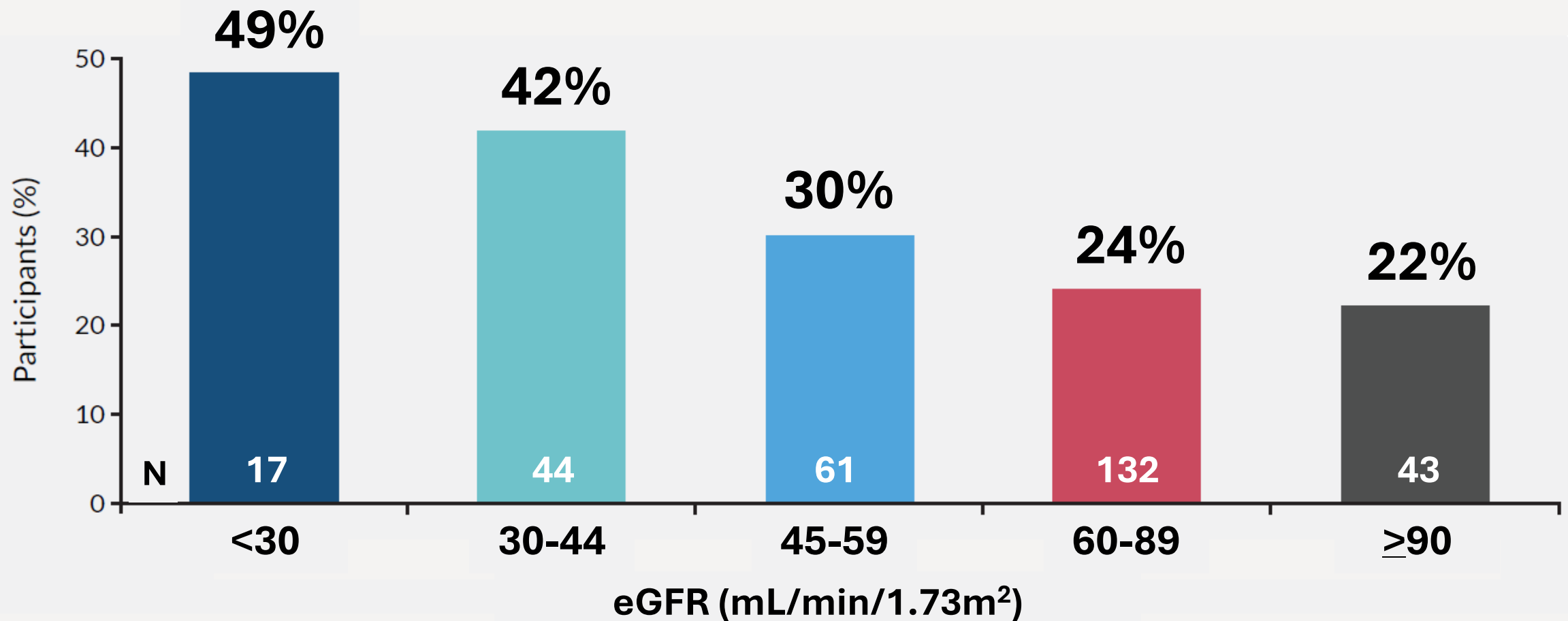
	eGFR <45 mL/min/1.73 m ² (n=140) ^a	eGFR 45–59 mL/min/1.73 m ² (n=203)	eGFR ≥60 mL/min/1.73 m ² (n=743)
Age, years	69.5	69.9	63.3
Female, (%)	44.3	55.7	51.1
Race, (%)			
Asian	1.4	5.4	3.0
Black or African American	30.0	31.0	39.2
White	66.4	62.1	54.0
Other ^b	2.1	1.5	3.9
Ethnicity not Hispanic/Latino, (%)	75.0	81.8	70.5
Weight, kg	92.2	93.2	95.1
BMI, kg/m ²	31.8	33.3	33.3
Waist circumference, cm	109.9	108.3	108.5
SBP, mmHg	137.4	137.8	141.9
DBP, mmHg	79.9	79.9	86.1

