Decree of the Rector n. 1254 of 29/11/2023  
Competition for awarding 1 research grant at the University of Udine

DISCLAIMER:
The official and legally binding call for applications is in Italian only.  
This document cannot be used for legal purposes and is only meant to provide information in English on the call for applications (Decree of the Rector n. 1254 of 29/11/2023). Please refer to the official call published on: https://www.uniud.it/it/albo-ufficiale

Any change and integration will be made available on the above mentioned web page. Therefore, no personal written communication regarding the examination date and/or competition results shall be provided to applicants.

Annex 1

Competition announcement for the assignment of 1 research grant at the University of Udine, entitled “Design, synthesis and investigations on new G-quadruplex aptamers against STAT3 and the interleukine-6 receptor (G4-APTASTAT)” SSD: BIO/10 (principal investigator, Valentina Rapozzi)

Research grant funded by the resources of the project PRIN 2022 - Prot. n. 2022KZ4JRM

Art. 1

A selection procedure is hereby launched for the award of 1 research grant at the University of Udine, as identified in Attachment A which constitutes an integral part of the present announcement. The research grant is linked to the research project and is subject and conditioned upon the relative funding. The fellowship may be renewed, in compliance with Art. 22, Law No. 240 of 30 December 2010 (as in the text in force before the implementation of the Conversion Law of the D.L. 36/2022, L. 79/2022), Law No. 11 of 27 February 2015, and the current regulations of the University of Udine for awarding research grants, issued with the Rector’s Decree No. 182 of 31 March 2021. The renewal is subject to the scientific coordinator’s positive assessment of the researcher’s activities, an adequate scientific rationale, and a corresponding financial covering.

The research fellowship does not give rise to any right with regards to accessing University posts.

Any personal communication to candidates related to this selection will be sent exclusively to the email address indicated when registering for the selection, as mentioned in Art. 5.

Art. 2

The research grant described in this competition announcement and the required qualifications to apply for the position are identified in Attachment A. The lack of the admission requirements leads to the automatic exclusion from the competition procedure.

Possession of a PhD or equivalent degree obtained abroad or, only for the interested areas, of a medical specialization accompanied by an adequate scientific production, constitutes a preferential
qualification for awarding the research fellowship of this selection, if it has not been provided as a mandatory requirement.

For the only purpose of the admission to the competition, the Examining Board (Art. 7) shall assess the equivalence of the qualification obtained abroad, except for the evaluation of the medical specialization qualification to which Article 38 of the Legislative Decree 165/2001 and subsequent modifications and additions, and EU regulations on the matter, shall be applied.

The Examining Board will proceed to the evaluation of the qualification obtained abroad according to the documentation attached to the application form. The Examining Board may exclude the candidate if the submitted documentation does not provide sufficient information for the assessment.

Therefore, applicants must enclose all the documentation in their possession relating to their qualification in order to provide the Examining Board with sufficient information for assessment.

Candidates holding a qualification issued by a European Research Area country, if successful, must submit, if not already attached to the application form one of the following options:
- Supplement Diploma in English issued by the competent University.
- CIMEA Certificate of comparability of the foreign qualification, issued by CIMEA (Information Centre on Academic Mobility and Equivalence) via the "diplome" service at https://cimea.diplo-me.eu/udine/#/auth/login

Candidates holding a qualification issued by a non-European Research Area country, if successful, must submit, if not already attached to the application form one of the following options:
- Declaration of the on-site value of the qualification and the certificate relating to the degree with examinations and grades. A certificate in a language other than Italian or English must be accompanied by an official translation into one of these languages (certified by the competent diplomatic-consular authority or certified by a court in Italy).
- CIMEA Certificate of comparability of the foreign qualification, issued by CIMEA (Information Centre on Academic Mobility and Equivalence) via the "diplome" service at https://cimea.diplo-me.eu/udine/#/auth/login

If the Supplement Diploma or the statement/attestation of comparability are not available when signing the contract, the applicant must demonstrate that he/she has requested the documentation and submit it as soon as possible.

