

Definizione della sindrome fibromialgica, aspetti epidemiologici e collocazione nosografica



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Diagnostic and therapeutic care pathway for fibromyalgia

P. Sarzi-Puttini¹, V. Giorgi¹, F. Atzeni², R. Gorla³, E. Kosek^{4,5}, E.H. Choy⁶, L. Bazzichi⁷,
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M. Di Carlo²⁵, G. Cassisi²⁶, A. Batticciotto²⁷

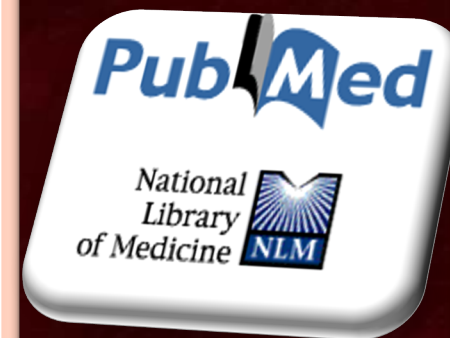
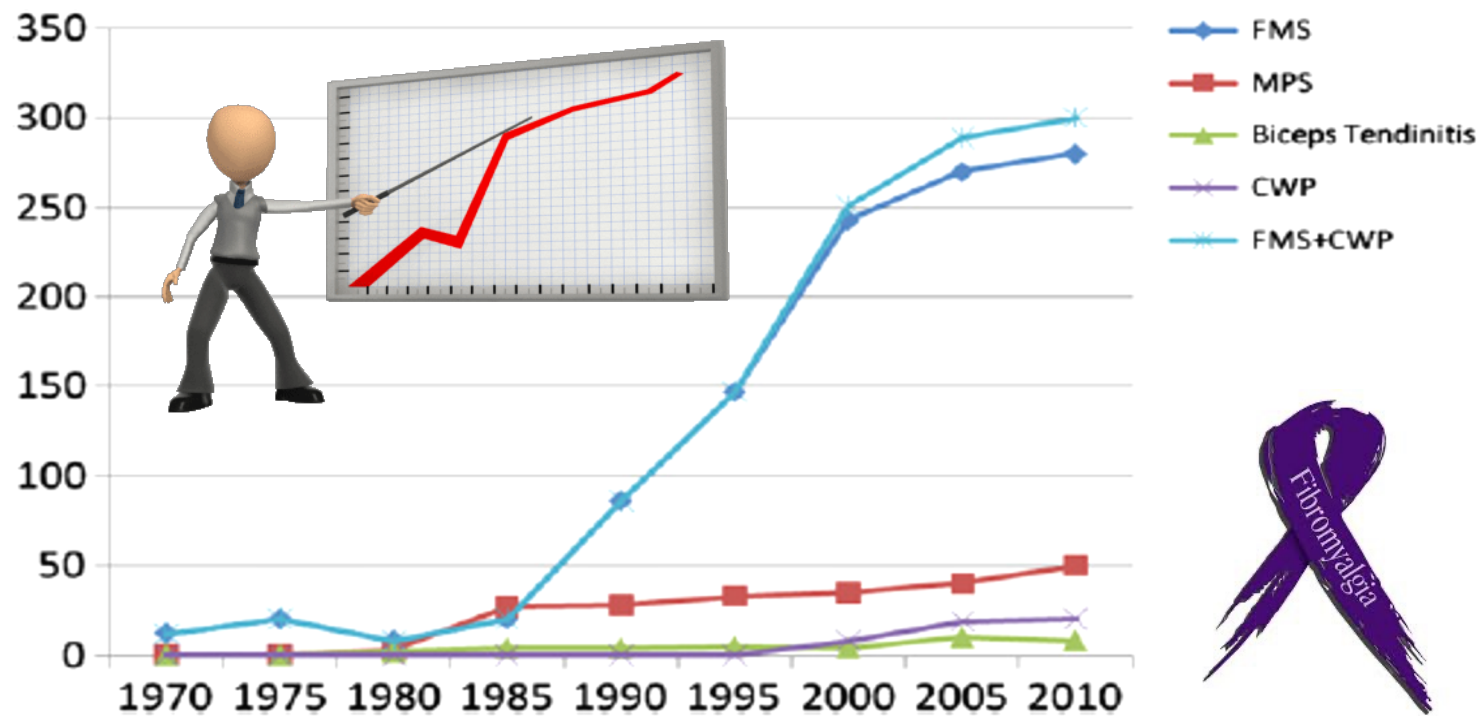
Fibromyalgia position paper

P. Sarzi-Puttini¹, V. Giorgi¹, F. Atzeni², R. Gorla³, E. Kosek^{4,5}, E.H. Choy⁶, L. Bazzichi⁷,
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La fibromialgia è una sindrome da sensibilizzazione centrale caratterizzata dalla disfunzione dei neuro-circuiti preposti alla percezione, trasmissione e processazione delle afferenze nocicettive, con prevalente estrinsecazione del dolore a livello dell'apparato muscoloscheletrico.

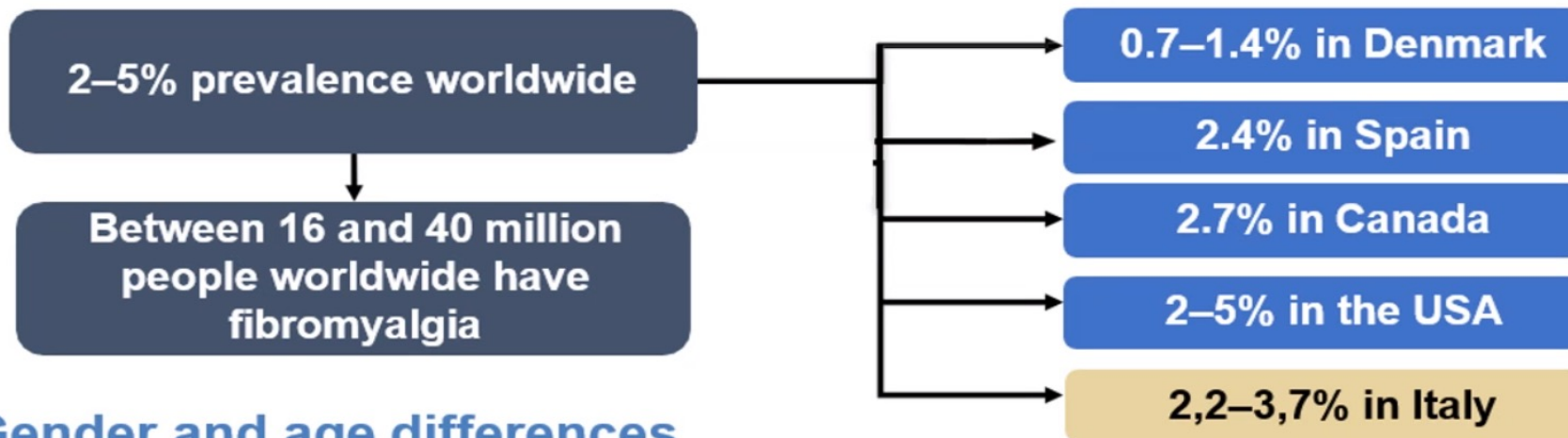
Oltre al dolore possono essere presenti molteplici sintomi di accompagnamento (astenia, disturbi del sonno, dolori addominali...) comuni ad altre sindromi da sensibilizzazione centrale.

Numbers of Medline-referenced publications during nine 12-month periods at five year intervals from 1970 to 2010. The data illustrate a dramatic increase in publications about FMS beginning in 1990 when the American College of Rheumatology approved Classification Criteria for FMS.



Epidemiology of Fibromyalgia

Fibromyalgia is the most common chronic widespread pain condition



Gender and age differences

- **This condition affects women 10 times more frequently than men**
- **Majority of patients are aged 35–60 years (working age)**

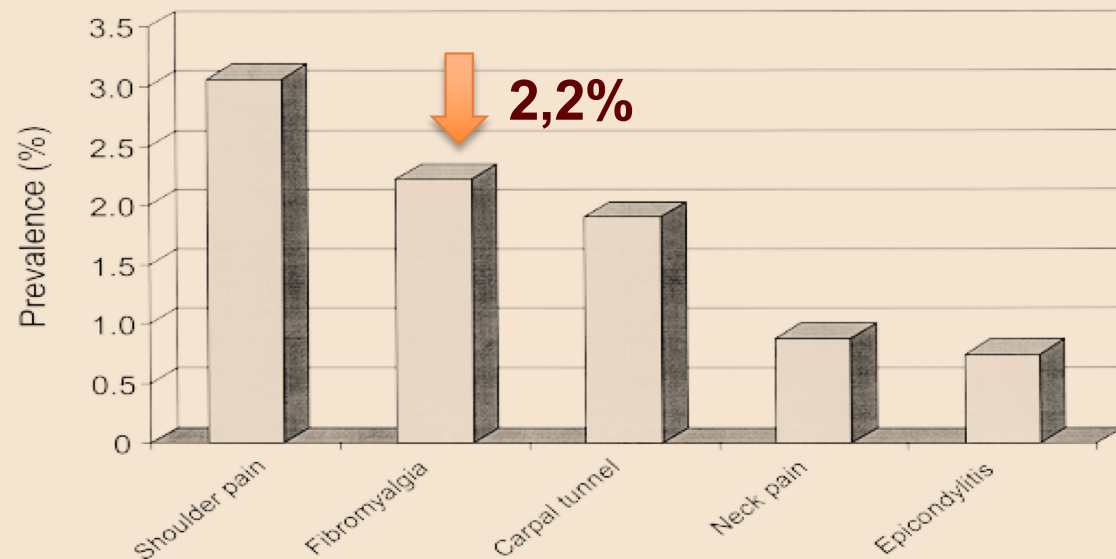
Prevalence of musculoskeletal conditions in an Italian population sample: results of a regional community-based study.

I. The MAPPING study

F. Salaffi, R. De Angelis, W. Grassi, on behalf of the MArche Pain Prevalence INvestigation Group (MAPPING) Study*









Clinical and Experimental Rheumatology 2005; 23: 819-828.

Prevalence in the total target adult population of the soft-tissue disorders: shoulder pain, fibromyalgia, carpal tunnel syndrome, neck pain, and lateral epicondylitis.



