

Epicardial and pericardial adiposity without myocardial steatosis in Cushing's syndrome

Peter Wolf^{1,2}, Benjamin Marty³, Khaoula Bouazizi^{4,5}, Nadjia Kachenoura^{4,5}, Céline Piedvache⁶, Anne Blanchard⁷, Sylvie Salenave¹, Mikaël Prigent⁴, Christel Jublanc⁸, Christiane Ajzenberg⁹, Céline Droumaguet⁹, Jacques Young¹, Anne-Lise Lecoq¹, Emmanuelle Kuhn¹, Hélène Agostini⁶, Séverine Trabado¹⁰, Pierre G. Carlier³, Bruno Fève¹¹, Alban Redheuil^{5,12}, Philippe Chanson¹, Peter Kamenický¹

¹ Université Paris-Saclay, Inserm, Physiologie et Physiopathologie Endocrinienne, Assistance Publique-Hôpitaux de Paris, Hôpital Bicêtre, Service d'Endocrinologie et des Maladies de la Reproduction, Centre de Référence des Maladies Rares de l'Hypophyse, 94275 Le Kremlin-Bicêtre, France; ² Medical University of Vienna, Department of Internal Medicine III, Division of Endocrinology and Metabolism, 1090 Vienna, Austria; ³ Institut de Myologie, CEA, Laboratoire de Résonance Magnétique Nucléaire, 75013 Paris, France; ⁴ Institut de Cardiometabolisme et Nutrition (ICAN), 75013 Paris, France; ⁵ Sorbonne Université, CNRS, Inserm, Laboratoire d'Imagerie Biomédicale, LIB, 75006 Paris, France; ⁶ Assistance Publique-Hôpitaux de Paris, Hôpital Bicêtre, Unité de Recherche Clinique, 94275 Le Kremlin-Bicêtre, France; ⁷ Université de Paris, Assistance Publique des Hôpitaux de Paris, Hôpital Européen Georges-Pompidou, Centre d'Investigations Cliniques, Inserm CIC1318 et UMR 1138, 75013 Paris, France; ⁸ Assistance Publique-Hôpitaux de Paris, Hôpital Pitié-Salpêtrière, Service d'Endocrinologie-Métabolisme, 75013 Paris, France; ⁹ Assistance Publique-Hôpitaux de Paris, Hôpital Henri-Mondor, Service de Médecine Interne, 94000 Créteil, France; ¹⁰ Assistance Publique-Hôpitaux de Paris, Hôpital Bicêtre, Service de Génétique Moléculaire, Pharmacogénétique et Hormonologie, 94275 Le Kremlin-Bicêtre, France; ¹¹ Assistance Publique-Hôpitaux de Paris, Hôpital Saint-Antoine, Service d'Endocrinologie et Métabolisme, Centre de Référence des Maladies Rares de l'Insulino-Scrétion et de l'Insulino-Sensibilité (PRYSIS), Sorbonne Université, Inserm, Centre de Recherche Saint-Antoine, UMR-S938, IHU ICAN, 75012 Paris, France; ¹² Assistance Publique-Hôpitaux de Paris, Hôpital Pitié-Salpêtrière, Unité d'Imagerie Cardiovasculaire et Thoracique, APHP.SU, 75013 Paris, France.

Objective:

Cardiovascular disease is the leading cause of death in patients with Cushing's syndrome. Cortisol excess and adverse metabolic profile could increase cardiac fat, which can subsequently impair cardiac structure and function. We aimed to evaluate cardiac fat mass and distribution in patients with Cushing's syndrome.