Any exclusion from the selection procedure due to lack of eligibility requirements, absence of required documents, failure to sign the selection application or submission of the selection application in a manner different from what is provided for in this call for applications will be communicated to applicants exclusively at the email address indicated in the application form.

**Art. 3**

The research grant referred to in this call for applications cannot be awarded:

a. to employees of Universities and the entities referred to in Article 22, section 1, of Italian Law no. 240 of 30 December 2010 (in the text prior to the reform introduced by Law no. 79 of 29 June 2022);

b. to those who have already been awarded research grants pursuant to Italian Law no. 240 of 30 December 2010 (prior to the reform introduced by Law no. 79 of 29 June 2022) for the maximum period provided by law, even if not continuously, excluding the period in which the grant was used in conjunction with the doctorate, up to the legal term of the relative course;
c. to those who have already benefited from research grants and fixed-term researcher contracts provided for, respectively, in Articles 22 and 24 of Italian Law no. 240 of 30 December 2010 (in the text prior to the reform introduced by Law no. 79 of 29 June 2022), for a total of 12 years, even if not consecutive;
d. to anyone who has a degree of kinship or affinity, up to and including the fourth degree, with:
- the Rector, the Director General or a member of the Board of Directors of the University of Udine;
- the scientific supervisor or a professor/researcher belonging to the department or organisation hosting the research grant in question.

The research grant provided for in this call for applications cannot be combined:

a) with scholarships of any kind, except for those granted by Italian or foreign institutions to supplement, by means of stays abroad, the fellow's training or research activities;
b) with other research grants;
c) with an employment relationship, even if part-time, without prejudice to the relevant provisions for employees of public administrations.

The grant awarded under this call for applications is also incompatible with simultaneous attendance at university degree courses, either Bachelor's degree or Master's degree courses, research Doctorates with scholarships and medical specializations, in Italy or abroad.

### Art. 4

Applicants must enclose with their application, under penalty of exclusion, the following documents:

a) their professional scientific CV, highlighting the candidate’s aptitude for carrying out and implementing the research project (Attachment A);
b) their identity card, their passport or any other identification document¹;
c) (for candidates with a foreign qualification only) certification or self-certification of both the academic qualification required for the admission to the selection, and of the exams (with evaluation) took during the period of study abroad, and of any other document that can be useful to the evaluation of the degree by the Examining Board.

Applicants can attach to the application, publications and any other certification considered useful to demonstrate the qualification based on the research program (Attachment A) and to certify any research activity accomplished at public or private institutes (indicating the starting and ending date and the duration).

The documents and qualifications mentioned above must be submitted in Italian or English. Those that are not as requested will not be evaluated. Documents originally written in a language other than Italian or English must come with a translation in Italian or English, that the candidate will do on its own responsibility. The translation can be an abstract concerning the thesis.

Italian and Community candidates wishing to submit qualifications referring to conditions and facts attested by Public Administrations must proceed exclusively with self-certification.
Non-EU citizens legally residing in Italy may self-certify only data that can be verified or certified by Italian public bodies. They may also use declarations in lieu when provided for by an international convention between Italy and the declarant's country of origin.
Non-EU citizens not residing in Italy cannot self-certify.

Only the qualifications possessed by the candidate on the date the application form is submitted are

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¹ Please be aware that the residence permit is not an identification document.
submitted in accordance with the procedures set out in Article 5 will be assessed.

Failure to submit mandatory documents provided for in this article will constitute grounds for exclusion from the selection.

Art. 5
The submission of the applications for the present call starts on December 7, 2023 at 2:00 pm (Italian time) and ends on January 26, 2024 at 2:00 pm (Italian time).

The application to take part in the selection must be completed, under penalty of exclusion, using the appropriate online procedure, available at the link [https://pica.cineca.it/](https://pica.cineca.it/)
The procedure involves an applicant registration step, for those who do not already have an account, and then an application completion step.