Post-DST Cortisol & ACTH Across eGFR Groups

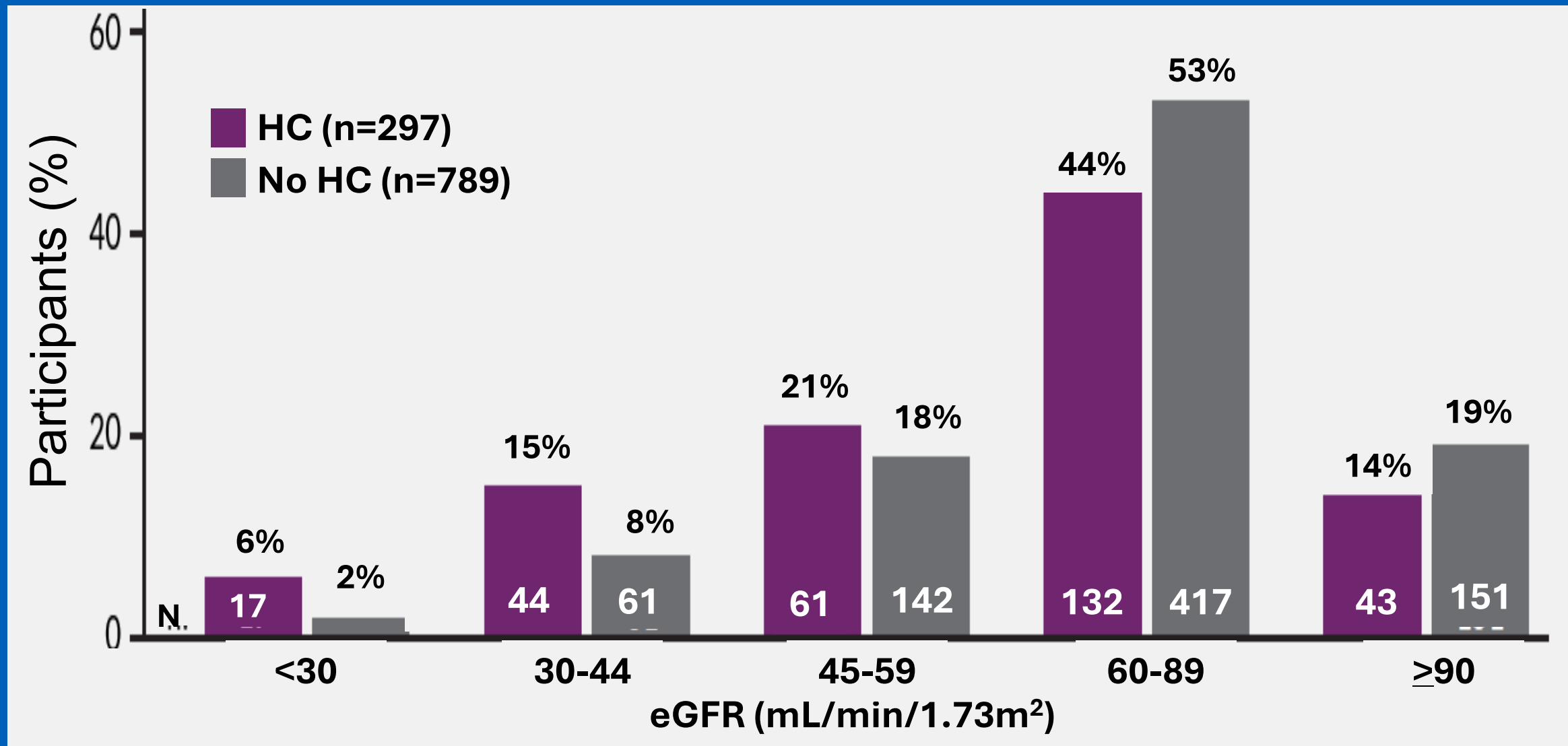
- ACTH was similar across eGFR groups (overall and in those with hypercortisolism)
- Post-DST cortisol was higher in those with lower eGFR; mean values were higher in MOMENTUM vs Garg¹

	eGFR <45 mL/min/1.73 m ² (n=140)	eGFR 45–59 mL/min/1.73 m ² (n=203)	eGFR ≥60 mL/min/1.73 m ² (n=743)
Post-DST cortisol, mg/dL, mean			
Overall	2.4	2.1	1.9
Hypercortisolism	3.9 (44%)	4.1 (30%)	4.4 (24%)
Post-DST ACTH, ng/L, mean			
Overall	5.3	4.9	4.9
Hypercortisolism	7.3	7.5	8.5
8 AM ACTH in participants with HC, ng/L	23.3	18.8	21.9