The estimated rate of disease prevalence for fibromyalgia was 2.22% (95% CI 1.36-3.19)

Definition of fibromyalgia severity: findings from a cross-sectional survey of 2339 Italian patients

Fausto Salaffi ¹, Marco Di Carlo ¹, Laura Bazzichi², Fabiola Atzeni ³, Marcello Govoni⁴, Giovanni Biasi⁵, Manuela Di Franco⁶, Flavio Mozzani⁷, Elisa Gremese⁸, Lorenzo Dagna⁹, Alberto Batticciotto¹⁰, Fabio Fischetti¹¹, Roberto Giacomelli¹², Serena Guiducci¹³, Giuliana Guggino ¹⁴, Mario Bentivegna¹⁵, Roberto Gerli ¹⁶, Carlo Salvarani¹⁷, Gianluigi Bajocchi¹⁸, Marco Ghini¹⁹, Florenzo Iannone ²⁰, Valeria Giorgi²¹, Sonia Farah ¹, Mariateresa Cirillo³, Sara Bonazza⁴, Stefano Barbagli⁵, Chiara Gioia⁶, Daniele Santilli⁷, Annunziata Capacci⁸, Giulio Cavalli⁹, Francesco Carubbi¹², Francesca Nacci¹³, Ilenia Riccucci¹⁶, Luigi Sinigaglia²², Maurizio Masullo²³, Bianca Maria Polizzi²³, Maurizio Cutolo²⁴ and Piercarlo Sarzi-Puttini ²¹

FIQR, FAS 2019mod and PDS cut-off values for FM severity states

Rheumatology 2020;0:1-9

FM severity state	Cut-off values		
	FIQR	FAS 2019mod	PDS
Remission	≤23	≤12	≤5
Mild	>23 and ≤40	>12 and ≤20	>5 and ≤15
Moderate	>40 and ≤63	>20 and ≤28	>15 and ≤20
Severe	>63 and ≤82	>28 and ≤33	>20 and ≤25
Very severe	>82	>33	>25

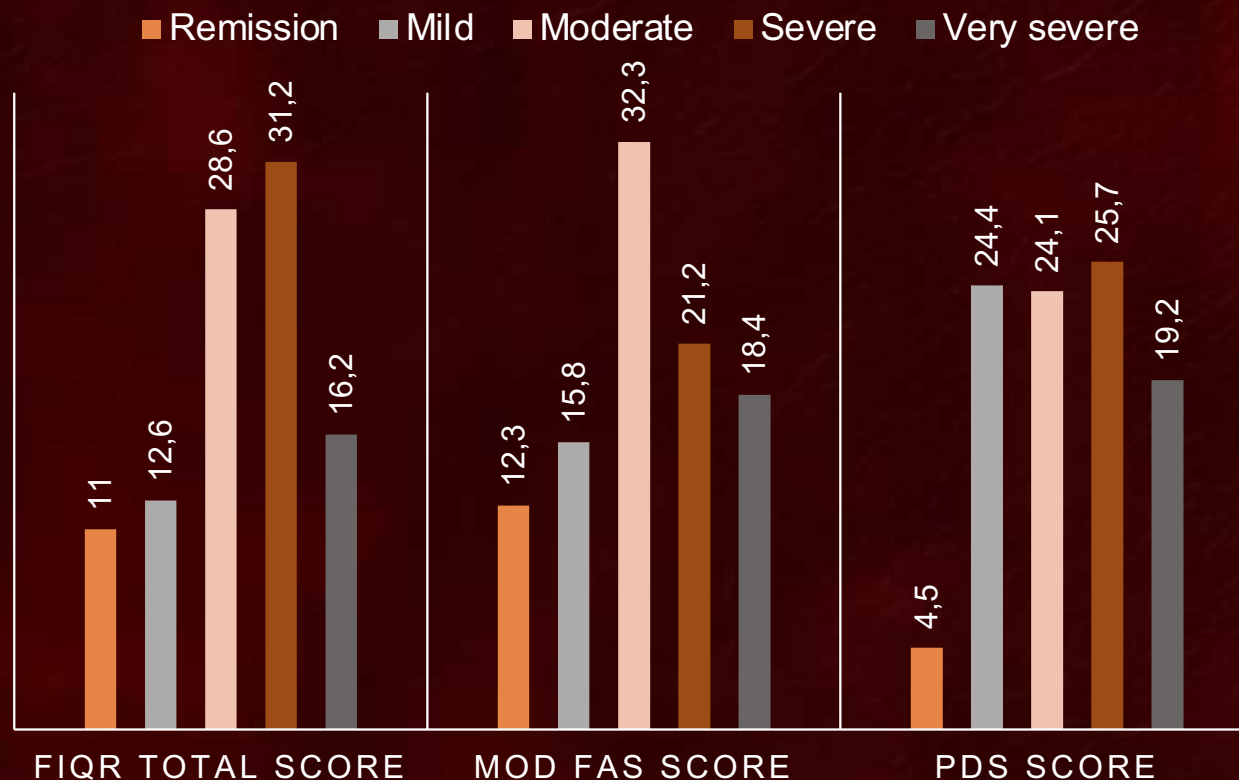
FAS 2019mod: modified Fibromyalgia Assessment Status; FIQR: revised Fibromyalgia Impact Questionnaire; PDS: Polysymptomatic Distress Scale.

CUTOFF DISTRIBUTION

The application of the calculated cutoff points for FIQR, Mod FAS and PDS showed the following distribution: remission in 11.0%, 12.3% and 4.5% of the patients, mild severity in 12.6% 15.8% and 26.4%, moderate severity in 28.6%, 32.3% and 24.1%, high severity in 31.2%, 21.2% and 25.7% and very high severity in 16.2%, 18.4% and 19.2% of the patients, respectively.

According to the FIQR cut-off points, 47.4% of the patients resulted in high or very high disease severity, while according to the Mod FAS and PDS cut-off points these percentages were 39.6% and 44.9%, respectively.

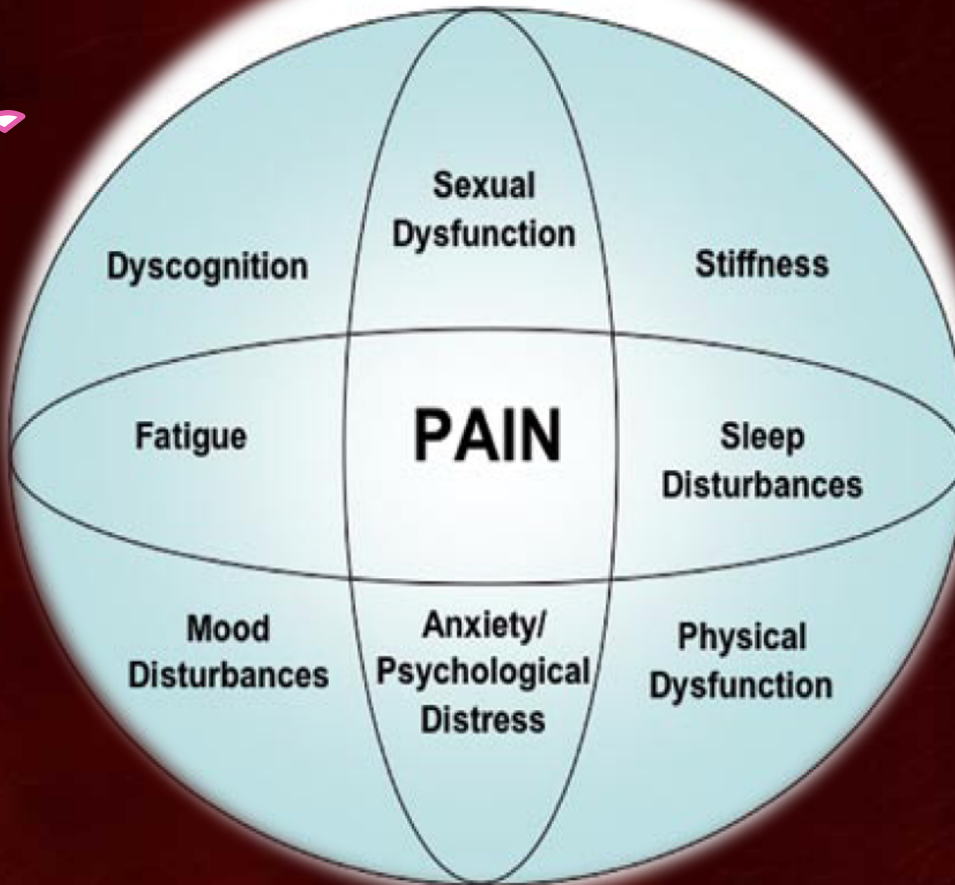
F Salaffi et al. 2020



Questionnaires	Remission	Mild	Moderate	Severe	Very severe
FIQR total score	11	12,6	28,6	31,2	16,2
Mod FAS score	12,3	15,8	32,3	21,2	18,4
PDS score	4,5	24,4	24,1	25,7	19,2



PAIN



Fibromyalgia domains



Preliminary identification of key clinical domains for outcome evaluation in fibromyalgia using Delphi methods: the Italian experience

F. Salaffi¹, A. Ciapetti¹, P. Sarzi Puttini², F. Atzeni², C. Iannuccelli³, M. Di Franco³, M. Cazzola⁴, L. Bazzichi⁵

Reumatismo, 2012; 64 (1): 28-35

Domini classificati per rilevanza dai pazienti

Dominio	Items	Frequency	Mean importance (MI)	Frequency importance product (FIP)
1. Dolore	Dolore o disagio fisico; articolazioni dolenti; dolorabilità alla palpazione	97.3	2.9	282.2
2. Fatica	Stanchezza; scarsa energia	93.6	2.7	252.7
3. Qualità del sonno	Difficoltà ad addormentarsi; insonnia; risvegli frequenti	90.1	2.6	234.3
4. Funzione multidimensionale	Difficoltà nei movimenti, nel camminare o svolgere esercizi; difficoltà nello svolgere normali attività, compromissione dell'attività lavorativa, scolastica ed impatto nella vita quotidiana	89,8	2.6	233,5
5. Depressione	Sentirsi tristi, demotivati, pessimisti, isolati, svogliati	81.5	2.4	195.6
6. Sensibilità a stimoli esterni	Sensibilità a suoni, luci, odori e/o al freddo	78.3	2.4	187.9
7. Ansia	Sentirsi frustrati; essere preoccupati; avere paura	76.1	2.3	175.0
8. Disturbi cognitivi	Difficoltà nel ricordare o pensare; perdita di memoria; difficoltà nel concentrarsi	74.7	2.1	156.9



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Reumatismo, 2012; 64 (1): 28-35

Domini classificati per rilevanza dai clinici.