Once completed, the online application must be signed in the manner described in the online procedure (manual signature with attached identity document or digital signature), under penalty of exclusion from selection. The application does not have to be signed if you access the above-mentioned online procedure using your SPID ID.

The qualifications referred to in Article 4 must be attached to the application in .pdf format. Individual .pdf files may not exceed 30MB.

The application for participation in the selection is automatically sent to the University of Udine with the definitive closing of the online procedure.

The University Administration:
- is not responsible if it is impossible to read the submitted documentation in electronic format due to damaged files;
- shall not accept or take into consideration qualifications or documents received in paper form or by any means other than what is specified in this article.

Reference to documents or publications already submitted in connection with other competitions is not allowed.

The Administration is not responsible for any missing document or communication because of inaccurate indication of residence and/or address submitted by the candidate during the application. Also, the Administration is not responsible if the candidate has not communicated changes in this information, or has communicated them too late. The Administration is also not responsible for any postal or telegraphic problems not attributable to the Administration itself.

Applicants are advised not to wait until the last few days before the closing date to submit their application. The University accepts no responsibility for any malfunctions due to technical problems and/or overloading of the communication line and/or application systems.

Art. 6
The selection procedure is held in accordance with the modality indicated in Attachment A.

The test will aim to assess the general preparation, experience and aptitude for research of the candidate. It will consist in the evaluation of the professional scientific curriculum, of the publications and qualifications presented, and of the interview, where foreseen.
Art. 7

The Examining board for the competition is identified in Attachment A of the present competition announcement, of which it is an integral part.

At its first meeting, the Examining board shall appoint its President and Secretary, and establish the criteria and methods for evaluating the qualifications and the interview, where foreseen.

The results of the qualifications assessment must be disclosed to applicants during the interview, where foreseen.

The Examining board can award a maximum of 100 points (one hundred out of one hundred) to the selection.

At the end of the evaluation procedure, the Examining board shall formulate the general merit list based on the overall score of each candidate, and draw up the minutes of the whole competition procedure.

Based on the ranking list, the assignment is awarded to candidates who have obtained a minimum overall score of 70/100 (seventy out of one hundred).

The Examining board's judgement is final.

The ranking list will be made public exclusively through publication on the University's official website.

Applicants will not be notified of the outcome of the evaluation.

Those who do not declare their acceptance of the research grant and do not present themselves at the research centre within the deadline communicated by the latter, even if not formally, shall lose the right to receive it. Exceptions to this term will only be granted in cases of documented force majeure.

Art. 8

The research activity cannot be started before signing the contract defining the terms and conditions of the collaboration.

The activity covered by the research grant must have the following characteristics:

a) it must be carried out as part of the research programme covered by the grant and not be a merely technical support to it;

b) it must have a close connection with the realization of the research program for which the winner of the grant has been awarded the contract;

c) it must be continuous and, in any case, temporally defined, not merely occasional, and in coordination with the overall activity of the University;

d) it must be carried out autonomously, solely within the limits of the programme prepared by the programme supervisor, without predetermined working hours.

The researcher is required to submit a detailed written report on the work carried out and the results achieved, accompanied by the opinion of the scientific supervisor, to the reference organisation at the intervals set out in the contract. The researcher must also submit interim reports and timesheets, if requested by the reference organisation.
Either the fellow or the reference organisation may withdraw from the contract. The reference organisation may terminate the contract not only in the cases referred to in Article 9, sections 2 and 3, of the "Internal rules for awarding research grants pursuant to law 240 of 30 December 2010" of the University of Udine, but also in the event the research project and therefore the financial coverage on which the research grant is based cease to exist.

**Art. 9**

The following legal dispositions shall apply to the grant referred to in this call for applications:
- for tax matters, the provisions of Article 4 of Italian Law no. 476 of 13 August 1984, as subsequently amended and supplemented;
- for social security matters, the provisions of Article 2(26) et seq. of Italian Law no. 335 of 8 August 1995, as subsequently amended and supplemented;
- for mandatory maternity leave, the provisions of the Italian Ministerial Decree of 12 July 2007;
- with regard to sick leave, the provisions of Article 1(788) of Italian Law no. 296 of 27 December 2006 and subsequent amendments.