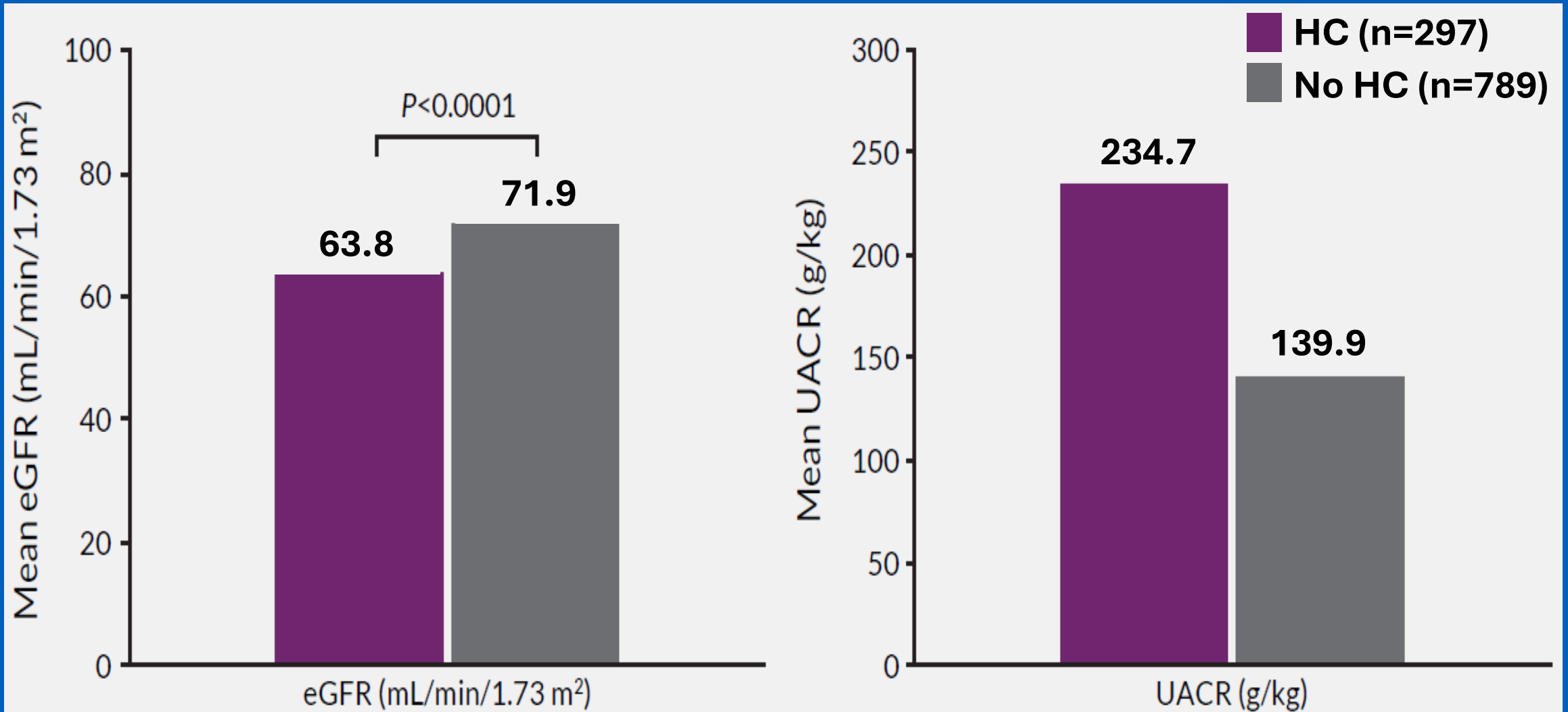
Relationship between Percentage of Participants With Post-DST Cortisol $>1.8 \mu\text{g/dL}$ and eGFR