Dominio	Frequency	Mean importance (MI)	Frequency importance product (FIP)
1. Dolore	79,0	2,85	225,2
2. Fatica	78,6	2,65	208,3
3. Disturbi del sonno	74,6	2,70	201,4
4. Salute-qualità della vita (HRQL)	73,8	2,60	191,9
5. Depressione	70,2	2,60	182,5
6. Ansia	71,8	2,50	179,5
7. Problemi di memoria e di concentrazione	69,1	2,40	165,8
8. Tender points	60,2	2,20	126,4



The IASP classification of chronic pain for *ICD-11*: chronic primary pain

PAIN 160 (2019) 28–37

Michael Nicholas^a, Johan W.S. Vlaeyen^{b,c,d}, Winfried Rief^e, Antonia Barke^e, Qasim Aziz^f, Rafael Benoliel^g, Milton Cohen^h, Stefan Eversⁱ, Maria Adele Giamberardino^j, Andreas Goebel^k, Beatrice Korwisi^e, Serge Perrot^l, Peter Svensson^{m,n}, Shuu-Jiun Wang^{o,p}, Rolf-Detlef Treede^{q,*}, The IASP Taskforce for the Classification of Chronic Pain

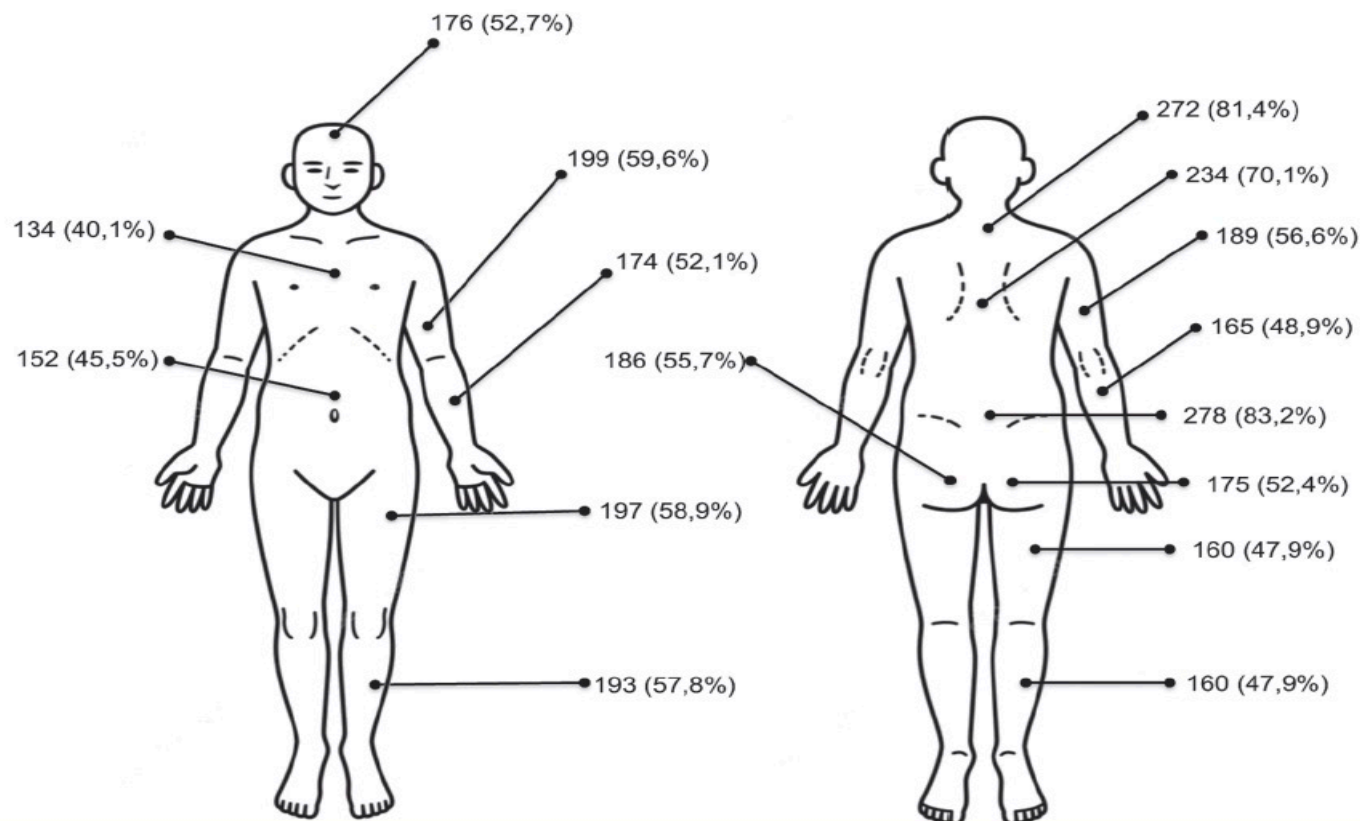
Chronic primary pain is chosen when pain has persisted for more than 3 months and is associated with significant emotional distress and/or functional disability, and the pain is not better accounted for by another condition.



Chronic pain was defined as pain that lasts or recurs for longer than 3 months.

Fausto Salaffi¹
Flavio Mozzani²
Antonella Draghessi¹
Fabiola Atzeni³
Rosita Catellani²
Alessandro Ciapetti⁴
Marco Di Carlo¹
Piercarlo Sarzi-Puttini⁵

Identifying the symptom and functional domains in patients with fibromyalgia: results of a cross-sectional Internet-based survey in Italy



Il paziente descrive il dolore con espressioni del tipo “mi fa male dappertutto”, Si tratta di un dolore “centrale”, che non ha una localizzazione ed un’entità costante, ma migra e può aumentare o diminuire durante l’arco della giornata

Criteri diagnostici proposti dall'American College of Rheumatology (ACR) del 2016 per la diagnosi di fibromialgia

Secondo tale proposta, per la diagnosi di FM devono essere soddisfatti contemporaneamente 3 criteri:

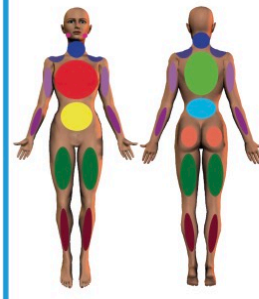
- dolore diffuso in specifiche aree e regioni del corpo;
- presenza di sintomi caratteristici (astenia, sonno non ristoratore, problemi cognitivi, emicrania, dolore/crampi addominali, depressione) che compromettono la vita quotidiana;
- durata della sintomatologia pari ad almeno 3 mesi

F. Salaffi et al. (2019) www.rheumalab.it

Indice di Diffusione del Dolore (Widespread Pain Index-WPI)

(1 punto per ogni casella spuntata; Range di punteggio: 0-19)

1- Indicare se si è provato dolore o indolenzimento nel corso degli ultimi 7 giorni, nelle aree riportate in figura. Spuntare le caselle in corrispondenza di ogni area interessata dal dolore o dall'indolenzimento.



<input type="checkbox"/> Cingolo scapolare sinistro	<input type="checkbox"/> Anca (gluteo, trocantere) sinistra	<input type="checkbox"/> Mascella sinistra
<input type="checkbox"/> Cingolo scapolare destro	<input type="checkbox"/> Anca (gluteo, trocantere) destra	<input type="checkbox"/> Mascella destra
<input type="checkbox"/> Braccio sinistro	<input type="checkbox"/> Coscia sinistra	<input type="checkbox"/> Torace
<input type="checkbox"/> Braccio destro	<input type="checkbox"/> Coscia destra	<input type="checkbox"/> Addome
<input type="checkbox"/> Avambraccio sinistro	<input type="checkbox"/> Gamba sinistra	<input type="checkbox"/> Area Dorsale
<input type="checkbox"/> Avambraccio destro	<input type="checkbox"/> Gamba destra	<input type="checkbox"/> Area Lombare
<input type="checkbox"/> Collo	Punteggio: ____ /19 aree	

Indice di Gravità dei Sintomi (Symptom Severity Scale-SSS)

(Range di punteggio: 0-12)

2- Per ognuno dei sintomi elencati esprimere la loro gravità nel corso degli ultimi 7 giorni, utilizzando la scala di seguito riportata.

0= *Nessun problema*

1= *Problema lieve*: generalmente lieve e intermittente

2= *Problema moderato*: disturbo di considerevole entità; presente spesso e/o di grado moderato

3= *Problema grave*: continuo, che ostacola le attività della vita quotidiana

	0	1	2	3
A. Astenia, spossatezza	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
B. Difficoltà di concentrazione, perdita di memoria	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
C. Stanchezza al risveglio, sonno non ristoratore	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

3- Indicare la presenza o assenza dei 3 sintomi (D, E, F) negli ultimi sei mesi

	Assente	Presente
D. Dolori o crampi addominali	<input type="checkbox"/>	<input type="checkbox"/>
E. Depressione	<input type="checkbox"/>	<input type="checkbox"/>
F. Emicrania, cefalea	<input type="checkbox"/>	<input type="checkbox"/>