During the period of mandatory maternity leave, the allowance paid by INPS according to Art. 5 of the Italian Ministerial Decree of 12 July 2007 is supplemented by the University up to the full amount of the research grant.

The grant will be paid in monthly instalments.

**Art. 10**

The data collected as part of the procedure referred to in Art. 5 are necessary to properly manage the selection procedure, for any subsequent management of the research grant and for purposes related to managing services provided by the University. The University of Udine is the Data Controller. At any time, the data subject may request access, rectification and, depending on the University's institutional purposes, cancellation and restriction of processing or oppose the processing of their data. The data subject can always lodge a complaint with the Italian Data Protection Authority. The complete disclosure is available on the University of Udine website in the "Privacy" section, accessible from the home page www.uniud.it Direct Link: https://www.uniud.it/it/pagine-speciali/guida/privacy

**Art. 11**

For all matters not expressly mentioned in this call for applications, refer to the regulations in force on the subject cited in the introduction and to the "Internal rules for awarding research grants pursuant to Italian Law no. 240 of 30 December 2010" of the University of Udine, issued by Rector's Decree no. 182 of 31 March 2021.

**Art. 12**

The procedure supervisor is Dr Sandra Salvador, Head of the Research Services Area of the University of Udine.

The Responsible office at the University of Udine is "Area Servizi per la Ricerca - Ufficio Formazione per la Ricerca", via Mantica n. 31 - 33100 Udine, Italia.

To request information about the call for applications, please complete the following form available on the University of Udine website: https://helpdesk.uniud.it/SubmitSR.jsp?type=req&accountId=universityofudine&populateSR_id=42105
Attacco A

Responsabile scientifico della ricerca / Principal investigator:

Nome e cognome / Name and surname: Valentina Rapozzi
Qualifica / Position: Professoresa Associata / Associate Professor
Dipartimento / Department: Area Medica / Medicine
Area MUR / Research field: 05 – Scienze biologiche / Biological sciences
Settore concorsuale e Settore scientifico disciplinare / Scientific sector: 05/E1; BIO/10 – Biochimica

Titolo dell’assegno di ricerca / Topic of the research fellowship “assegno di ricerca”:
I bandi sono consultabili dal sito dell’Ateneo, del MUR e di Euraxess / The calls are available on the University, MUR and Euraxess websites

Testo in italiano:
Progettazione, sintesi e studio di nuovi aptameri G-quadruplex contro STAT3 e il recettore dell’interleuchina-6 (G4-APTASTAT).

Testo in English:
Design, synthesis and investigations on new G-quadruplex aptamers against STAT3 and the interleukine-6 receptor (G4-APTASTAT).

Obiettivi previsti e risultati attesi del programma di ricerca in cui si colloca l’attività dell’assegno di ricerca / Foreseen objectives and results of the research programme performed by the research fellow “assegno di ricerca”:
I bandi sono consultabili dal sito dell’Ateneo, del MUR e di Euraxess / The calls are available on the University, MUR and Euraxess websites

Testo in italiano:
STAT3 è un fattore attivato in risposta al legame di un certo numero di ligandi, tra cui citochine (ad es. IL-6) e altri fattori, ai loro recettori di superficie cellulare affini (ad es. IL-6R). Questa proteina è stata identificata come un importante bersaglio per la terapia del cancro, poiché partecipa all’oncogenesi attraverso la sovraregolazione di geni che codificano per inibitori dell’apoptosi (Bcl-xL, Mcl-1 e survivina), regolatori del ciclo cellulare (ciclina D1 e c-myc) e induttori dell’angiogenesi. Evidenze crescenti hanno dimostrato che STAT3 è anche costitutivamente attivato in molti tumori umani. Inoltre, studi recenti hanno evidenziato il significato prognostico dei livelli di STAT3 nel carcinoma mammario invasivo e nei cordomi. Pertanto, il targetting di questa proteina attraverso ligandi con elevata affinità e specificità ha un potenziale sia terapeutico che diagnostico. T40214 è un aptamero oligonucleotidico G-quadruplex (G4) [(G3C)46] in grado di legare STAT3, che si è dimostrato efficace contro diverse linee cellulari tumorali e xenotrapianti tumorali. Tuttavia, T40214 richiede la formazione di un complesso con polietilenimmina per essere erogato in modo efficiente nelle cellule e protrarre la degradazione.