Percent of Participants With and Without Hypercortisolism by eGFR Status

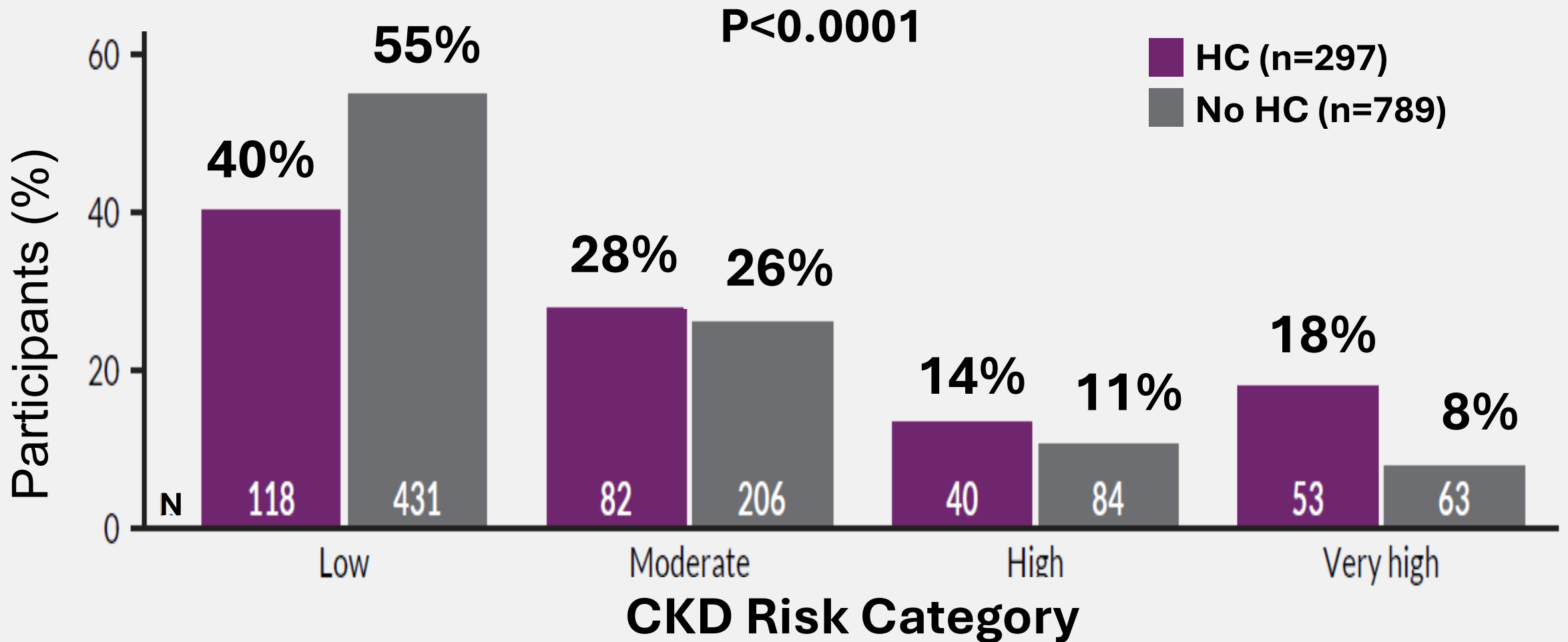


Markers of Kidney Function (eGFR and UACR) in Participants with and Without Hypercortisolism



eGFR, estimated glomerular filtration rate; HC, hypercortisolism; UACR, urine albumin-to-creatinine ratio.

Percentage of Participants With and Without Hypercortisolism by CKD Progression Risk Category



Summary & Conclusions

- In the overall MOMENTUM population, **hypercortisolism prevalence was 27.3%**
- **Hypercortisolism prevalence increased with lower eGFR**
- MOMENTUM could not determine whether this association reflects a true disruption in the HPA axis or a prolongation in the clearance of cortisol
- **ACTH was similar and in the low normal range across eGFR groups, making a disturbance in the hypothalamic-pituitary axis an unlikely cause of the hypercortisolism**
- UACR trended higher in people with hypercortisolism, as did CKD progression risk

▶ **These results support the need for targeted hypercortisolism screening in people with rHTN**