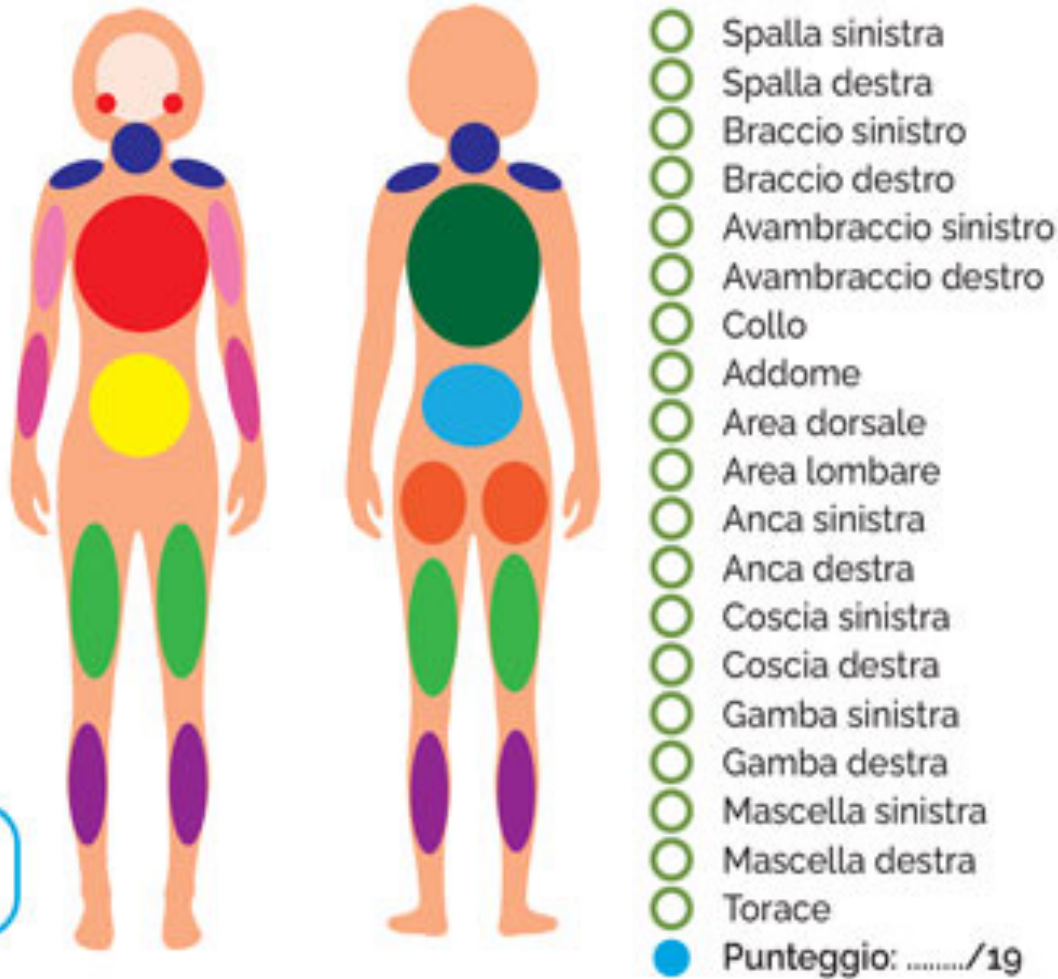
Criteri aggiuntivi (senza punteggio)

4- I sintomi delle domande 2 e 3 e il dolore diffuso sono stati avvertiti per almeno tre mesi? Sì No

5- E' presente una patologia che può motivare il dolore avvertito? Sì No

Punteggio Totale ____ /31

Indichi nelle rispettive caselle se ha provato dolore nelle aree riportate nella figura qui sotto nel corso degli ultimi 7 giorni



La combinazione del punteggio della scala di severità dei sintomi (SS) e dell'indice del dolore diffuso (WPI) definisce la diagnosi di fibromialgia (WPI \geq 7 e SS \geq 5) oppure (WPI 3-6 e SS \geq 9).

Fonte: Salaffi F, Farah S, <https://sindromefibromialgica.it/>

Diagnosis of fibromyalgia: comparison of the 2011/2016 ACR and AAPT criteria and validation of the modified Fibromyalgia Assessment Status

Fausto Salaffi¹, Marco Di Carlo ¹, Sonia Farah¹, Fabiola Atzeni², Dan Buskila³, Jacob N. Ablin⁴, Winfried Häuser⁵ and Piercarlo Sarzi-Puttini⁶

Rheumatology 2020;0:1–8

With the publication of the ACR 2010 Cr and ACR 2011 Cr, the definition of FM moved from a predominantly chronic pain condition to a multi-symptom disorder, and the tender point exam has been eliminated as a requirement for diagnosis



There is a considerable agreement between criteria-based diagnoses of FM

Understanding key types of pain

NOCICEPTIVE

NEUROPATHIC

DISFUNCTIONAL/
NOCIPLASTIC

Do we need a third mechanistic descriptor for chronic pain states?

PAIN 157 (2016) 1382–1386

Eva Kosek^{a,*}, Milton Cohen^b, Ralf Baron^c, Gerald F. Gebhart^d, Juan-Antonio Mico^e, Andrew S.C. Rice^f, Winfried Rief^g, A. Kathleen Sluka^h

Nociplastic pain: Pain that arises from altered nociceptive function.



Central sensitization: a biopsychosocial explanation for chronic widespread pain in patients with fibromyalgia and chronic fatigue syndrome

Mira Meeus • Jo Nijs

Clin Rheumatol (2007) 26:465–473

First, **lower pain thresholds at different sites (hyperalgesia)** are reported in patients with FS. The lack of peripheral tissue damage and the lack of a distinct localization of the pain complaints are suggestive of a central abnormality responsible for the chronic widespread pain.

A second important argument in the central sensitization theory for FS concerns the **cognitive, psychological, and behavioral changes** in patients diagnosed with FS. FS patients often present with depression

Finally, brain imaging already provided evidence for **altered brain activity in CFS**.



Neuroinflammation and Central Sensitization in Chronic and Widespread Pain

Ru-Rong Ji, Ph.D., Andrea Nackley, Ph.D., Yul Huh, B.S., M.S., Niccolò Terrando, Ph.D., William Maixner, D.D.S., Ph.D.

Accumulating evidence suggests that central sensitization is **driven by neuroinflammation in the peripheral and central nervous system**. A characteristic feature of neuroinflammation is the activation of glial cells, such as microglia and astrocytes, in the spinal cord and brain, leading to the release of proinflammatory cytokines and chemokines



Recentemente studi di neuroimaging hanno documentato alterazioni consistenti e ricorrenti nell'ambito del Sistema Nervoso Centrale in pazienti affetti da fibromialgia (FM)

In generale, è stata riscontrata una significativa riduzione della densità neuronale ovvero una diminuzione della sostanza grigia cerebrale in numerose aree cerebrali deputate all'elaborazione dell'input nocicettivo (*e.g., talamo, corteccia somatosensoriale primaria, corteccia prefrontale, corteccia cingolata anteriore*), espressione verosimile di un invecchiamento precoce cerebrale, non Alzheimer o età anagrafica – correlato. Anche aree coinvolte nei processi mnestici (*come le aree paraippocampali*) sono spesso interessate.



Functional Magnetic Resonance Imaging Evidence of Augmented Pain Processing in Fibromyalgia

ARTHRITIS & RHEUMATISM
Vol. 46, No. 5, May 2002, pp 1333-1343

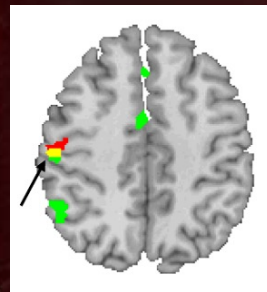
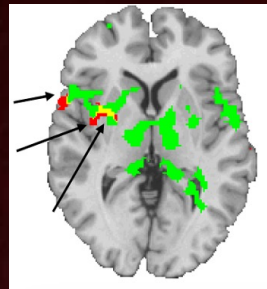
Richard H. Gracely,¹ Frank Petzke,² Julie M. Wolf,³ and Daniel J. Clauw²

Biology and therapy of fibromyalgia

Functional magnetic resonance imaging findings in fibromyalgia

David A Williams^{1,2} and Richard H Gracely^{1,3}

Arthritis Research & Therapy 2006, 8:224



Functional magnetic resonance imaging (fMRI) have helped to provide insights into the role of supraspinal mechanisms in pain perception.

Magnetic resonance spectroscopy, obtains spectra of multiple selected regions and determines the ratio of concentrations of metabolites such as N-acetyl-aspartate, creatine, choline, lactate, glucose and glutamate.

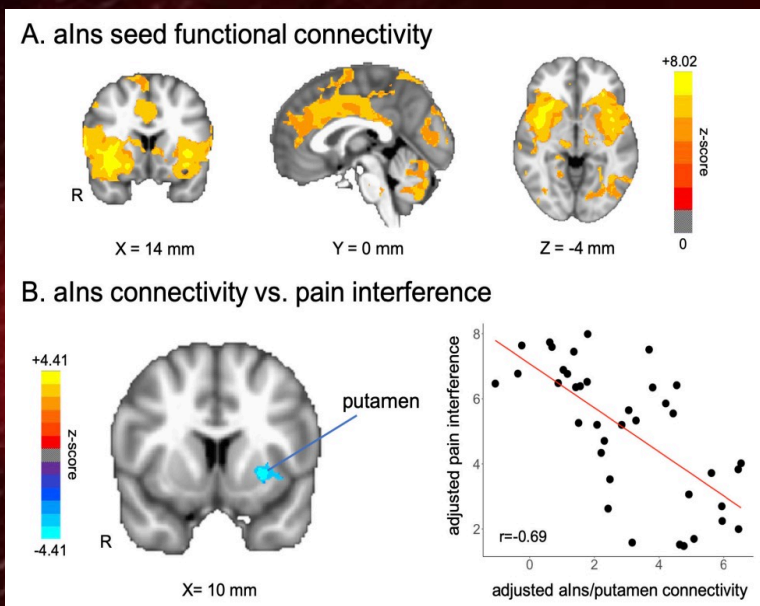
Magnetic resonance perfusion can assess cerebral blood flow and cerebral blood volume, providing measures of baseline differences similar to that currently provided by PET.

Magnetic resonance imaging of neuroinflammation in chronic pain: a role for astrogliosis?

Pain. 2020 July ; 161(7): 1555–1564.