Inoltre, la cinetica di ripiegamento per la formazione della conformazione di G4 attiva e nucleasi resistente è piuttosto lenta, considerando la bassa concentrazione di K+ (necessaria per il ripiegamento della struttura G4) presente a livello extracellulare. La progettazione e l’implementazione di opportune modifiche per migliorare le proprietà generali di T40214 richiede informazioni dettagliate riguardanti sia la conformazione attiva dell’aptamero che la sua interazione con la proteina bersaglio. Sebbene sia stata proposta una conformazione G4 antiparallela per T40214, questa struttura è stata severamente messa in discussione da recenti indagini che hanno dimostrato che T40214 adotta una conformazione G4 parallela. Queste considerazioni richiedono una revisione approfondita delle indagini riguardanti il complesso T40214/STAT3, in quanto mancano completamente studi vincolanti riguardanti la loro interazione in sistemi isolati. Un altro aptamero correlato a T40214 e che ha come bersaglio STAT3, vale a dire T40231 [G2T(G3T)2G3], ha anche mostrato proprietà biologiche similia
STAT3 is a factor activated in response to the binding of a number of ligands, including cytokines (e.g. IL-6) and other factors, to their cognate cell surface receptors (e.g. IL-6R). This protein has been identified as an important target for cancer therapy, since it participates in oncogenesis through the upregulation of genes encoding apoptosis inhibitors (Bcl-xL, Mcl-1 and survivin), cell-cycle regulators (cyclin D1 and c-myc), and inducers of angiogenesis. Growing evidences have shown that STAT3 is also constitutively activated in many human cancers. Furthermore, recent studies have highlighted the prognostic significance of STAT3 levels in invasive breast cancer and chordomas. Therefore, targeting this protein through ligands with high affinity and specificity has both therapeutic and diagnostic potential. T40214 is an oligonucleotide G-quadruplex (G4) aptamer [(G3C)4] able to link STAT3, which has been proven efficient against several cancer cell lines and tumor xenografts. However, T40214 requires the formation of a complex with polyethylenimine to be efficiently delivered in cells and protected against degradation. Moreover, the folding kinetic for the formation of the active and nuclease resistant G4 conformation is quite slow, considering the low K+ concentration (required for the folding of the G4 structure) present extracellularly. The design and implementation of appropriate modifications to enhance the general properties of T40214 requires detailed information concerning both the active conformation of the aptamer and its interaction with the target protein. Although an antiparallel G4 conformation has been proposed for T40214, this structure has been severely questioned by recent investigations showing that T40214 adopts a parallel G4 conformation. These considerations require an in-depth revision of the investigations concerning the T40214/STAT3 complex, since binding studies regarding their interaction in isolated systems are completely lacking. Another aptamer related to T40214 and targeting STAT3, namely T40231 [(G2T(G3T)2G3], has also shown similar biological properties as T40214. Interestingly, another G4 aptamer with a sequence similar to T40231, namely T30923 [(G3T)4], has been proven to bind to IL-6R which in turn activates STAT3. Considering that T40214 and T30923 adopt similar parallel G4 structures, these data give us the unique opportunity to design and prepare an aptamer targeting both proteins, IL-6R and STAT3, involved in the same bio-molecular pathway.