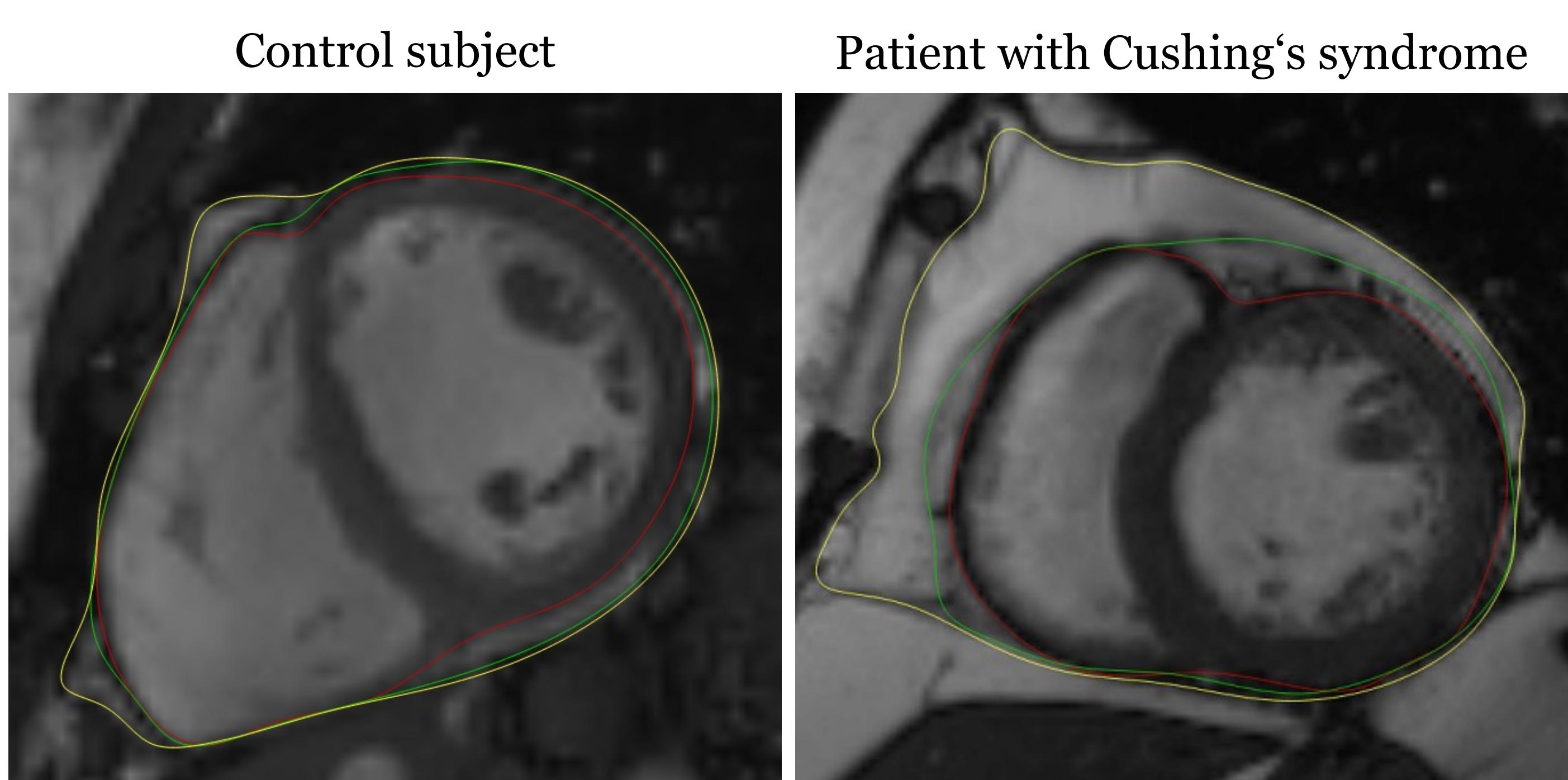
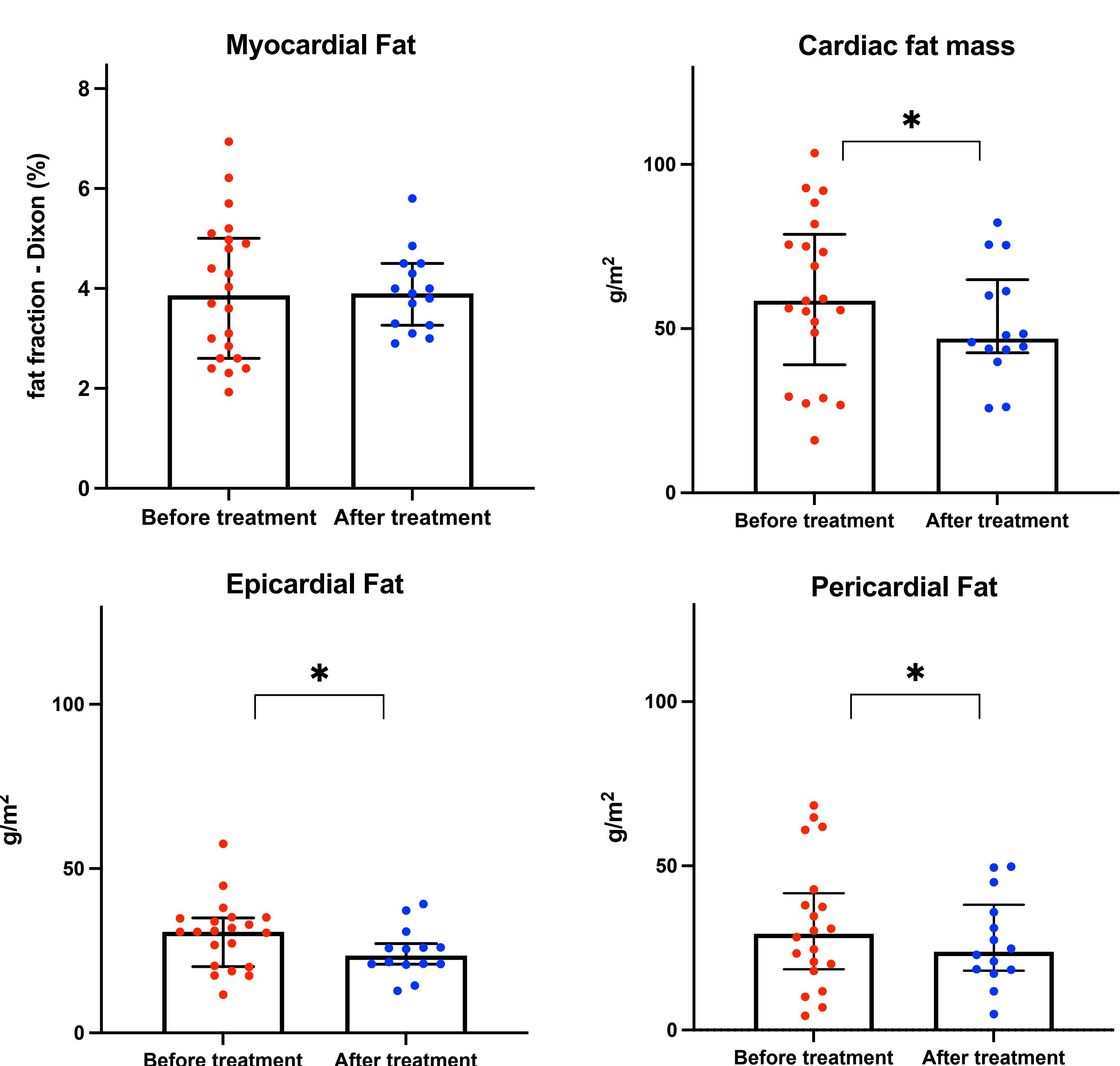
Methods:

- prospective cross-sectional study in 23 patients with Cushing's syndrome and 27 control subjects of comparable age, sex and body-mass-index
- cardiac magnetic resonance imaging and proton spectroscopy before and after biochemical disease remission
- myocardial fat measured by the Dixon method was the main outcome measure
- intramyocardial triglyceride/water ratio measured by spectroscopy and epicardial and pericardial fat volumes were secondary outcome measures

	Cushing's syndrome	Control group	p-value
Age (yrs)	38 (31; 52)	41 (33; 55)	0.53
Sex (Female, %)	20 (75%)	20 (75%)	NA
Body mass index (kg/m ²)	28 (24; 34)	27 (24; 32)	0.89
Systolic blood pressure (mmHg)	133 (123; 145)	120 (114; 136)	0.009
Diastolic blood pressure (mmHg)	83 (75; 90)	72 (65; 78)	0.002
HbA1c (%)	6.4 (5.7; 6.6)	6 (5.4; 6.1)	0.031
24h urinary free cortisol (ug/24h)	346 (198; 406)	29 (13; 43.3)	<0.001

Results:

	Cushing's syndrome	Control group	p-value
End-diastolic volume index (mL/m ²)	67.5 (64.9 ; 80.8)	81.3 (73.8 ; 91.6)	0.0209
End-systolic volume index (mL/m ²)	27.3 (24.8 ; 31.6)	33.1 (27.9 ; 35.9)	0.11
Ejection fraction (%)	61.8 (56.6 ; 66.3)	61.1 (59.1 ; 62.3)	0.73
Stroke volume index (mL/m ²)	41.7 (37.1 ; 49.3)	49.4 (45.1 ; 55.6)	0.0169
Left ventricular mass index (g/m ²)	56.3 (49.3 ; 59.3)	52.1 (45.3 ; 57.4)	0.13
End-diastolic mass / volume	0.77 (0.7 ; 0.9)	0.63 (0.6; 0.72)	0.0014
Myocardial fat - Dixon (%)	3.9 (2.6 ; 4.9)	3.9 (3.1 ; 4.3)	0.80
Myocardial fat - spectroscopy (%)	1.8 (0.7 ; 2.9)	2.2 (1.4 ; 2.9)	0.41
Cardiac adipose tissue index (g/m ²)	58.5 (48.8 ; 87.9)	30.4 (21.9 ; 42.4)	<.001
Epicardial adipose tissue index (g/m ²)	30.8 (20.4 ; 34.8)	17.2 (13.1 ; 23.5)	<.001
Pericardial adipose tissue index (g/m ²)	28.3 (17.9 ; 38.0)	11.4 (7.5 ; 19.4)	0.0035



Discussion:

Intramyocardial fat stores are not increased in patients with Cushing's syndrome, despite highly prevalent metabolic syndrome, suggesting increased cortisol-mediated lipid consumption. Cushing's syndrome is associated with marked accumulation of epicardial and pericardial fat. Epicardial adiposity may exert paracrine proinflammatory effects promoting cardiomyopathy.