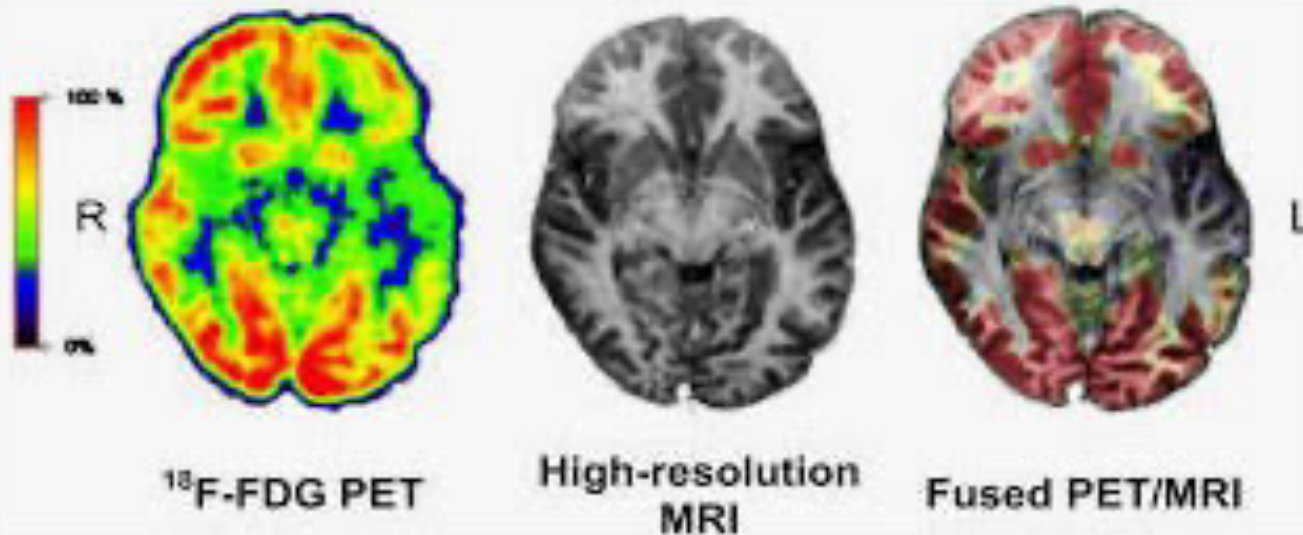
Changjin Jung^{1,2}, Eric Ichesco³, Eva-Maria Ratai^{1,4}, R. Gilberto Gonzalez^{1,4}, Tricia Burdo⁵, Marco L. Loggia^{1,4}, Richard E Harris^{3,*}, Vitaly Napadow^{1,4,6,*}

Several proton **Magnetic Resonance Spectroscopy** (1H-MRS) metabolites have been linked with glial activity (i.e. choline and myo-inositol) and found to be altered in chronic pain patients,



Results demonstrated that cortical choline levels were correlated with glial fibrillary acidic protein, a known marker for *astrogliosis*. Choline, a putative neuroinflammatory 1H-MRS-assessed metabolite elevated in FS and associated with pain interference, may be linked with astrogliosis in these patients.

Le nuove tecnologie, quali SPECT/TC, PET/TC e PET/MRI, uniscono la possibilità di studiare l'attività metabolica di organi e tessuti e la funzionalità delle cellule, insieme ad uno studio anatomico dettagliato.



Imaging integrato: dalla TC-PET alla PET-RM

L'attuale frontiera nell'utilizzo delle radiazioni (ionizzanti e non) in ambito clinico è quello di utilizzare tecniche combinate, in particolare modo accoppiando fra loro due apparecchiature di diagnostica per immagini, o anche una apparecchiatura di diagnostica per immagini e una di terapia radiante.

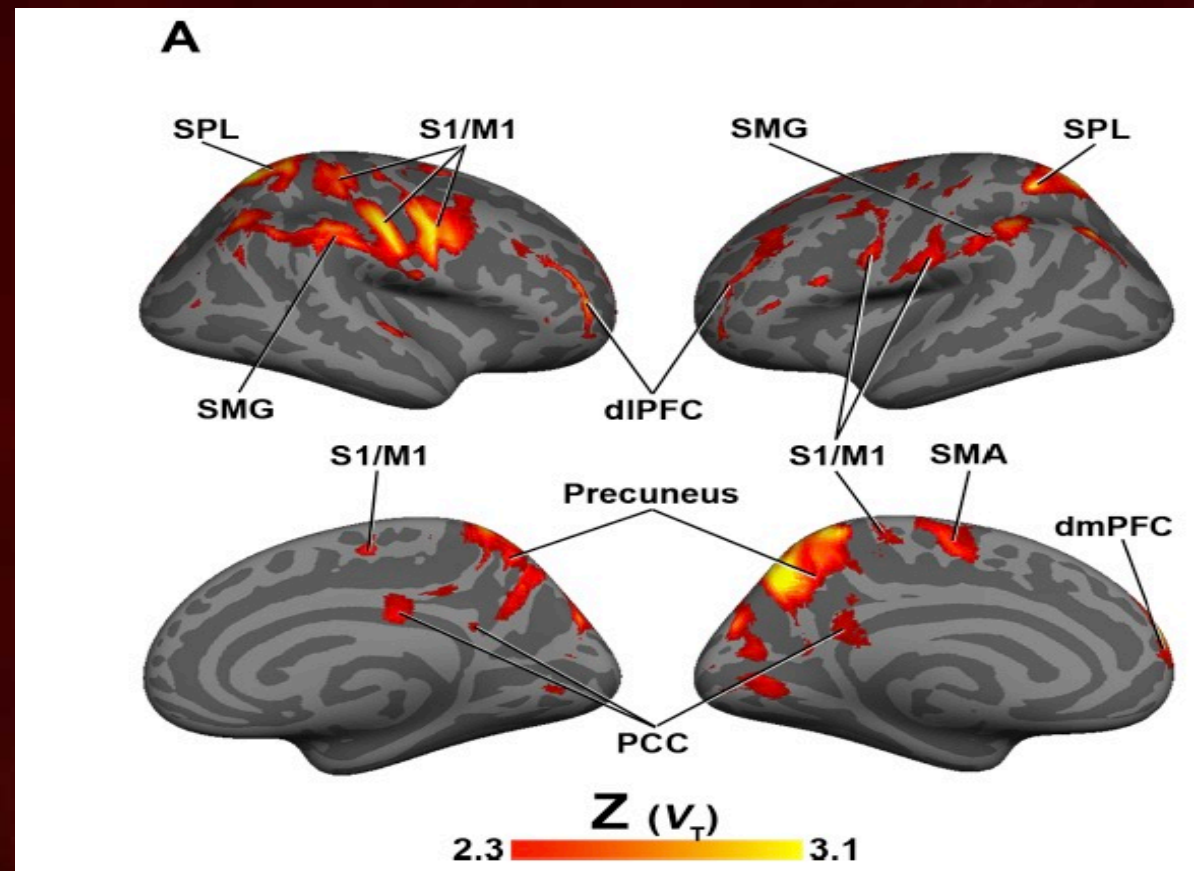
Brain glial activation in fibromyalgia – A multi-site positron emission tomography investigation

Brain, Behavior, and Immunity xxx (

Daniel S. Albrecht^{a,1}, Anton Forsberg^{b,1}, Angelica Sandström^{c,d}, Courtney Bergan^a, Diana Kadetoff^{c,d,e}, Ekaterina Protsenko^a, Jon Lampa^f, Yvonne C. Lee^{g,h}, Caroline Olgart Höglundⁱ, Ciprian Catana^a, Simon Cervenka^b, Oluwaseun Akeju^j, Mats Lekander^{c,d,k}, George Cohen^l, Christer Halldin^b, Norman Taylor^j, Minhae Kim^l, Jacob M. Hooker^l, Robert R. Edwards^m, Vitaly Napadow^{a,m}, Eva Kosek^{c,d,e,*,2}, Marco L. Loggia^{a,*,2}

Our work provides the first in vivo evidence supporting a role for glial activation in FM pathophysiology.

In this study, we conducted a Positron Emission Tomography (PET) study using [11C]PBR28, which binds to the **translocator protein (TSPO)**, a protein upregulated in activated microglia and astrocytes.



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Neuropathic pain component in patients with fibromyalgia

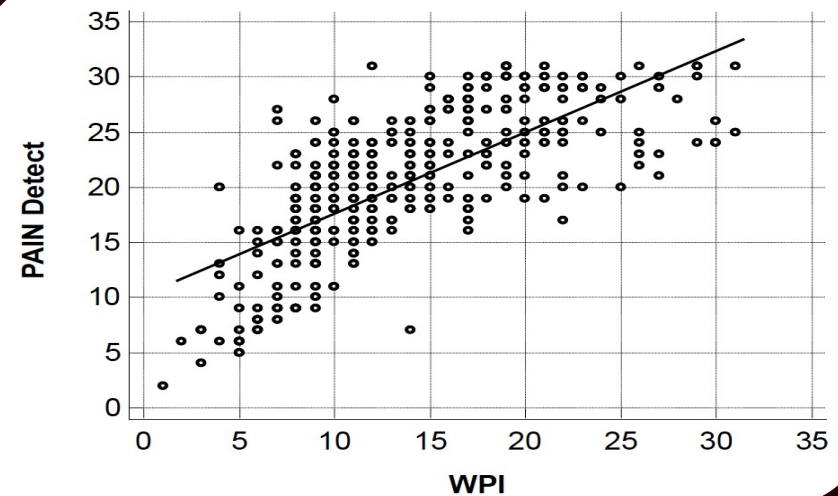
Fausto Salaffi, Sonia Farah, Marco Di Carlo

43%



The study was completed by 393 patients (90% females) whose PDQ scores indicated that 170 (43%) had a possible/likely NP in the PDQ. A strongly significant correlation was seen with the WPI ($r_s=0.66$, $p<0.0001$).

In the logistic regression model, widespread pain (coefficient 0.782; $p<0.0001$), was the only independent variable associated with PDQ.



Scatter plots with regression line illustrating the correlation ($r_s=0.66$, $p<0.0001$), between the Pain Detect questionnaire (PDQ) and the widespread pain (WPI)

Review

> [Neurol Sci.](#) 2022 Mar;43(3):1667-1684. doi: 10.1007/s10072-021-05806-x.

Epub 2022 Jan 14.