The main aims of this project are:

1) preparing and studying T40214 analogues with improved general stability properties;
2) investigating the interaction between T40214, its analogues and STAT3;
3) preparing and studying modified aptamers able to efficiently bind to both STAT3 and IL-6R;
4) exploring the possibility to engraft the designed aptamers into POPC liposomes in order to improve their delivery into cancer cells;
5) evaluating the biological activity of the most promising aptamers against pancreatic cancer adenocarcinoma cells and other cancer cell lines.
Struttura dell’Università di Udine presso la quale verrà sviluppata l’attività di ricerca / Department or other structure of the University of Udine where research activities will be carried out:

Dipartimento di Area Medica (DAME) / Department of Medicine

Importo dell’assegno di ricerca (al lordo oneri carico assegnista) / Total grant gross for the research fellowship:

€ 19,457.28

Durata dell’assegno di ricerca / Duration of the research fellowship “assegno di ricerca”:

12 mesi / months

Finanziamento / Financed by:


Requisiti di ammissione / Minimum qualifications necessary:

- Possesso di un diploma di laurea vecchio ordinamento (ante decreto 3 novembre 1999 n. 509) o di laurea specialistica/magistrale (ex decreto 3 novembre 1999 n. 509 e decreto 22 ottobre 2004 n. 270) o titolo equivalente conseguito all’estero;
- possesso di un curriculum scientifico professionale idoneo allo svolgimento dell’attività di ricerca contemplata.
- University degree obtained before Decree n. 509 of 3 November 1999 or specialistic/Master’s degree (post decree n. 509 of 3 November 1999 and decree n. 270 of 22 October 2004) or equivalent degree obtained abroad;
- professional scientific curriculum suitable for the research activity above mentioned.

Procedura selettiva / Competition procedure:

Valutazione per titoli e colloquio / Evaluation of titles and oral exam

I risultati della valutazione dei titoli saranno resi noti agli interessati nel corso del colloquio / The evaluation of the qualifications will be disclosed to candidates during the interview

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<tr>
<th>Calendario del colloquio / Calendar of the oral exam</th>
<th>Modalità / Modality</th>
<th>Videoconferenza / Videoconference</th>
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<tr>
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Per sostenere il colloquio i candidati devono esibire un valido documento di riconoscimento. / Candidates must come to the interview with a valid identity document.

Eventuali variazioni saranno rese note esclusivamente mediante pubblicazione all'albo ufficiale on line dell'Ateneo / Any change will be made public solely through publication on the University web site
http://web.uniud.it/ateneo/normativa/albo_ufficiale

Nota / Note: Le indicazioni sulle modalità di svolgimento della prova in modalità telematica saranno inviate ai candidati con successiva email da parte del Presidente della Commissione. Ai fini dell'identificazione e a pena di esclusione dalla procedura selettiva, ciascun candidato è tenuto ad identificarsi prima che il colloquio abbia inizio, esibendo il medesimo documento di identità allegato alla domanda di ammissione al concorso. Il candidato deve risultare reperibile nella giornata e all'orario indicato sul bando. Il mancato collegamento, l’irreperibilità del candidato nel giorno o nell’orario stabilito o la mancata esibizione del documento identificativo, sono motivo di esclusione dalla procedura selettiva. La registrazione delle prove orali è vietata. L'Ateneo adotterà pertanto tutti i provvedimenti in suo potere per tutelare i soggetti coinvolti qualora venissero diffuse tramite internet – o altri mezzi di diffusione pubblica – video, audio o immagini della procedura selettiva. / Instructions on how the video interviewing will be conducted will be provided to candidates by the Chairman of the Examining Board via email. For identification purposes, each candidate is required to identify him/herself before the interview by exhibiting the same identification document attached to the application. Candidates must be available on the day and time established by the call for applications. Failure of the candidate to establish a video connection, the unavailability of the candidate on the day and/or time established or failure of the candidate to provide the required identification document are all grounds for exclusion from the selection procedure. Recording of the video interviews is prohibited. The University will adopt all the measures within its power to protect all personnel involved as a result of dissemination via the internet or via other forms of public dissemination, of videos, audios or other pictures of the selection procedures.

Commissione giudicatrice / Examining Board:

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<td>Barbara Skerlavaj</td>
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