The puzzle of fibromyalgia between central sensitization syndrome and small fiber neuropathy: a narrative review on neurophysiological and morphological evidence

[Marina de Tommaso](#) ^{1 2}, [Eleonora Vecchio](#) ^{3 4}, [Maria Nolano](#) ^{3 4}



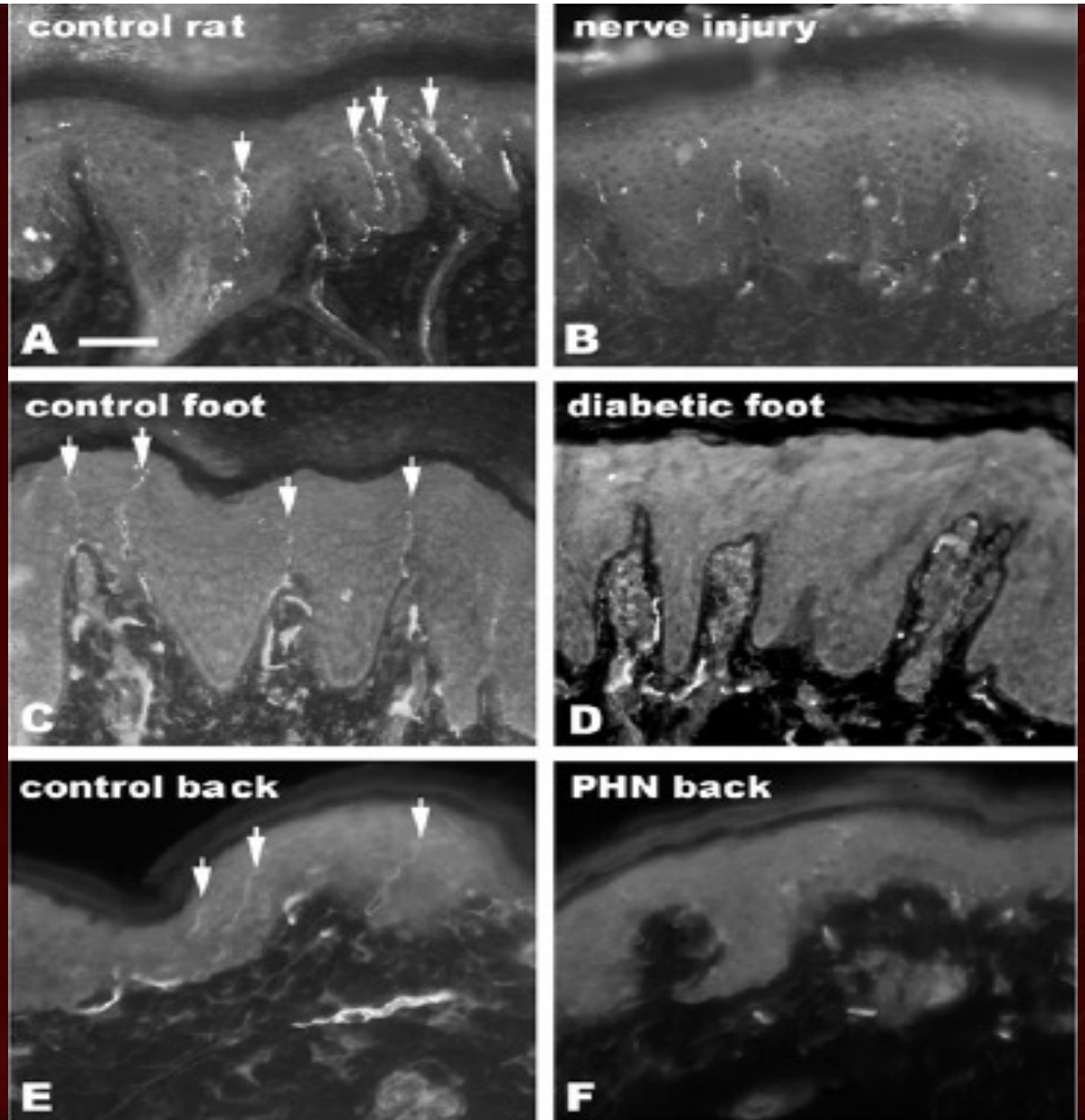
Evidence based on structural and functional neuroimaging methods, electrophysiological, and morphological - skin biopsy - features demonstrated a central and peripheral nervous system involvement.

Phillip J. Albrecht* and Frank L. Rice

Fibromyalgia syndrome pathology and environmental influences on afflictions with medically unexplained symptoms

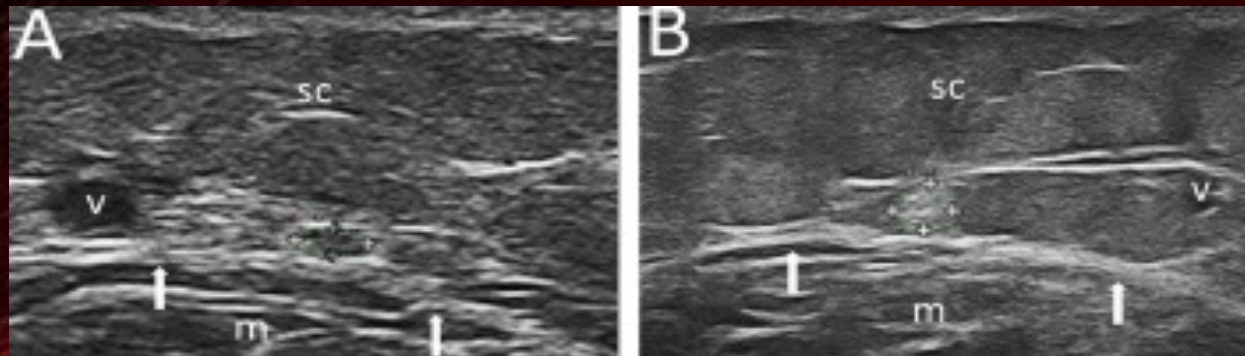
Rev Environ Health 2016; 31(2): 281-294

Intraepidermal nerve fiber (IENF) innervation loss is detected among numerous disorders characterized by chronic pain, including overt nerve trauma, metabolic disorders, and viral infections.



Sural Nerve Size in Fibromyalgia Syndrome: Study on Variables Associated With Cross-Sectional Area

Marco Di Carlo^{1*}, Claudio Ventura^{2†}, Pietro Cesaroni^{1†}, Marina Carotti^{2†}, Andrea Giovagnoni^{2†} and Fausto Salaffi^{1†}



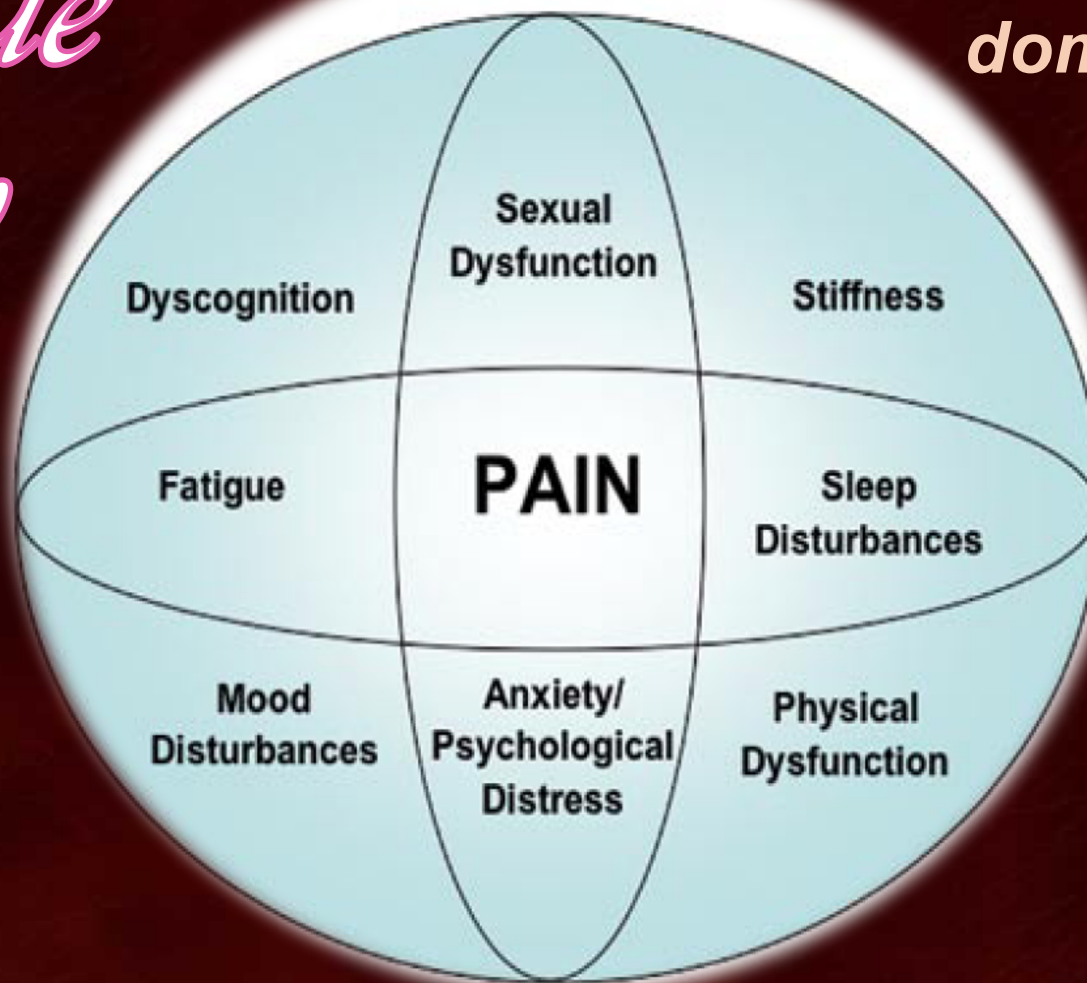
Ultrasound images of the sural nerve at the lateral and distal portion of the calf.

Increased cross-sectional area (CSA) of sural nerve, documented by ultrasound (US), has been revealed in small fibers neuropathy, condition present in about half of patients with fibromyalgia (FM).

Ultrasound examination of the sural nerve at calf level may reveal useful information in patients with FM, identifying a cluster of patients with peripheral nervous system alterations.

Fatigue Sleep

Fibromyalgia domains



Development and validation of the self-administered Fibromyalgia Assessment Status: a disease-specific composite measure for evaluating treatment effect

Fausto Salaffi¹, Piercarlo Sarzi-Puttini², Rita Girolimetti¹, Stefania Gasparini¹, Fabiola Atzeni² and Walter Grassi¹

Arthritis Research & Therapy 2009, 11:R125

Content validity index values for the individual key domains identified by clinicians

	Frequency	Mean importance	Frequency × importance product
Clinician-identified domains			
1. Pain	100	3.9	390.0
2. Fatigue	99	3.7	366.3
3. Sleep quality	93	3.5	325.5
4. Patient global assessment	86	3.4	292.4
5. Physical function	84	3.3	277.2
6. Depression	80	3.2	256.0
7. Anxiety	77	3.3	254.1
8. Clinician global assessment	68	3.3	224.4
9. Quality of life	67	3.2	214.4
10. Occupational dysfunction	64	3.2	204.8
11. Social dysfunction	62	3.2	198.4
12. Cognitive impairment	57	3.2	182.4

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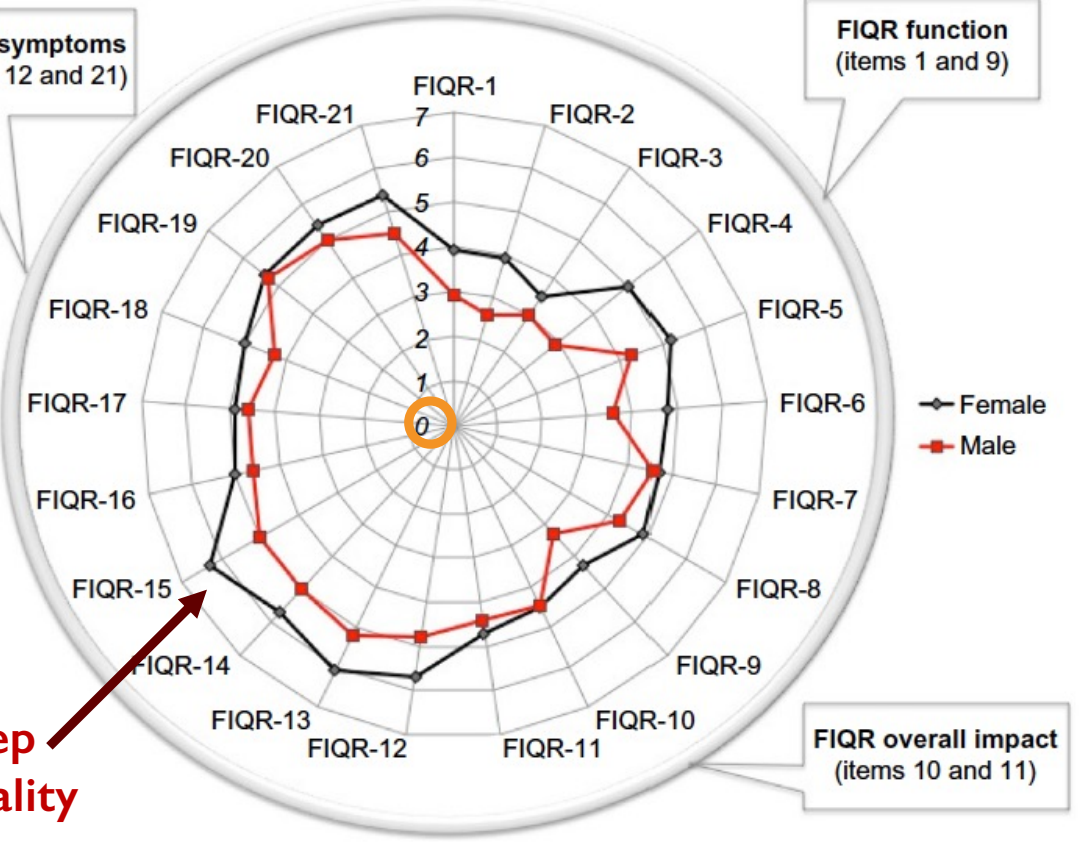
Fausto Salaffi¹
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 Piercarlo Sarzi Puttini⁵

Identifying the symptom and functional domains in patients with fibromyalgia: results of a cross-sectional Internet-based survey in Italy

FIQR symptoms
 (items 12 and 21)

FIQR function
 (items 1 and 9)

Sleep
 Quality



FIQR overall impact subtotal		9.16
12	Pain rating	5.69
13	Fatigue rating	6.04
14	Stiffness rating	5.64
15	Sleep quality	6.18
16	Depression level	4.94
17	Memory problems	4.90
18	Anxiety level	4.99
19	Tenderness level	5.42
20	Balance problems	5.33
21	Environmental sensitivity	5.32

The highest scoring items (those with the greatest disease impact) were the following symptoms related: sleep quality (FIQR 15), fatigue/energy (FIQR 13), pain (FIQR 12), stiffness (FIQR 14), tenderness (FIQR 19), balance problems (FIQR 20), and environmental sensitivity (FIQR 21).

Preliminary identification of key clinical domains for outcome evaluation in fibromyalgia using Delphi methods: the Italian experience

F. Salaffi¹, A. Ciapetti¹, P. Sarzi Puttini², F. Atzeni², C. Iannuccelli³, M. Di Franco³, M. Cazzola⁴, L. Bazzichi⁵

Reumatismo, 2012; 64 (1): 28-35

Domini classificati per rilevanza dai reumatologi.

Dominio	Frequency	Mean importance (MI)	Frequency importance product (FIP)
1. Dolore	79,0	2,85	225,2
2. Fatica	78,6	2,65	208,3
3. Disturbi del sonno	74,6	2,70	201,4
4. Salute-qualità della vita (HRQL)	73,8	2,60	191,9
5. Depressione	70,2	2,60	182,5
6. Ansia	71,8	2,50	179,5
7. Problemi di memoria e di concentrazione	69,1	2,40	165,8
8. Tender points	60,2	2,20	126,4



FIBROMIALGIA

Disturbi dell'umore

- **tassi di depressione maggiore variabili dal 20 all'80% con una mediana del 58%** (Hudson e Pope,1996)
- **tasso di depressione maggiore varia dal 14 al 36%** (Buskila,2007), **valori decisamente più alti di quelli osservabili nella popolazione generale (6,6%)** (Kessler,2003)

Anxiety and coping in patients with chronic work-related muscular pain and patients with fibromyalgia

European Journal of Pain (1998) 2: 309–319

Lillemor R.-M. Hallberg and Sven G. Carlsson



Depression, anxiety, health-related quality of life and pain in patients with chronic fibromyalgia and neuropathic pain

European Journal of Pain 14 (2010) 127.e1–127.e8

Lise Gormsen^{a,*}, Raben Rosenberg^b, Flemming W. Bach^c, Troels S. Jensen^a

EULAR recommendations underplay importance of severe anxiety and depression in fibromyalgia treatment

Ann Rheum Dis December 2017 Vol 76 No 12

The relationship between body mass index and pain, disease activity, depression and anxiety in women with fibromyalgia

Burhan Fatih Koçyiğit¹ and Ramazan Azim Okyay²

PeerJ 6:e4917; DOI 10.7717/peerj.4917

Distinct aberrations in cerebral pain processing differentiating patients with fibromyalgia from patients with rheumatoid arthritis

PAIN 00 (2021) 1–10

Angelica Sandström^{a,b,*}, Isabel Ellerbrock^{a,b}, Monika Löfgren^c, Reem Altawil^d, Indre Bileviciute-Ljungar^c, Jon Lampa^d, Eva Kosek^{a,b,e}

Chronic pain in FM is often associated with comorbidities such as anxiety resulting in a low health-related quality of life.

Fibrofog and fibromyalgia: a narrative review and implications for clinical practice

Howard M. Kravitz · Robert S. Katz

Rheumatol Int (2015) 35:1115–1125

FIBROMIALGIA

Fibrofog

- I pazienti con FM elaborano le informazioni alla stessa velocità dei controlli sani, ma mostrano dei deficit nella MEMORIA A LUNGO TERMINE e nella MEMORIA DI LAVORO (Grace, 1999)
- **La prestazione mnesica è uguale a quella di un soggetto non fibromialgico di 20 anni più anziano** (Park, 2001)



Validity of the Central Sensitization Inventory compared with traditional measures of disease severity in fibromyalgia

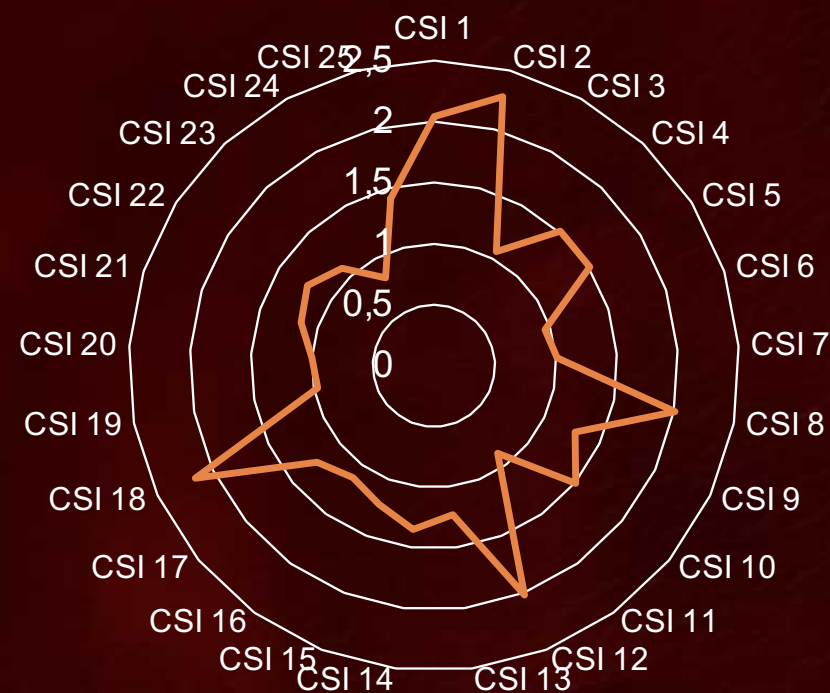
In press 2022

Authors and affiliations

¹Fausto Salaffi, ¹Sonia Farah, ¹Claudia Mariani, ²Piercarlo Sarzi-Puttini, ¹Marco Di Carlo

The study included 562 FM patients, 199 (35.4%) were classified as having central sensitization syndrome

The CSI items linked to fatigue/sleep problems (CSI-1, CSI-8, and CSI-12) and muscle pain/tension (CSI-2 and CSI-18) had the highest ratings (greatest impact). The middle score areas were bruxism (CSI-4), **gastrointestinal issues (CSI-5)**, (CSI-12), and headaches (CSI-10).



Central sensitization: Implications for the diagnosis and treatment of pain

Clifford J Woolf

Pain. 2011 March ; 152(3 Suppl): S2–15

In the **urological tract**, pain hypersensitivity is a feature of interstitial cystitis, chronic prostatitis, endometriosis, and vulvodynia, conditions whose pathophysiology and etiology is however, poorly understood. Although central sensitization has been hypothesized to contribute, not much data is available and few studies have been performed.



Men with chronic prostatitis have though heightened pain sensitivity in the perineum while women with vulvodynia have an enhanced post capsaicin allodynia and secondary hyperalgesia compared to controls

Interstitial Cystitis/Painful Bladder Syndrome and Associated Medical Conditions With an Emphasis on Irritable Bowel Syndrome, Fibromyalgia and Chronic Fatigue Syndrome


J. Curtis Nickel,^{*,†} Dean A. Tripp,[‡] Michel Pontari,[§] Robert Moldwin,^{||}
Robert Mayer,[¶] Lesley K. Carr, Ragi Doggweiler,^{**} Claire C. Yang, Nagendra Mishra
and Jorgen Nordling

0022-5347/10/1844-1358/0
THE JOURNAL OF UROLOGY®

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Printed in U.S.A.

updates

Lower urinary tract symptoms and perineal function in women with and without fibromyalgia: a cross-sectional study

Hellen Cristina Souza de Carvalho Fusco¹  • Marco Antônio Gonçalves Pontes Filho² • Jorge Milhem Haddad³ •
Miriam Raquel Diniz Zanetti⁴ • Amélia Pasqual Marques¹ • Elizabeth Alves Gonçalves Ferreira¹

Association between fibromyalgia and sexual dysfunction in women

Leonid Kalichman

Clin Rheumatol (2009) 28:365–369


Evaluation of sleep disorder and its effect on sexual dysfunction in patients with Fibromyalgia syndrome

Tuba Tülay Koca¹, Günseli Karaca Acet¹, Emrullah Tanrıkut², Burcu Talu³ | Turk J Obstet Gynecol 2016;13:167-71

Lower urinary tract symptoms and perineal function in women with and without fibromyalgia: a cross-sectional study

updates

Clinical Rheumatology (2019) 38:2885–2890

Hellen Cristina Souza de Carvalho Fusco¹  · Marco Antônio Gonçalves Pontes Filho² · Jorge Milhem Haddad³ · Miriam Raquel Diniz Zanetti⁴ · Amélia Pasqual Marques¹ · Elizabeth Alves Gonçalves Ferreira¹

Considering the impact of UI on quality of life, the women in FG scored higher in all the KHQ domains, indicating poorer perception of quality of life when compared with NFG.

OR=5,03

The odds of presenting lower urinary tract symptoms (LUTS) is 5.03 higher in women with FM

Table 4 Impact of urinary incontinence on the quality of life of women with and without fibromyalgia according to King Health Questionnaire (KHQ) domains

KHQ domains	FG (n = 62)	NFG (n = 64)	P value
General health perceptions	75 (25–75)	25 (25–25)	< 0.001
Incontinence impact	33 (0–67)	33 (0–33)	0.531
Role limitations	17 (0–50)	0 (0–33)	0.133
Physical limitations	33 (0–50)	17 (0–50)	0.164
Social limitations	0 (0–11)	0 (0–0)	0.089
Personal relationships	0 (0–17)	0 (0–0)	0.076
Emotion	11 (0–33)	0 (0–11)	0.056
Sleep and energy	17 (0–67)	0 (0–17)	0.003
Severity measures	50 (25–67)	33 (25–50)	0.353

Results represented by median (first-third quartiles). P value calculated using the Wilcoxon-Mann-Whitney test

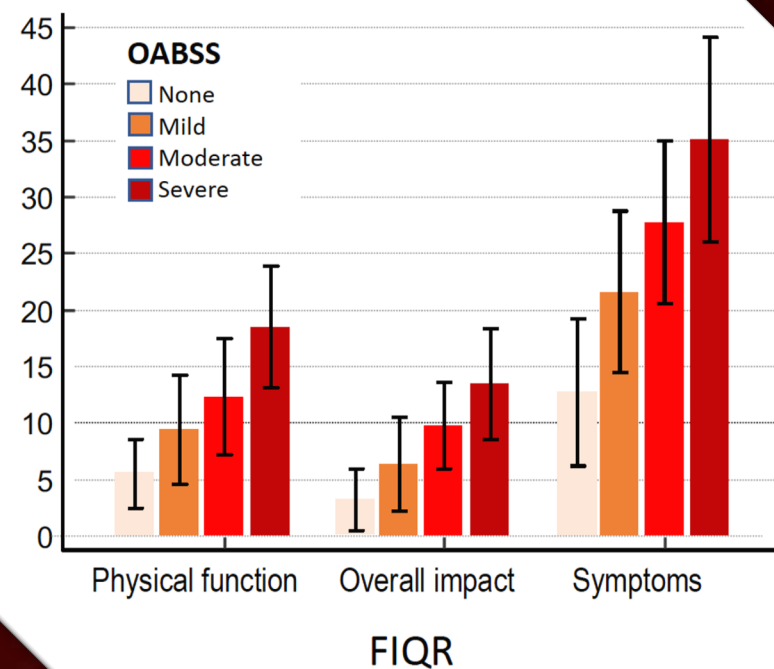
Overactive bladder syndrome and sexual dysfunction in women with fibromyalgia and their relationship with disease severity

F. Salaffi¹, M. Di Carlo¹, S. Farah¹, V. Giorgi², P. Sarzi-Puttini²

Clinical and Experimental Rheumatology 2021

The severity of OAB symptoms showed a direct association with FIQR, suggesting that the more extreme the OAB symptom, the higher the FIQR subdomains

The study included **481** patients, 116 (24.11%) had mild OAB, **82 patients (17.04%) had moderate OAB, and 34 patients had serious OAB (7.06%)**. In 14.17% of patients the bladder condition was causing them major issues in terms of discomfort. In 7.87% of patients the bladder condition was causing them significant problems



Overactive bladder syndrome and sexual dysfunction in women with fibromyalgia and their relationship with disease severity

F. Salaffi¹, M. Di Carlo¹, S. Farah¹, V. Giorgi², P. Sarzi-Puttini²

Clinical and Experimental Rheumatology 2021

Sexual dysfunctions were found in 91/481 of women with FM (18.91%).

19%

Using the FSFI as the dependent variable, multivariate analysis revealed a positive relationship between disease incidence as measured by the **FIQR ($p < 0.0001$)**, **PDQ ($p < 0,0001$)**, and the degree of pain as measured by the **WPI ($p = 0.0037$)**.



Does central sensitization help explain idiopathic overactive bladder?

Nat Rev Urol. 2016 August ; 13(8): 481–491.

W. Stuart Reynolds¹, Roger Dmochowski¹, Alan Wein², and Stephen Bruehl³

Central sensitization describes an induced state of spinal hypersensitivity that is associated with a variety of chronic pain disorders that share many attributes with OAB, albeit without the presence of pain.



An understanding of the pathophysiology and clinical manifestations of central sensitization, and the evidence that supports a role of central sensitization in OAB, including the potential implications of mechanisms of central sensitization for the treatment of patients with OAB could provide a novel approach to the treatment of patients with this disease.



Clinical Evidence of a Neurogenic Mechanism for the Co-existence of Urinary and Bowel Symptoms in Women with Bladder Pain Syndrome

J Urol. 2012 February ; 187(2): 503–507.

Lori Cory¹, Heidi S. Harvie², Gina Northington³, Anna Malykhina⁴, Kristene Whitmore⁵, and Lily Arya³

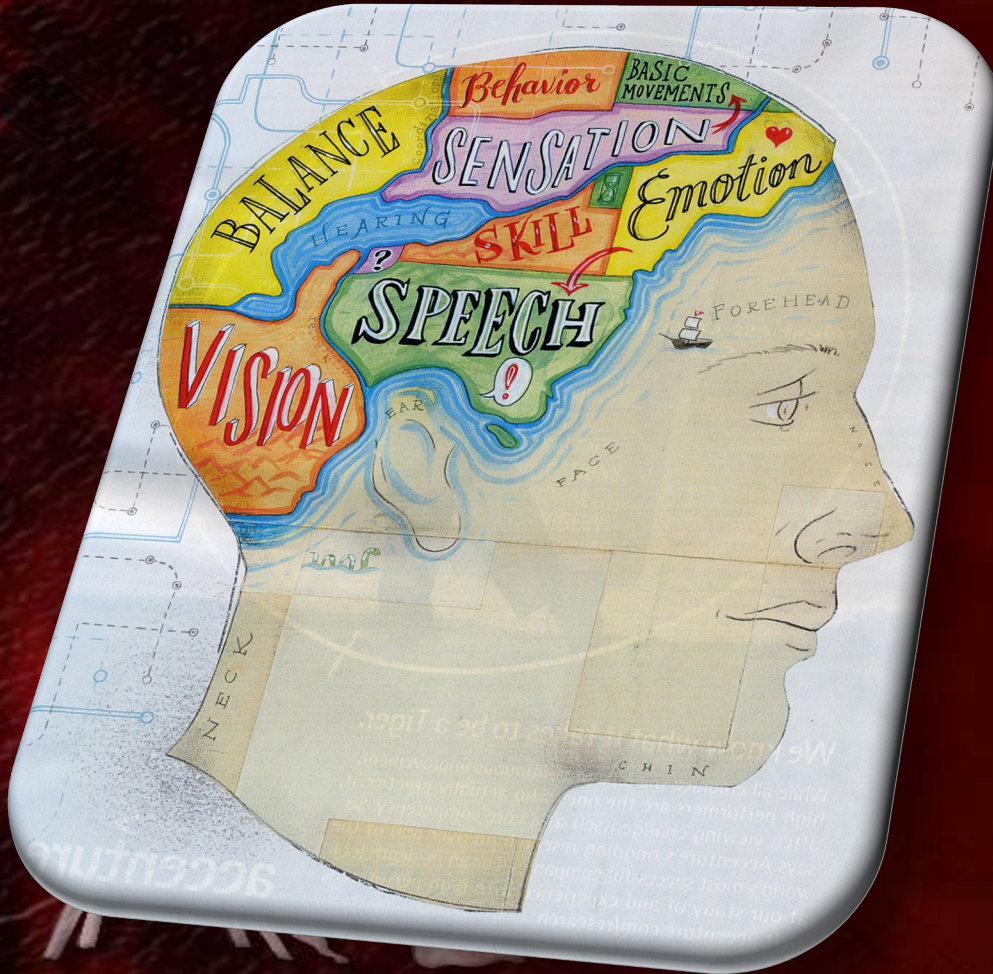
This study provides clinical evidence of a neurogenic mechanism for the co-existence of urinary and bowel symptoms in women with bladder pain syndrome.

Relationship of neuropathic pain to urinary and bowel symptoms and quality of life

	Total Neuropathic Pain Score [¥]	
	R value [*]	p-value
Total urinary symptom score (ICSI)	0.31	<0.001 ^{†‡}
Total quality of life score (ICPI)	0.28	<0.001 ^{†‡}
Total bowel symptom score (IBS)	0.49	<0.001 ^{†‡}

In women with bladder pain syndrome, the presence of neuropathic pain is significantly associated with the severity of bladder and bowel symptoms, quality of life and pain catastrophizing. A neurogenic mechanism may explain the co-existence of urinary and bowel symptoms in women with bladder pain syndrome.

GRAZIE per l'attenzione



- **Wolfe : “fibromyalgia will always exist regardless of the name given to the syndrome”.**

Wolfe E. The fibromyalgia problem. *J Rheumatol* 1997; 24:1247-9.